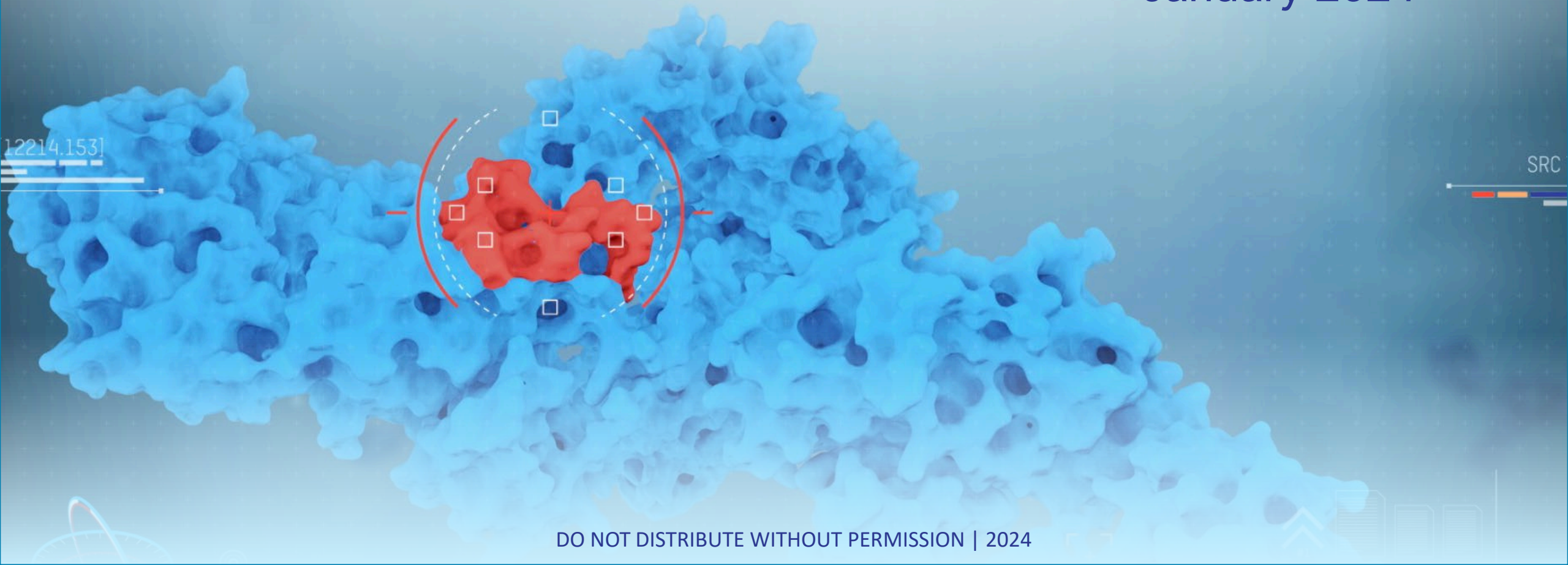


# Corporate Presentation

January 2024



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This presentation includes forward-looking statements regarding our drug candidates, the timing and outcome of regulatory decisions, future availability of clinical trial data, our collaborations for our product candidates and the maintenance of those collaborations, business and results from operations, and other matters. Actual results could differ materially from those contained in any forward-looking statements as a result of various factors, including without limitation: that our drug candidates do not advance in development or result in approved products on a timely or cost effective basis or at all; the cost, timing and results of clinical trials; that many drug candidates that have completed early-stage trials do not become approved drugs on a timely or cost effective basis or at all; the ability to enroll patients in clinical trials; possible safety and efficacy concerns; regulatory developments; our ability to protect our intellectual property rights, and unexpected costs, charges or expenses that reduce cash runway. Our pipeline programs are in various stages of pre-clinical and clinical development, and the process by which such pre-clinical or clinical therapeutic candidates could potentially lead to an approved therapeutic is long and subject to significant risks and uncertainties. These and other risks and uncertainties that we face are described in our most recent Annual Report on Form 10-K, and in other filings that we make with the Securities and Exchange Commission from time to time. We undertake no obligation to update forward-looking statements as a result of new information or otherwise.

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and growth and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions, and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

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# Corporate Highlights

We are a clinical stage biotech company developing antibody therapeutics for cancer

## LEAD ASSETS



CTX-009: DLL4 x VEGF-A bispecific antibody

CTX-471: CD137 agonist antibody

CTX-8371: PD-1 x PD-L1 bispecific antibody

## CORE SCIENCE



StitchMabs™ platform designed to identify synergistic bispecific antibodies

Common Light Chain technology enables multi-specificity and manufacturability

Focus on Translational research

## RESOURCES



Cash runway into mid-2026 (YE 2023: \$152M)

Funded by leading life-science investors

~32 FTEs based in Boston, MA with experienced leadership team

# Focused Pipeline with Multiple Value Inflection Points

Program	Target	Discovery	IND Enabling Pre-Clinical	Phase 1	Phase 2	Phase 3	Next Milestone
CTX-009	DLL4 x VEGF-A	COMPANION-002: BTC					Top line data in U.S. H2 2024
		COMPANION-003: Colorectal					Top line data in U.S. Mid-2024
		COMPANION-004: TBD					Initiate in U.S. H2 2024
CTX-471	CD137	CD137 agonist (monotherapy)					Fully enrolled
		CD137 + PD-1 (combination)*					Top line data in U.S. H1 2025
CTX-8371	PD-1 x PD-L1	Solid Tumors					Initiate Phase 1 Q1 2024

\*Clinical collaboration with Merck & Co. Inc., Rahway NJ USA in combination with anti-PD-1 therapy KEYTRUDA®

