



DEFEATING
CANCER:
The Challenge.
Our Mission.

December 2020

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Executing on Our Mission

At Deciphera, we are focused on discovering, developing, and bringing important new medicines to patients for the treatment of cancer.



Successfully **launch** QINLOCK® (riporetinib) in the U.S.



Drive the development of QINLOCK in **2nd line GIST** (INTRIGUE)













Rapidly advancing wholly-owned clinical-stage portfolio⁽¹⁾



Novel switch control kinase inhibitor discovery platform **fuels the pipeline**

Robust Pipeline of Novel Switch-Control Kinase Inhibitors

		PRECLINICAL	PHASE 1	PHASE 1B/2	PHASE 3	REGULATORY SUBMISSION	APPROVED	COMMERCIAL RIGHTS
QINLOCK (ripretinib) 50 mg tablets Broad Spectrum Inhibitor of KIT & PDGFRA	GIST ≥4 th -Line (INVICTUS Study)						 ⁽¹⁾  ⁽²⁾   	 ⁽³⁾
	GIST 2 nd -Line (INTRIGUE Study)							
DCC-3014 Selective Inhibitor of CSF1R	Tenosynovial Giant Cell Tumor							
Rebastinib Selective Inhibitor of TIE2	Solid Tumors in Combination with Paclitaxel (includes breast, ovarian and endometrial cancers)							
	Solid Tumors in Combination with Carboplatin (includes mesothelioma, ovarian and breast cancers)							
DCC-3116 Selective Inhibitor of ULK	Autophagy Inhibitor for Targeting RAS Mutant Cancers							
Additional Programs	Undisclosed							



2020

Expected Milestones for the Year Ahead

QINLOCK
(ripretinib) 50 mg tablets

- ✓ FDA approval and U.S. commercial launch in 4th line GIST
- ✓ Present Phase 1 study expansion data
- ✓ Submit EU Marketing Authorisation Application to EMA
- ✓ Complete enrollment in the INTRIGUE Phase 3 study in 2nd line GIST

DCC-3014

- ✓ Select Phase 2 dose for tenosynovial giant cell tumor (TGCT) and initiate the expansion portion of study
- ✓ Update Phase 1 data in TGCT patients

Rebastinib

- ✓ Select Phase 2 dose and activate Part 2 of Phase 1b/2 study in combination with carboplatin
- ✓ Present Phase 1b/2 data in combination with paclitaxel
- ✓ Present Phase 1b/2 data in combination with carboplatin

DCC-3116

- Submit IND application to FDA

QINLOCK®: Now Approved in the United States



U.S. FDA Approval & Commercial Launch

- Approved by U.S. FDA in 4th line GIST on May 15, 2020
- Net sales in first full quarter of \$14.7MM (3Q20); \$19.5MM (launch to 3Q20)

Multiple Global Regulatory Filings and Approvals

- Approved by Health Canada (June 2020) and Australian TGA (July 2020)
- Accepted NDA to China National Medical Products Administration (July 2020)¹
- Submitted MAA to European Medicines Agency (Validated 4Q20)

Ongoing Global Phase 3 Study in 2nd Line GIST (INTRIGUE)

- Completed target enrollment of 426 patients
- Top-line data expected in 2H21

QINLOCK is the first approved TKI designed specifically for GIST regardless of patients' mutational status

QINLOCK[®]: 4 Strategic Objectives For Launch

1	Educate and raise awareness	<ul style="list-style-type: none">• Extensive mutational heterogeneity drives resistance to established therapies
2	Differentiate QINLOCK[®]	<ul style="list-style-type: none">• Novel switch control mechanism of action• Potent inhibition of broad spectrum of mutations <i>in vitro</i>• Potentially practice-changing efficacy• Favorable tolerability with low dose modifications due to AEs
3	Reach and impact GIST prescribers	<ul style="list-style-type: none">• Academic centers of excellence• Community practices
4	Optimize patient access	<ul style="list-style-type: none">• Deciphera AccessPoint[™]• Comprehensive patient support programs and resources

