## 0 COMPASS

## Compass Therapeutics Presentation



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## Company Overview

## We are a clinical stage biotech company developing antibody therapeutics for cancer \& autoimmune diseases

## CORE SCIENCE

- StitchMabs ${ }^{\text {TM }}$ platform designed to identify synergistic bispecific antibodies
- Common Light Chain technology enables multi-specificity and manufacturability
- Our understanding of Cancer Biology informs the design of our proprietary drug candidates


## FOCUS

- Next Generation novel antibody therapeutics
- Key programs: CTX-009, CTX-471 and CTX-8371
- Let the science guide the therapeutic format: monoclonal vs. multispecific


## RESOURCES

- ~25 FTEs, based in Boston, MA
- Capable of rapidly advancing candidates from idea to clinical proof-of-concept
- Funded by leading life-science investors


## Seasoned Leadership Team

Senior Executive Team


Board of Directors $\qquad$

## Carl L. Gordon, Chair

OrbiMed
## Ellen Chiniara

Phil Ferneau


Mary Ann Gray
Independent Board Member
Alexion

Peter Moesta Interim Head of CMC



Stephen Squinto
(8) OrbiMed

## Compass Pipeline



## Key Events Expected in the Next 24 Months

| CTX-009 |
| :---: |
| DLL4 x VEGF-A + Taxol |
| Biliary Tract Cancer |
| CTX-009 |
| DLL4 x VEGF-A |
| Advanced Colorectal |
| CTX-009 |
| DLL4VEGF-A + Chemo <br> Ovarian |



| CTX-471+ PD-1 |
| :---: |

Post PD-1/PD-L1

## CTX-8371

PD-1 x PD-L1
Multiple Indications

## Targeted Therapy

## CTX-009 - DLL4 X VEGF-A bispecific antibody

## CTX-009: Vision and Potential

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

$>$ Phase $2 / 3$ ready, with parallel active development in S. Korea and China - potential to turn studies into global studies

## $>$ Oncology:

- Has demonstrated compelling activity in the $3^{\text {rd }}$ line and $4^{\text {th }}$ line settings in Cholangiocarcinoma, Colorectal Cancer, Gastric Cancer and Pancreatic Cancer
> Ovarian Cancer - strong potential based on DLL4 expression and other agents
- Could become front line therapy in multiple solid tumors
- Other indications based on DLL4 expression
> Ophthalmology:
- Potential to address AMD and DME based on mechanism
- Consideration for partnership


## CTX-009 - Phase 1 Clinical Summary

> Two Phase 1 Studies (S. Korea)

- Ph 1a dose-escalation monotherapy, including cohort expansion at projected RP2Ds
$>$ Ph 1b combination study with irinotecan and paclitaxel
- Safety: well-tolerated; MTD has not been determined
- Activity: 8 PRs, 6 confirmed by RECIST in 33 advanced solid tumor patients treated at the therapeutic doses

Responses as a monotherapy: colorectal and gastric cancers
Responses in combination with chemotherapy: cholangiocarcinoma, pancreatic
Cholangio ORR=50\%; Clinical benefit rate $=75 \%$ with a median duration of response of 9.7 months

## Phase 1a Monotherapy Waterfall Plot (all doses)



## Phase 1a Monotherapy Safety Data (n=45)

| Drug-related adverse events | Total <br> (n) | Total <br> $(\%)$ | Grade 3 <br> $(\mathrm{n})$ | Grade 3 <br> $(\%)$ |
| :--- | :---: | :---: | :---: | :---: | :---: |
| Hypertension* | 17 | 37.8 | 7 | 15.6 |
| General disorders (fatigue, fever, asthenia, <br> edema, etc.) | 7 | 15.6 | 1 | 2.2 |
| Nervous system disorders (headache, <br> dizziness) | 7 | 15.6 | 1 | 2.2 |
| Gastrointestinal disorders (nausea, <br> vomiting, etc.) | 6 | 13.3 | 2 | 4.4 |
| Pulmonary hypertension | 4 | 8.9 | 0 | 0 |
| Proteinuria | 3 | 6.7 | 0 | 0 |