# Leadership Team Experienced in Drug Discovery and Development



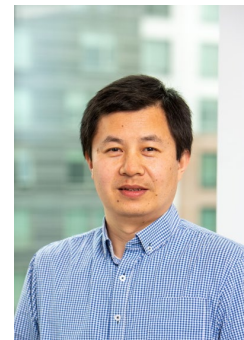
**Jon Anderman**  
VP, Head of Legal



**Vered Bisker-Leib, PhD, MBA**  
CEO



**Ian Chia, PhD**  
VP, Business Development



**Bing Gong, PhD**  
VP, Protein Sciences



**Karin Herrera**  
VP, Clinical Operations



**James Kranz, PhD**  
VP, CMC



**Neil Lerner, CPA, MIM**  
VP, Finance



**Minori Rosales, MD, PhD**  
SVP, Head of Clinical Development



**Kris Sachsenmeier, PhD**  
VP, Translational Science



**Thomas J. Schuetz, MD, PhD**  
President of R&D and Vice Chairman

**CTX-009**

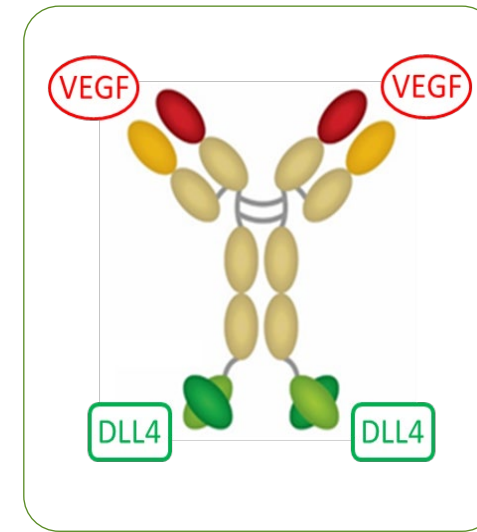
DLL4 X VEGF-A bispecific antibody



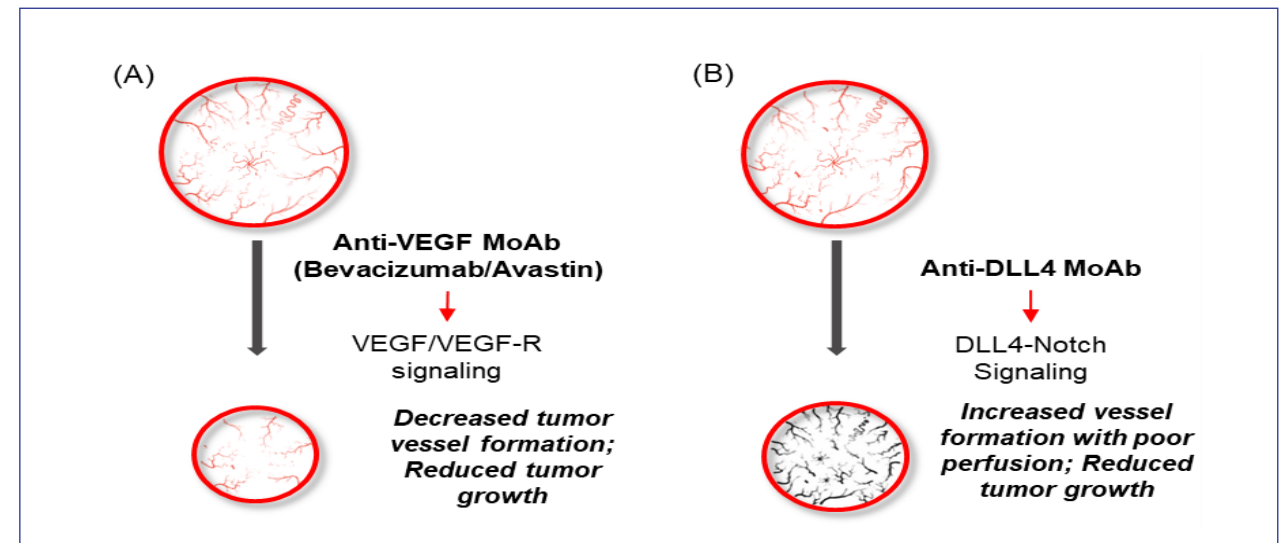


# Overview of CTX-009

- Bispecific antibody blocking DLL4 (Notch-1 ligand) and VEGF-A (soluble ligand)
- Does not lead to ADCC, Fc inactive
- Binds to its targets with 2:2 valency
- At 10 mg/Kg, CTX-009 has approximately the same VEGF-A capturing ability as bevacizumab (Avastin)
- The only DLL4 X VEGF bispecific that demonstrated monotherapy activity in the clinic in colorectal and gastric cancer
- Durable responses in patients with cholangiocarcinoma seen in Phase 1b study of CTX-009 in combination with paclitaxel



## Dual blockade of DLL4 and VEGF overcomes VEGF resistance



# CTX-009 – Vision and Potential



## Best-in-class DLL4 x VEGF-A bispecific

- Phase 3 ongoing in BTC
- Phase 2 ongoing in CRC

### Oncology

Has demonstrated compelling activity in the 3<sup>rd</sup> line and 4<sup>th</sup> line settings in patients with Cholangiocarcinoma, Colorectal Cancer, Gastric Cancer and Pancreatic Cancer

Could become front line therapy in multiple solid tumors

Other potential indications based on DLL4 expression such as Ovarian Cancer & Renal Cell

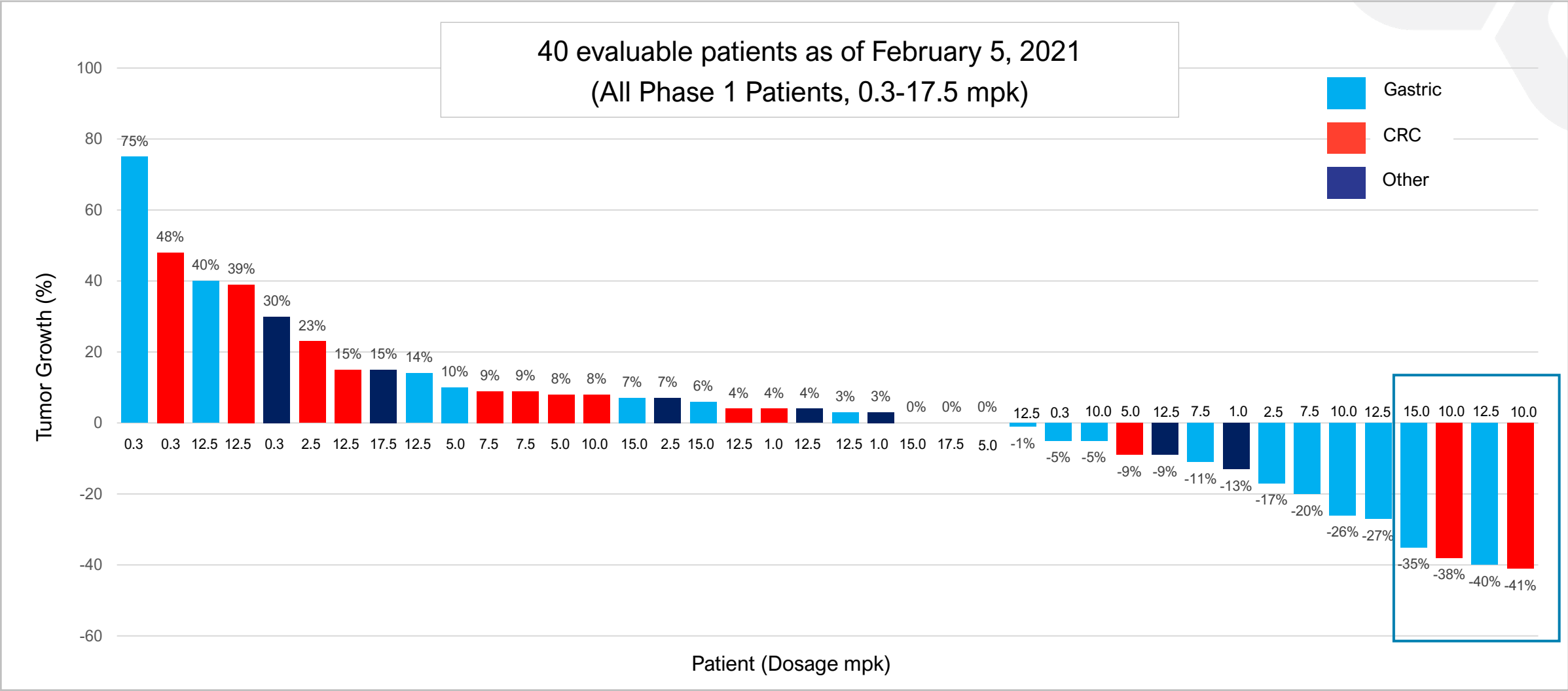
### Ophthalmology

Potential to address AMD and DME based on mechanism

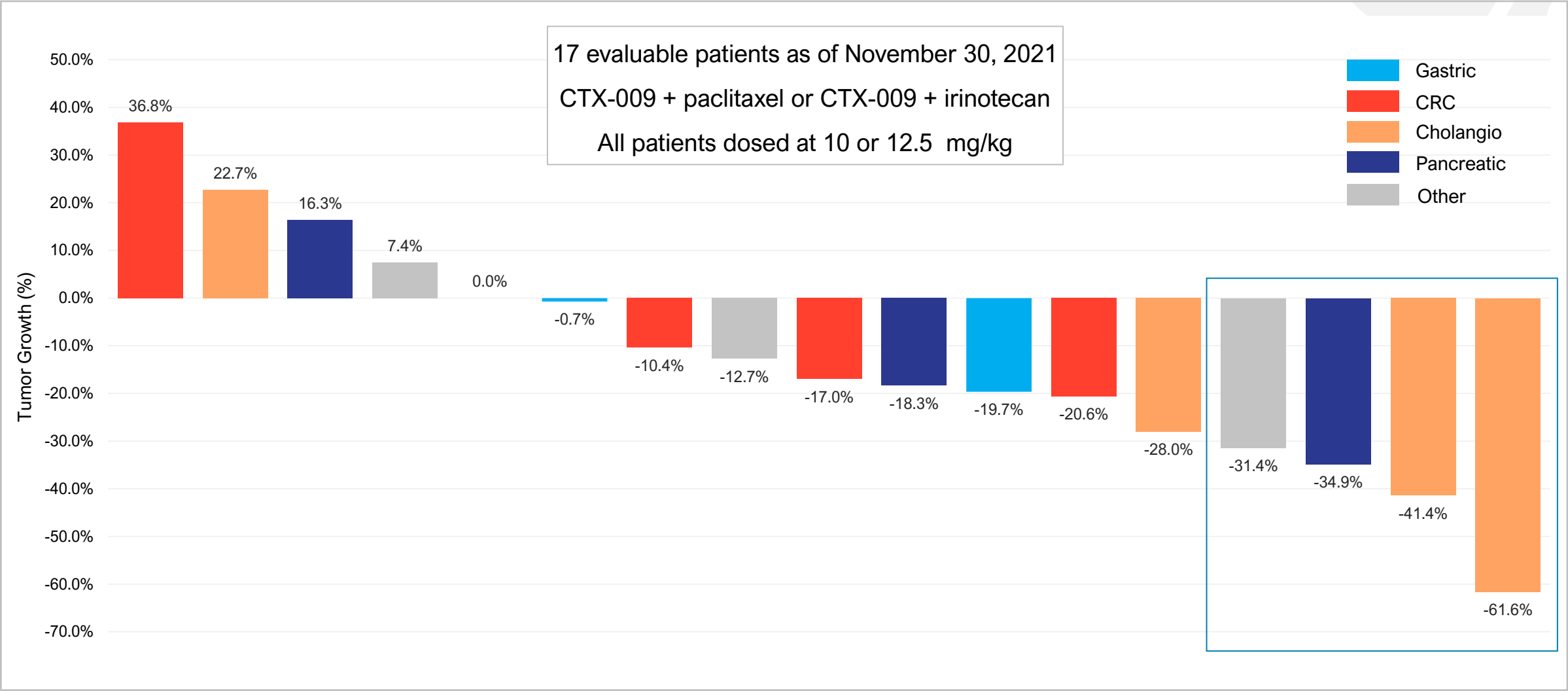
Consideration for partnership



# Phase 1a CTX-009 Monotherapy (all doses)



# Phase 1b CTX-009 Combination Study



# Phase 1 CTX-009 Safety Data

## Phase 1a Monotherapy (n=45)

Drug-related adverse events observed in > 5% of patients	Total (n)	Total (%)	Grade 3 (n)	Grade 3 (%)
Hypertension*	17	38	7	16
General disorders (fatigue, fever, asthenia, edema, etc.)	7	16	1	2
Nervous system disorders (headache, dizziness)	7	16	1	2
Gastrointestinal disorders (nausea, vomiting, etc.)	6	13	2	4
Pulmonary hypertension	4	9	0	0
Proteinuria	3	7	0	0