QINLOCK® U.S. Launch Success in 3Q20

Net Sales First
Full Quarter
(3Q20)
\$14.7
MILLION

Unique
Prescribers
(Launch-to-Date)
250+

Unique
Institutions
(Launch-to-Date)
200+

Initial Launch Details

- Continued strong demand, new prescriber growth, and broad access
- New prescribers more than doubled during the quarter
- Strong awareness, product perceptions, and intent to prescribe among GIST treaters with 4th line patients

*Community
Accounts*



Prescribers





Patients

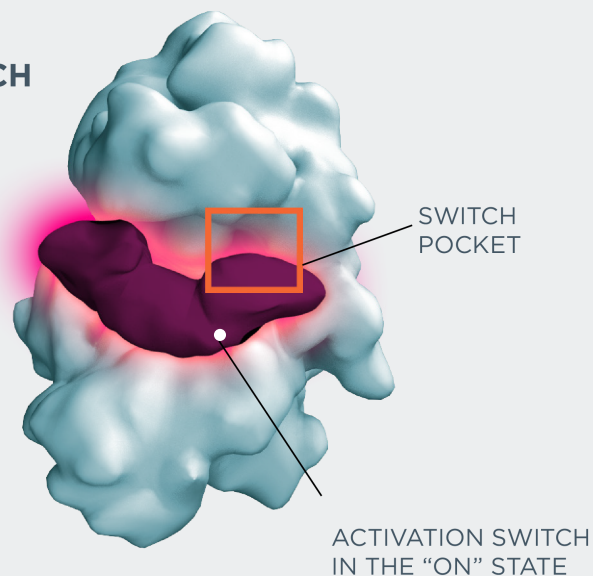
QINLOCK®: A Novel Switch-Control Kinase Inhibitor Engineered to Block the Drivers of Resistance in Advanced GIST

Switched on: Kinase active

Kinase activation requires the interaction of two critical regions:


-  **ACTIVATION SWITCH**
-  **SWITCH POCKET**


TYROSINE KINASE



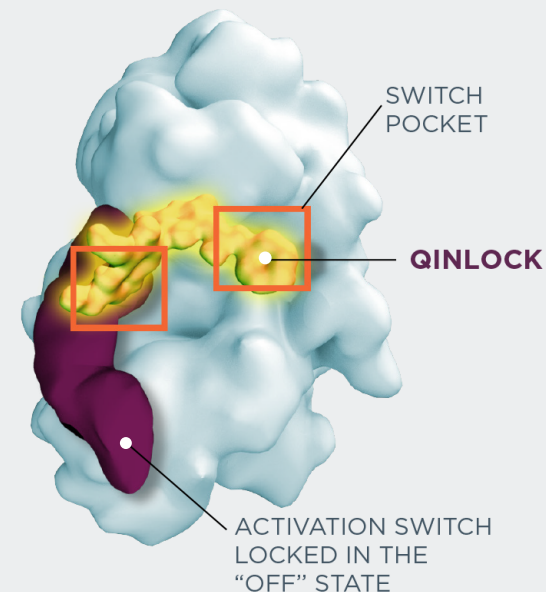
Switched off: Kinase inactive

As shown in preclinical studies, QINLOCK

-  **BINDS**
to both the activation switch and switch pocket, regardless of where mutations arise

-  **LOCKS**
the kinase in the inactive ("off") state, inhibiting downstream signaling and cancer cell proliferation