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


## Phase 1b Combination Study

P Phase 1b: combination study with chemotherapy ( $\mathrm{N}=17$ )
> 4 arms:
> 10.0 and $12.5 \mathrm{mg} / \mathrm{kg}$ CTX-009
> Irinotecan or paclitaxel
> Activity:
>4 PRs, 3 confirmed, including a confirmed PR in pancreatic cancer; PR rate: 23.5\%
> 9 SD; SD rate: 52.9\%
> Overall Response Rate (ORR): 23.5\%

- Clinical Benefit Rate (CBR): 76.5\%
- Phase 2a combination study in BTC has began


## Phase 1b Combination Study Waterfall Plot


80.0\%
70.0\%
60.0\%
50.0\%
$40.0 \%$
$30.0 \%$

- $30.0 \%$
20.0\%
10.0\%
0.0\%
$-10.0 \%$
-20.0\%
30.0\%
-40.0\%
-50.0\%
-60.0\%
-70.0\%
-80.0\%

$0.0 \%$


Waterfall of CTX-009 (ABL001) Combination study as of November 30, 2021
-61.6\%
(All Phase 1b Patients, $10 \mathrm{mg} / \mathrm{kg}$ or $12.5 \mathrm{mg} / \mathrm{kg}$ )

## CTX-009 - Phase 1 Studies Clinical Summary

## Overall Response Rate at the Efficacious Dose <br> ( $10-12.5 \mathrm{mg} / \mathrm{kg}$ ):

- Monotherapy: 18.8\% ORR (3/16)
- Combination: 23.5\% ORR (4/17)

Clinical Benefit Rate at the Efficacious Dose ( $10-12.5 \mathrm{mg} / \mathrm{kg}$ ):

- Monotherapy: 68.8\% (11/16)
- Combination: 76.5\% (13/17)


## Phase 2 Combination Study: CTX-009 Plus Paclitaxel

Phase 2 Study Design:

- Patients with biliary tract cancers after one or two prior therapies
- CTX-009 at $10 \mathrm{mg} / \mathrm{kg}$ biweekly plus paclitaxel $80 \mathrm{mg} / \mathrm{m}^{2}$ weekly 3 of 4 weeks
- Simon 2 Stage adaptive design:
- Stage 1: 21 patients $\rightarrow$ ORR

Stage 2: if 3 or more PRs $\rightarrow$ Stage 2: 45 additional patients

## Phase 2: Patient Baseline and Demographics

|  | 24 Total Patients |
| :--- | :--- |
| Age |  |
| Median (years) | 61.5 |
| Gender, $n(\%)$ <br> Male | $14(58 \%)$ |
| Female | $10(42 \%)$ |
| ECOG performance status, $n(\%)$ |  |
| 0 | $13(54 \%)$ |
| 1 | $11(46 \%)$ |


|  | 24 Total Patients |
| :--- | :--- |
| Prior systemic therapies, $\mathrm{n}(\%)$ |  |
| 1 | $11(46 \%)$ |
| 2 | $13(54 \%)$ |
| Prior Gem/Cis regimen | $23(96 \%)$ |
| BTC subtype, $\mathrm{n}(\%)$ |  |
| Intrahepatic cholangiocarcinoma | $9(38 \%)$ |
| Extrahepatic cholangiocarcinoma | $3(13 \%)$ |
| Gallbladder cancer | $7(29 \%)$ |
| Ampullary cancer | $5(21 \%)$ |

Phase 2 Waterfall: ORR = 42\%; CBR = 92\%


## Swimmer Plot: Median Time on Study ~ 6 Months



## Safety Data: Treatment-Related $\geq$ Grade 3 Adverse Events

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

| Event | 24 total Patients N(\%) |
| :--- | :---: |
| Neutropenia | $12(50.0 \%)$ |
| Hypertension | $4(16.7 \%)$ |
| Anemia | $3(12.5 \%)$ |
| Thrombocytopenia | $2(8.3 \%)$ |