## Phase 1b Combination (n=17)

Drug-related adverse events observed in > 1 patient	Total (n)	Total (%)	Grade 3 (n)	Grade 3 (%)
Hypertension	8	47	4	24
Nausea	8	47	1	6
Fatigue	6	35	1	6
Neutropenia**	6	35	2	12
Anemia**	4	24	3	18
Thrombocytopenia**	2	12	2	12
Diarrhea	5	29	0	0
Anorexia	5	29	0	0
Proteinuria	5	29	0	0
Pulmonary hypertension (all grade 1)	5	29	0	0
Dyspnea	4	24	0	0
Gingival edema (mucositis)	2	12	0	0
Anal hemorrhage	2	12	0	0

\* In clinical trials of bevacizumab, incidence of Grade 3-4 hypertension ranged between 5%-18% (Avastin label). It is typically managed by anti-hypertensive drugs

\*\*Labeled Grade 3/4 cytopenia events for concomitant chemotherapy agent:

Irinotecan: 31.4% neutropenia, 4.5% anemia, 1.7% thrombocytopenia. Paclitaxel: 52% neutropenia, 16% anemia, 7% thrombocytopenia

# CTX-009 – Phase 1 Clinical Studies Summary

## Overall Response Rate at the Efficacious Dose (10-12.5 mg/kg)

Monotherapy

**18.8% ORR (3/16)**

Combination

**23.5% ORR (4/17)**

## Clinical Benefit Rate at the Efficacious Dose (10-12.5 mg/kg)

Monotherapy

**68.8% (11/16)**

Combination

**76.5% (13/17)**

# Phase 2 CTX-009 Combination Study (S. Korea)

Patients with biliary tract cancers after one or two prior therapies

## Simon 2 Stage adaptive design

### Stage 1

CTX-009 at 10 mg/kg biweekly  
Paclitaxel 80 mg/m<sup>2</sup> weekly 3 of 4 weeks

**N = 21**



**3 or  
more PRs**

### Stage 2

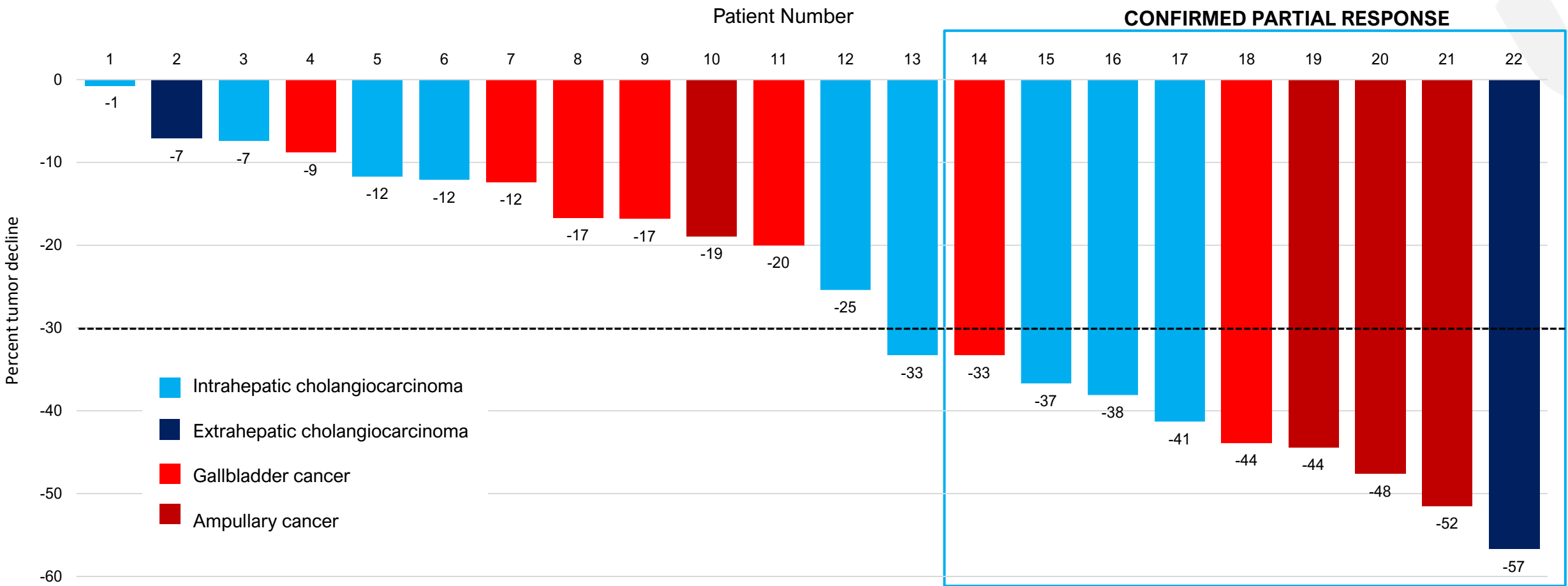
CTX-009 at 10 mg/kg biweekly  
Paclitaxel 80 mg/m<sup>2</sup> weekly 3 of 4 weeks