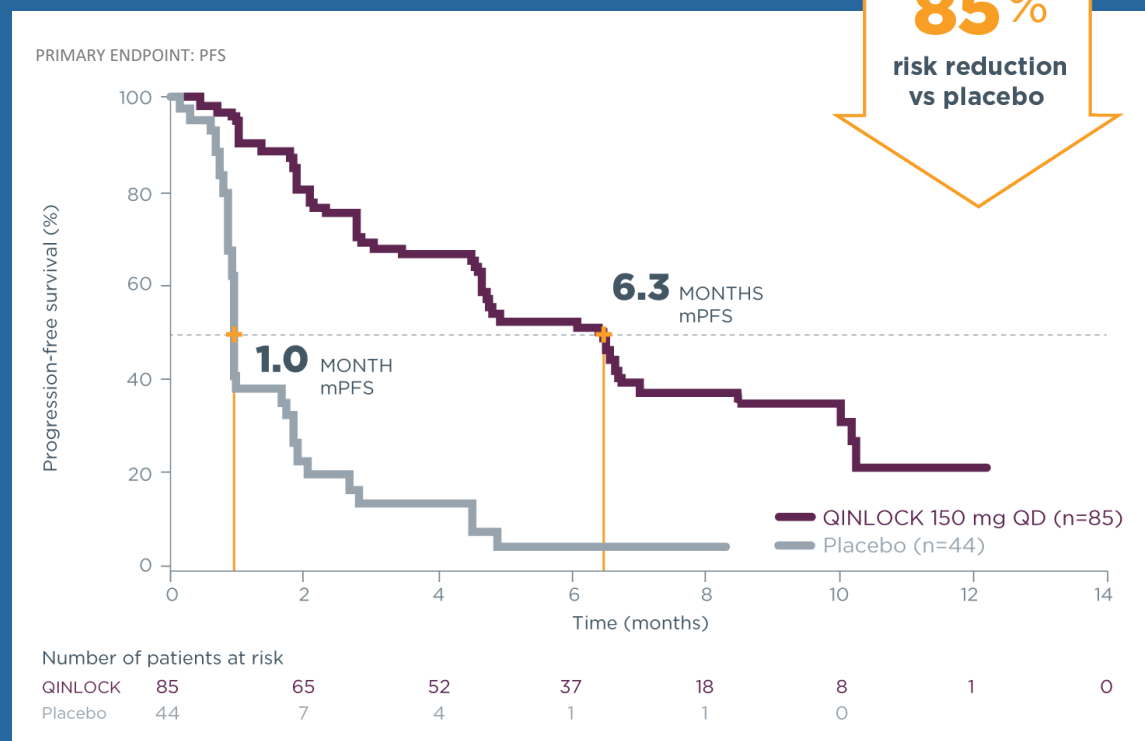
In vitro studies not designed to assess clinical efficacy



The dual MOA of QINLOCK provided broad-spectrum inhibition of KIT and PDGFR α kinase signaling *in vitro*, including multiple primary and secondary mutations and wild type GIST