Additional events observed in 1 patient: Intestinal perforation, Asthenia, Catheter site hemorrhage, Fatigue, Cholangitis, Abdominal infection, Bacterial gastritis, Pneumonia (fatal), Postprocedure hemorrhage, Decreased appetite, Cerebral hemorrhage, Proteinuria, Embolism

## Avastin and paclitaxel label information

| Event | Avastin (label) | Paclitaxel (label) |
| :--- | :--- | :---: |
| Neutropenia | $5-18 \%$ | $52 \%$ |
| Hypertension |  |  |
| Anemia |  |  |
| Thrombocytopenia |  | 7\%\% |
|  | Additional events: <br> Gl perforation, <br> wound healing <br> complications, <br> Proteinuria, <br> hemorrhage | Additional events: <br> Hypersensitivity <br> reactions, <br> infections, bleeding, <br> neuropathy |

## CTX-009: Interim Phase 2 Study Summary and Next Steps

> 10 partial responses (PRs) for a 42\% ORR in patients treated in the second- and third-line settings
> Median time on study approximately 6 months, with 7 patients ongoing

- Adverse event profile similar to Phase 1
> Other regimens in BTC:
- FOLFOX (NCCN guidelines): 5\% ORR in the second-line setting
- TOPAZ-1 (Phase 3 development): $26.7 \%$ ORR for Gem/Cis/Durvalumab (anti-PD-L1) in the firstline setting
> Initiate Stage 2 of the Phase 2 BTC study in the US in early Q3
> Initiate Phase 2/3 study in patients with colorectal cancer in the third line setting in the US in Q4 2022


## T Cell and NK Cell Agonist

## CTX-471 - CD137

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

NOVEL EPITOPE WITH DIFFERENTIATED ACTIVITY OBSERVED IN EXTENSIVE PRECLINICAL DATA*

## > CTX-471: next generation CD137 agonist

- Fully human, IgG4, optimized affinity for agonistic antibody
> Unique epitope: non-ligand blocking


## - Phase 1 Study Update



## Partial 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
> Multiple metastases: Largest mass (RUL Lung) shown below, $\sim 4 \mathrm{~cm}$ at baseline $\rightarrow 40 \%$ total decline
> Confirmed and durable PR at Month 18


Month 4


Month 8


## CTX-471: Clinical Development Plans

- Phase 1b study nearing completion
- Well tolerated
- Three partial responses as a monotherapy agent in the post PD-1/PD-L1 patient population
> Small cell lung cancer and melanoma (two patients)
- Initiate a Phase 1b combination study with a PD-1 inhibitor
- Abbreviated CTX-471 dose escalation into the PD-1 regimen, followed by a cohort expansion
- Post PD-1/PD-L1 Salvage Study
- Progression on a labeled PD-1/PD-L1 regimen
- Followed by PD-1 salvage plus CTX-471


## Bispecific Checkpoint Blocker

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

## StitchMabs ${ }^{\top M}$ Platform was Utilized to Identify CTX-8371:



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 Activity in MC38-hPD-L1 Model Implanted in hPD-1/hPD-L1 Transgenic Mice






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

[^0]
## CTX-8371: Development Status

> IND enabling studies underway

- Primate dose range finding study completed $\rightarrow$ PD-1 shedding confirmed in vivo
- Manufacturing initiated
- Pre-IND meeting completed
> Phase 1 study planning
- Multiple ascending dose dose-escalation study
> 5 doses planned: 0.1, $0.3,1.0,3.0$, and $10 \mathrm{mg} / \mathrm{kg}$
- Post PD-1 or PD-L1 patient population
- PD-1 shedding on peripheral T cells as a pharmacodynamic marker
> IND submission targeted for Q1 2023
> Potential for proprietary combination regimens with CTX-009 and CTX-471


## Key Milestones Expected Across Our Portfolio in 2022




[^0]:    Days post treatment