**N = 45 additional patients**

# Phase 2 CTX-009 Data

Responses achieved across multiple BTC subclasses. Data as of November 9, 2022

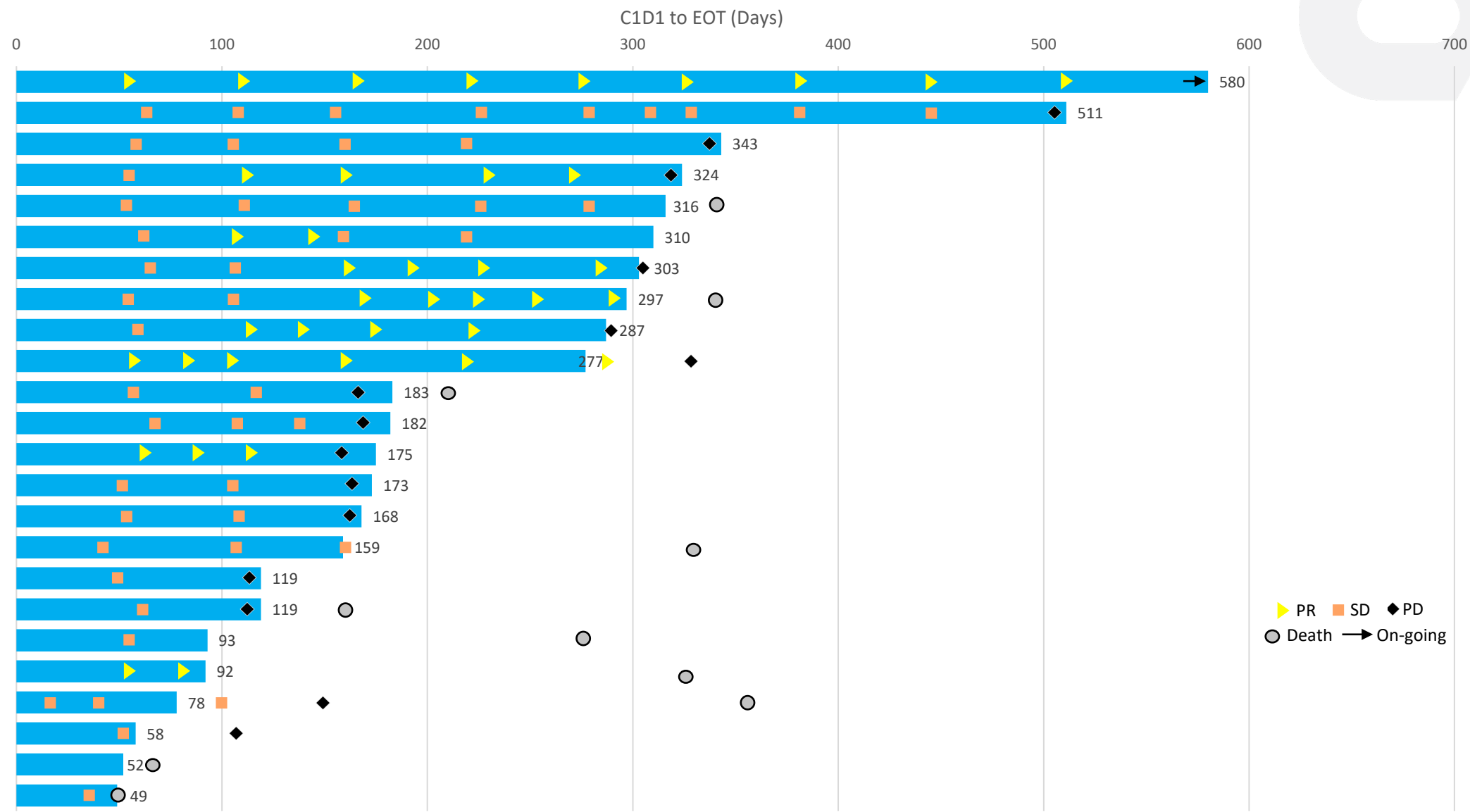
ORR = 37.5%  
CBR = 91.5%



2 patients were not evaluable (one missing post baseline scan; one had a scan outside of protocol window)



# CTX-009 Swimmer Plot



# CTX-009 Phase 2 Results (Median follow-up of 12.1 months)

- 24 patients enrolled and dosed
- 1 patient remains on study

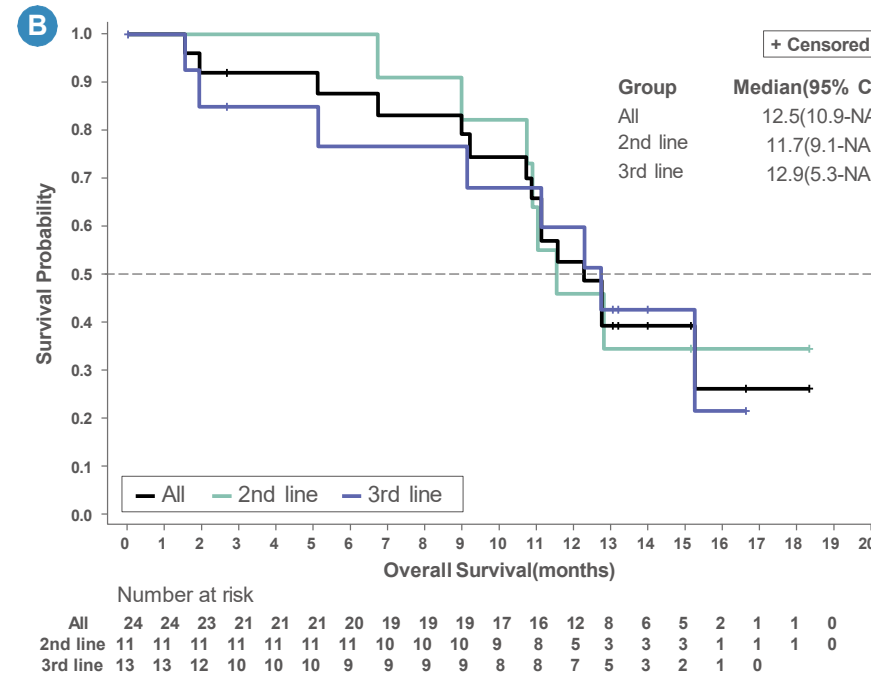
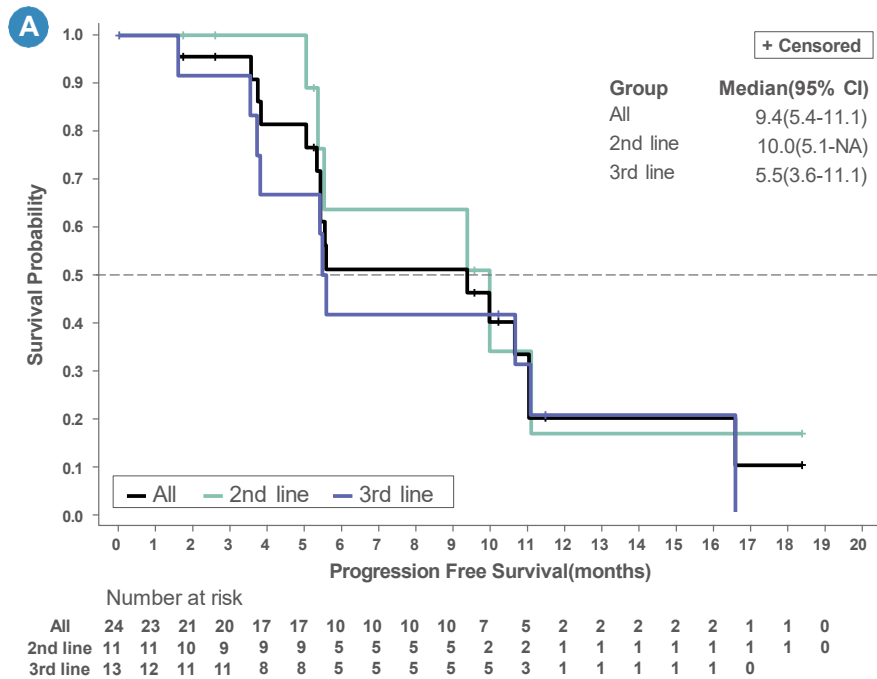
Endpoint	Value (95% CI)
Overall Response Rate (ORR)	37.5%
Stable Disease (SD)	54.2%
Progression Free Survival (PFS)	9.4 m (5.4 – 11.1)
Overall Survival (OS)	12.5 m (10.9 – NA)
Duration of Response	6.9 m (3.5 – NA)

Number of previous systemic therapies	ORR
Pts treated in the 2L [n=11]	7/11 (63.6%)
Pts treated in the 3L [n=13]	2/13 (15.4%)

# Secondary Endpoints: PFS and OS

- Median PFS: 9.40 m (5.4-11.1)

- Median OS: 12.5 m (10.9-NA)



# Treatment-Emergent $\geq$ Grade 3 Adverse Events (>10% of patients)

## Phase 2 BTC study of CTX-009 plus paclitaxel

Event	24 total Patients N (%)
Neutropenia	20 (83.3%)
Anemia	5 (20.8%)
Hypertension	4 (16.7%)
Thrombocytopenia	3 (12.5%)
TEAE leading to discontinuation: confusion, embolism, pneumonia (grade 5), biliary fistula, large intestine perforation, blood creatinine increased, and blood urea nitrogen increased	

## Bevacizumab and paclitaxel label information

Event	Bevacizumab (label)	Paclitaxel (label)
Neutropenia		52%
Hypertension	5-18%	
Anemia		16%
Thrombocytopenia		7%
	Additional events: GI perforation, wound healing complications, Proteinuria, hemorrhage	Additional events: Hypersensitivity reactions, infections, bleeding, neuropathy

# How Does CTX-009 Data Compared to Other BTC Studies?

Parameter	CTX-009 Mixed 2L and 3L N=24	FOLFOX (ABC-06) <sup>1</sup> Only 2L N=81	Gem/Cis <sup>2</sup> 1L N=204	Gem/Cis + Durv <sup>3</sup> Only 1L N=341
<b>ORR</b>	37.5% [64% 2L; 15% 3L]	5%	26%	26.7%
<b>OS</b>	12.5 m	6.2 m	11.7 m	12.9 m
<b>PFS</b>	9.4 m	4.0 m	8.0 m	7.2 m
<b>Any AE</b>	100%	99%	55%	99.4%
<b>Gr 3/4 AEs</b>	92%	60%	71%	74%
<b>Deaths (as Gr 5)</b>	1 (4%)	10 (12%)	17 (8%)	13 (4%)
<b>AEs leading to discontinuation</b>	25%	~ 12%	10%	13%

# CTX-009 Phase 2 Study Summary

## 24 patients with BTC have been enrolled and dosed

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9 partial responses (PRs) for a 37.5% ORR in patients treated in the second- and third-line settings (**64% ORR** of patients treated in the 2<sup>nd</sup> line setting)

Median PFS 9.4 months

Median OS 12.5 months

Adverse event profile similar to Phase 1 studies

## Other regimens in BTC

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### **FOLFOX** (NCCN guidelines):

5% ORR in the second-line setting

4.0 month median PFS

6.2 month median OS

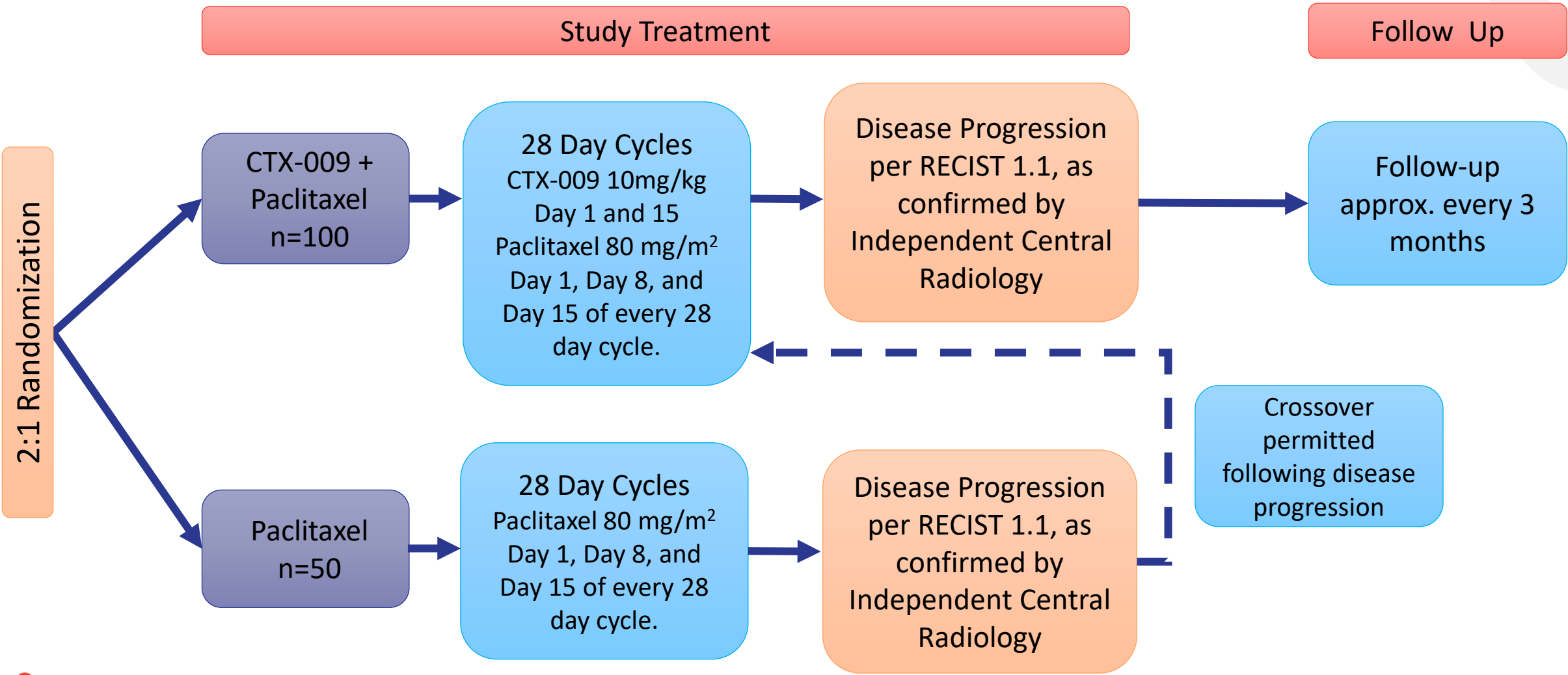
### **TOPAZ-1** (Phase 3 study):

26.7% ORR for Gem/Cis/Durvalumab (anti PD-L1) in the first-line setting



# COMPANION-002: Phase 2/3 U.S. BTC Study

Patients who have received one prior line of therapy



# CTX-009: BTC Patient Demographics and Current Treatments

	US	EU5	Japan	Worldwide
Incident Cases	18,400 <sup>1</sup>	21,800 <sup>2</sup>	14,329 <sup>2</sup>	>200,000 <sup>3</sup>

## 1L Treatment

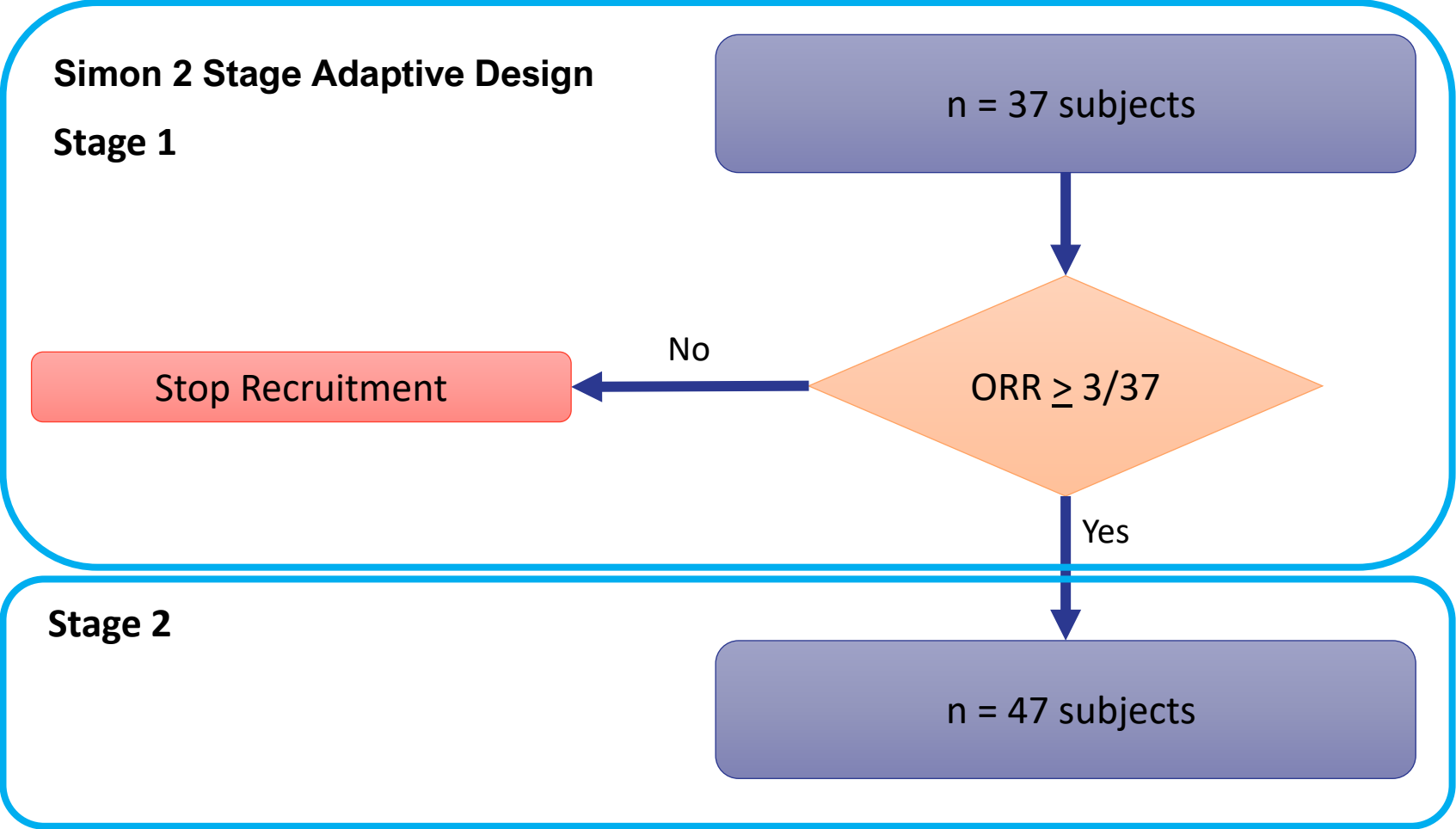
Doublet chemo of gemcitabine +  
cisplatin (ABC-02 study)  
Or  
Gemcitabine/cisplatin + durvalumab  
(recently approved for 1L)

## 2L Treatment

FOLFOX 5% ORR 0.9 Mos OS Δ	<u>FGFR2 mutation</u> Pemigatinib (10-15% of CCA)	<u>IDH1 mutation</u> Ivosidenib (1-3% of BTC)	<u>MSI-H tumors</u> PD-1 Inhibitor (<1% of BTC)	Clinical trial
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1. NCI Surveillance, Epidemiology, and End Results (SEER) program  
2. Delveinsight/company estimates  
3. International Agency for Research on Cancer/GLOBOCAN

# COMPANION-003: Phase 2 U.S. Colorectal Cancer (CRC) Study



# CTX-009: CRC Patient Demographics and Current Treatments

	US	EU5	Japan	Worldwide
Incident Cases	153,020 <sup>1</sup>	246,734 <sup>2</sup>	148,505 <sup>2</sup>	1,931,590 <sup>2</sup>
~50% Metastatic <sup>3</sup> 50-70% reach 3L <sup>4</sup>	38,000-53,000 patients			

## 1L Treatment

Chemotherapy  
FOLFOX/FOLFIRI

Bevacizumab or  
EGFR inhibitor +  
chemotherapy

Anti-PD-1 with  
MSI-H/dMMR  
mutation

~5% of CRC

## 2L Treatment

Bevacizumab or  
EGFR + chemo

BRAF/EGFR with  
V600E mutation

5-8% of CRC

## 3L Treatment

Regorafenib

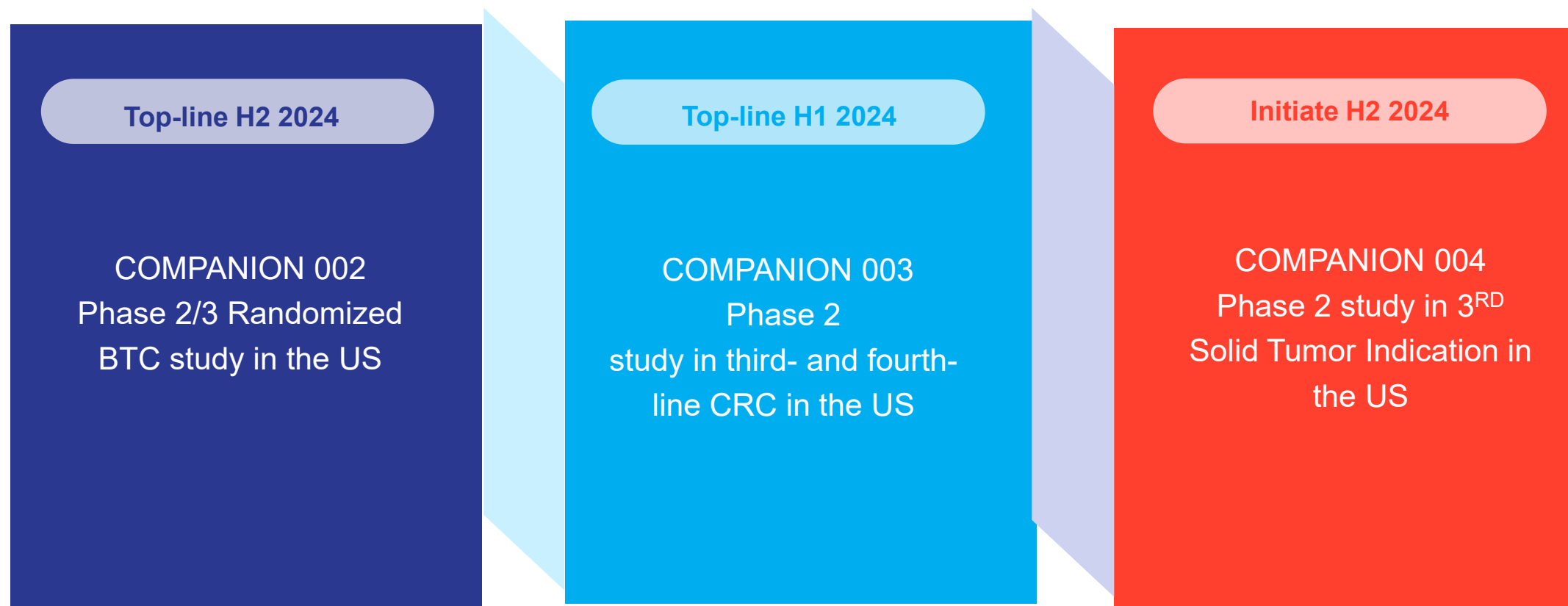
ORR 1%, Median  
PFS 2.0 months

Trifluridine/  
tipiracil

ORR 1-2%  
Median  
PFS ~2 months

1. NCI Surveillance, Epidemiology, and End Results (SEER) program  
2. International Agency for Research on Cancer/GLOBOCAN  
3. L Biller, D Schrag , JAMA 2021 Feb 16  
4. Bekaii-Saab, Clin advances in Hem and Onc, Supp Jan 2021

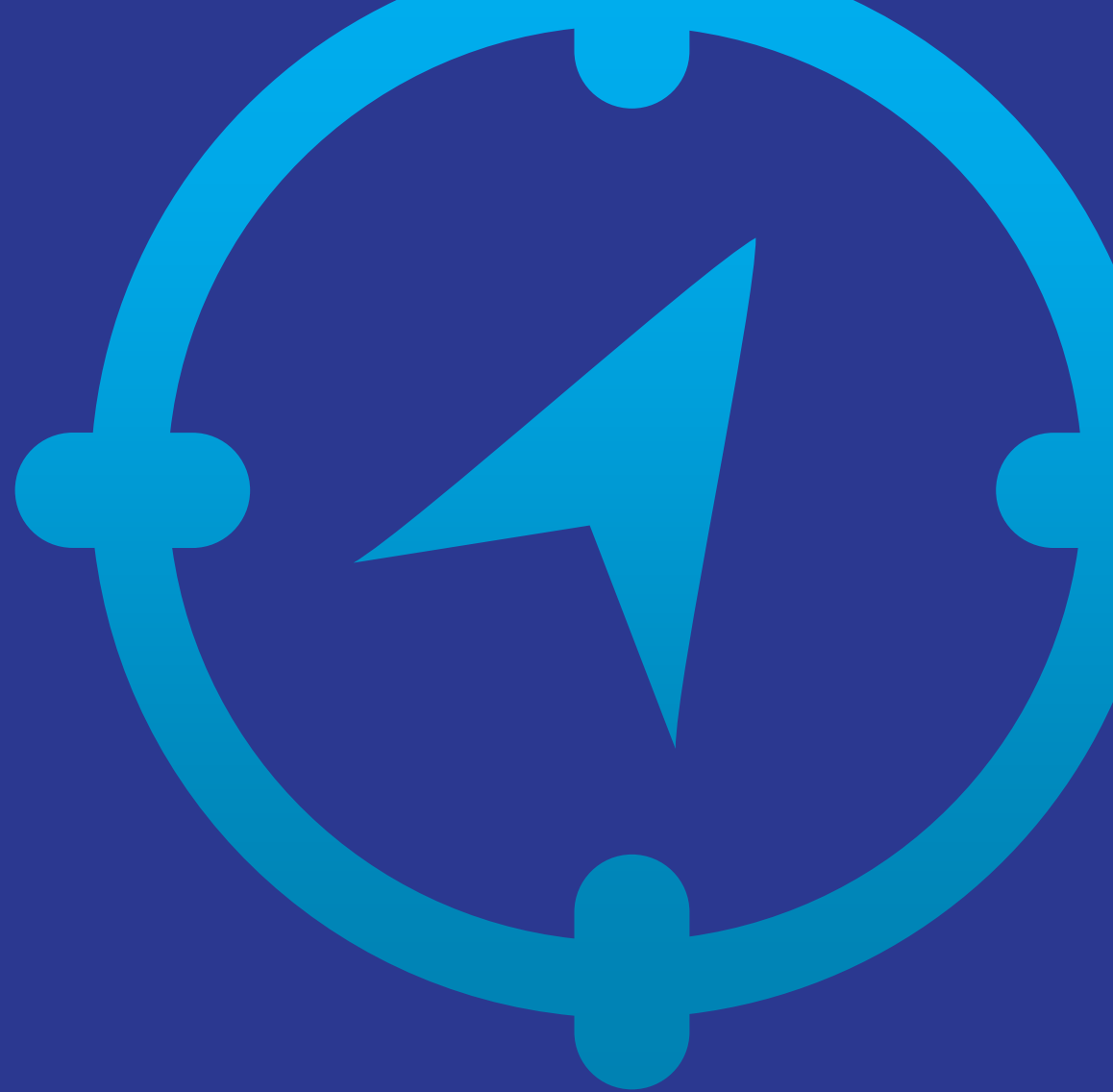
# The COMPANION (COMPASS ANTI-ANGIOGENESIS) Studies



Evaluating additional indications for CTX-009 both as a monotherapy and in combination with chemotherapy

**CTX-471**

CD137 monoclonal antibody





# CTX-471: Potential Best-in-Class CD137 Agonist

## CTX-471: next generation CD137 agonist

Fully human, IgG4, optimized affinity for agonistic antibody

Unique epitope: non-ligand blocking

## Phase 1 Study Update

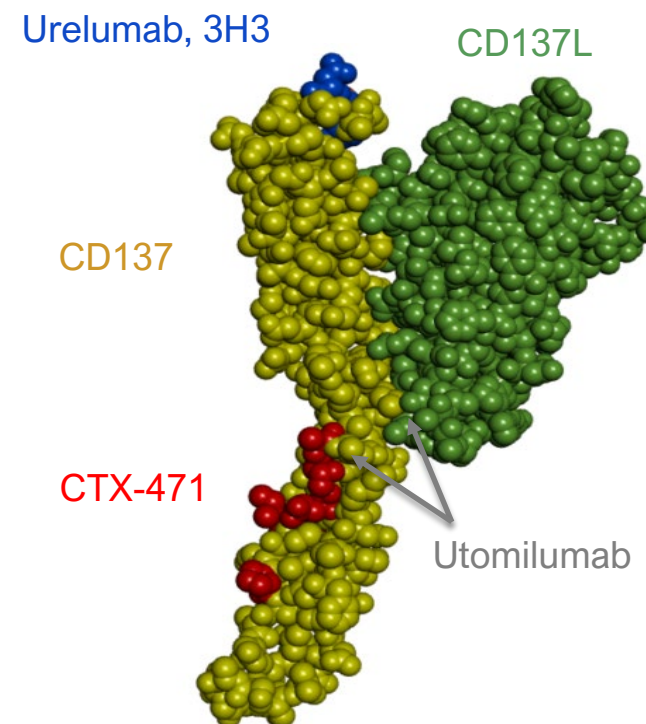
Monotherapy in post checkpoint inhibitor patients

Monotherapy Phase 1a multiple ascending dose study completed

- MTD defined by immune thrombocytopenia

Monotherapy Phase 1b dose expansion study completed

- 1 CR: small cell lung cancer
- 4 PRs observed: melanoma (3 of 11) and mesothelioma (1 of 4)

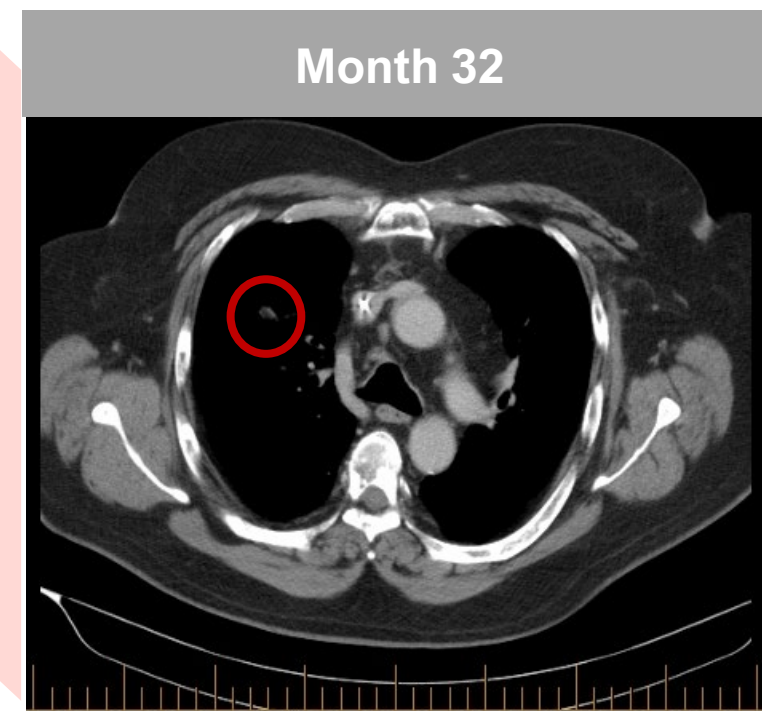
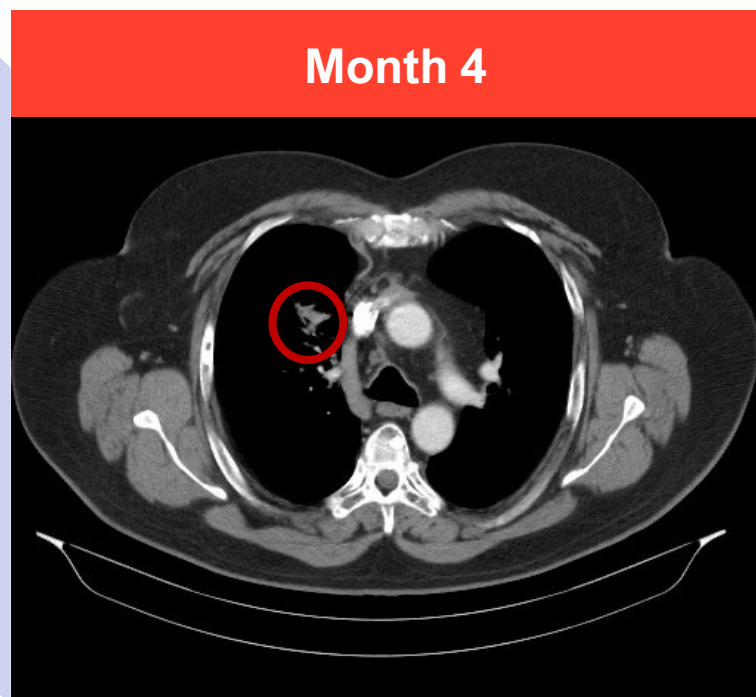
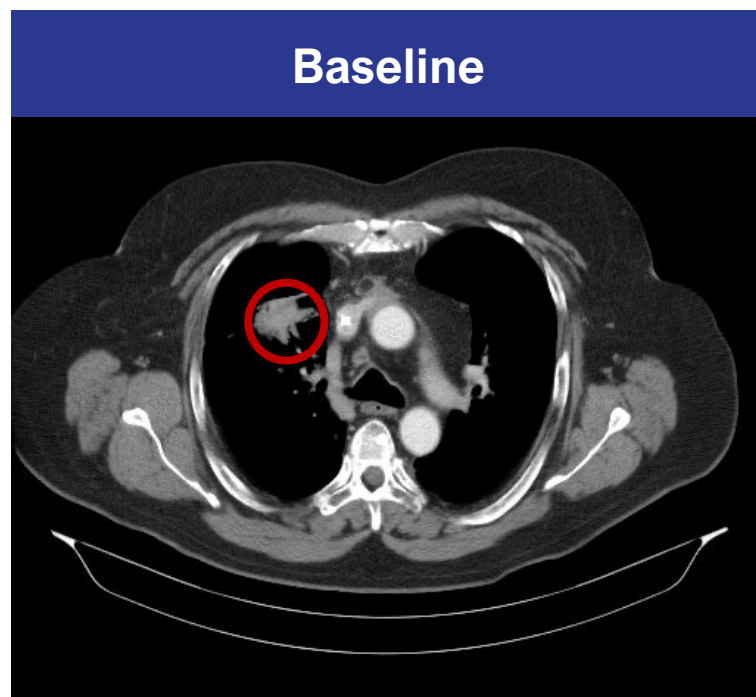


*JCI Insight.* 2020;5(5):e133647

# CTX-471: Complete Response in a Patient with Small Cell Lung Cancer

After progression on atezolizumab/chemo and nivolumab

- » 66 y.o. man with advanced SCLC: Carboplatin/etoposide plus atezolizumab first line; nivolumab second line
- » Confirmed, complete response (CR) by PET ~ 3 years on therapy



# CTX-471 Clinical Development Plans

## Phase 1b Monotherapy Study

Generally well tolerated

A complete response and four partial responses in the post PD-1/PD-L1 patient population

Small cell lung cancer, mesothelioma, and melanoma (three patients)

## Phase 1b of CTX-471 with KEYTRUDA<sup>®</sup> in collaboration with Merck

Abbreviated CTX-471 dose escalation into the PD-1 regimen, followed by a cohort expansion

Post PD-1/PD-L1 Salvage Study

Dose escalation complete, no DLT

Dose expansion is ongoing

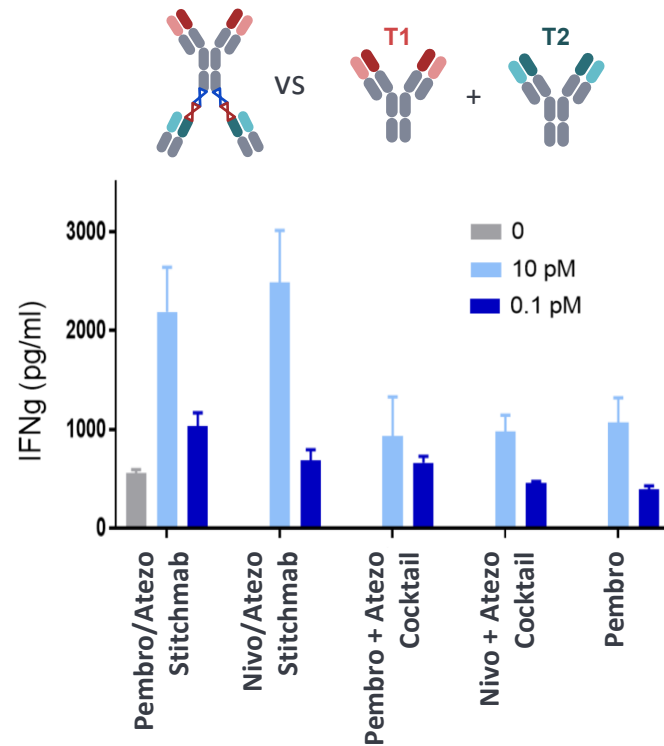
# CTX-8371

PD-1 x PD-L1 bispecific antibody



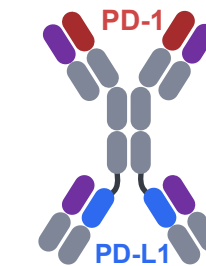
# StitchMabs™ Platform was Utilized to Identify CTX-8371

Unexpected synergistic activity of PD-1/PD-L1 combination in bispecific Stitchmab format

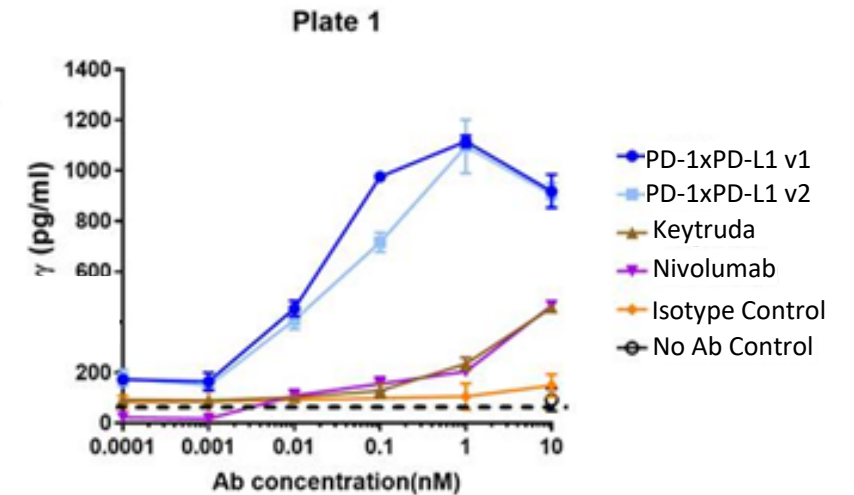


Mixed lymphocyte reaction (MLR) assay

Common Light Chain bispecifics were generated to test therapeutic hypothesis



Our PD-1xPD-L1 bispecifics observed to outperform PD-1 blockers in T-cell activation assay

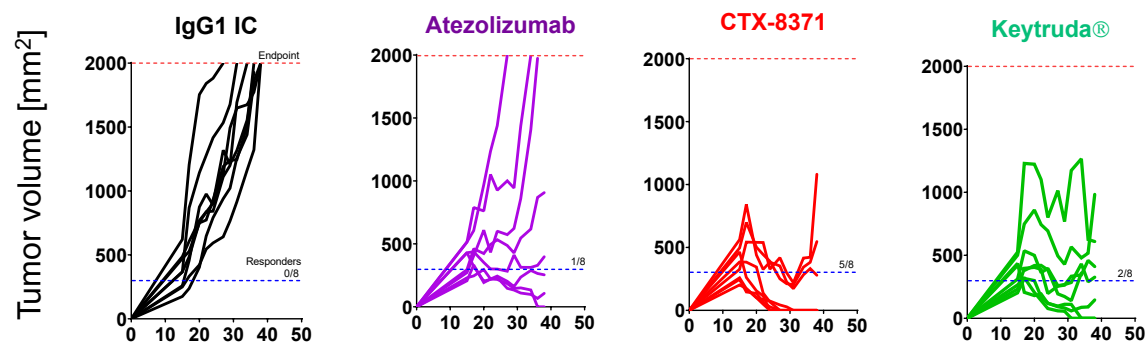
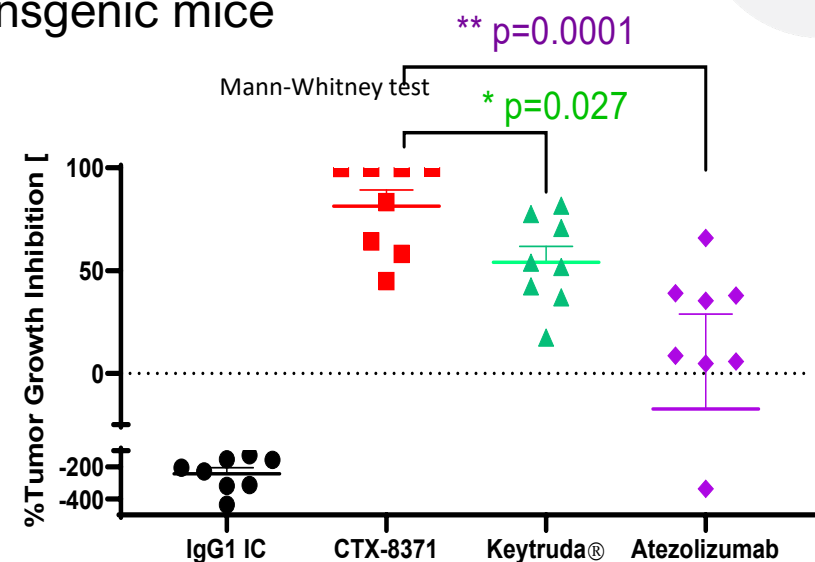
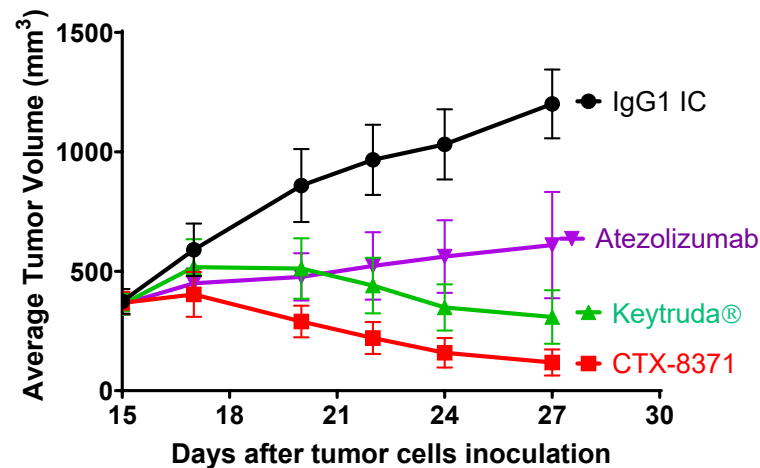






# CTX-8371 Pre-Clinical Proof of Concept

Activity in MC38-hPD-L1 model implanted in hPD-1/hPD-L1 transgenic mice



Group	% Cured	Tumor free / total
CTX-8371	62.5	5/8
Atezolizumab	12.5	1/8
IgG1 IC	0	0/8
Keytruda	25	2/8

# CTX-8371: Development Status

IND was accepted

Currently opening clinical sites

First patient dosing expected in  
1Q 2024

## Phase 1 study design

Multiple ascending dose, dose-escalation study

5 doses planned: 0.1, 0.3, 1.0, 3.0, and 10 mg/kg

Post PD-1 or PD-L1 patient population: Melanoma, NSCLC, HNSCC, Hodgkin's Lymphoma, TNBC

First patient dosing targeted for Q1 2024

Potential for proprietary combination regimens with CTX-009 and CTX-471

# Compass Therapeutics

## Summary



# Program Summary

## »»» **CTX-009** Novel DLL4 x VEGF-A bispecific antibody with both combination and monotherapy activity

Phase 1: Dose response established – responses in multiple indications

BTC Phase 2 results: 24 patients: 37.5% ORR (2L/3L), 63.6% (2L), median PFS 9.4 months, OS 12.5 months

COMPANION-002: BTC Phase 3 randomized study ongoing; top-line data expected H2 2024

COMPANION-003: CRC Phase 2 monotherapy study ongoing; top-line data expected mid-2024

## »»» **CTX-471** Potential best-in-class CD137 agonist antibody with monotherapy activity

Phase 1 monotherapy study complete:

1 complete response (CR): small cell lung cancer (1 of 3), and 4 partial responses (PRs) in post PD-1 population: metastatic melanoma (3 of 11) and mesothelioma (1 of 4)

CTX-471 in combination with KEYTRUDA® dose escalation complete, dose expansion ongoing

## »»» **CTX-8371** Next generation PD-1 x PD-L1 bispecific antibody

Unique MOA – enhances T-cell activation

IND cleared, currently opening clinical sites

# Key 12 Month Milestones