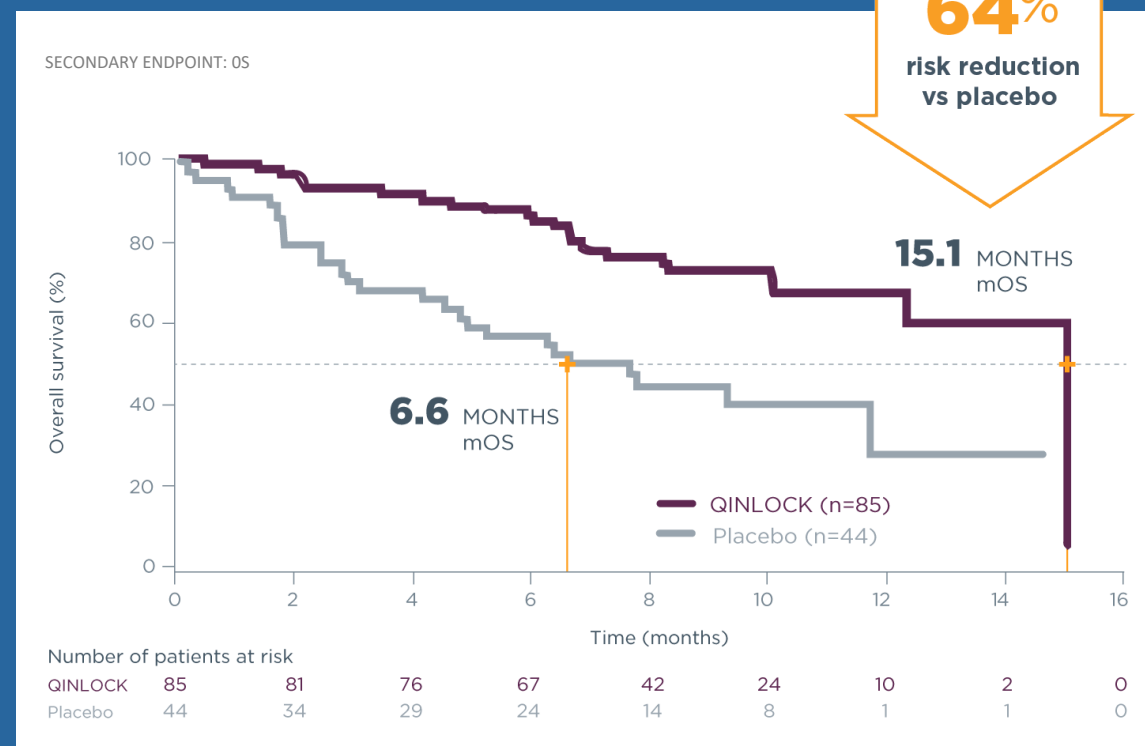
QINLOCK®: A Potential Best-In-Class Treatment for Advanced GIST

Progression-free survival



(median PFS of 6.3 months vs. 1.0 month; HR=0.15, 95% CI [0.09-0.25], $P<0.0001$)

Overall survival



(median OS of 15.1 months vs. 6.6 months; HR=0.36, 95% CI [0.21-0.62])

Key secondary endpoint of objective response rate was 9.4% compared with 0% for placebo ($P=0.0504$)

QINLOCK®: Safety Highlights from the Prescribing Information

Most Common Adverse Reactions (≥20%; Any Grade)

- Alopecia, fatigue, nausea, abdominal pain, constipation, myalgia, diarrhea, decreased appetite, palmar-plantar erythrodysesthesia syndrome, and vomiting

Warnings and Precautions

- Palmar-plantar erythrodysesthesia syndrome
- New primary cutaneous malignancies
- Hypertension
- Cardiac dysfunction
- Risk of impaired wound healing
- Embryo-fetal toxicity

Dose Modifications from INVICTUS Phase 3 Study⁽¹⁾

Any adverse reaction leading to...	RIPRETINIB (n=85)	PLACEBO (n=43) ⁽²⁾
Treatment discontinuation	7 (8%)	5 (12%)
Dose interruption	20 (24%)	9 (21%)
Dose reduction	6 (7%)	1 (2%)

Significant Unmet Medical Need Post-Imatinib

Estimated Incidence of GIST, U.S.: 4,000-6,000⁽¹⁾

1st line therapy
Imatinib⁽²⁾

18.9
months mPFS*

51.4%
ORR

49.0
months mOS*

*Results for imatinib 400 mg. This study compared imatinib 400 mg to imatinib 800 mg.⁽³⁾

2nd line therapy
Sunitinib^(4,5)

5.6
months mPFS
HR=0.33

6.8%
ORR

17.0
months mOS
HR=0.83

~2,000
U.S. incident patients
eligible for treatment⁽⁶⁾

3rd line therapy
Regorafenib⁽⁷⁾

4.8
months mPFS
HR=0.27

4.5%
ORR

17.4
months mOS
HR=0.91

~1,400–1,600
U.S. incident patients
eligible for treatment⁽⁶⁾

4th line therapy⁽⁸⁾

QINLOCK[®]
(ripretinib) 50 mg tablets

6.3
months mPFS
HR=0.15

9.4%
ORR

15.1
months mOS
HR=0.36

~1,000–1,300
U.S. incident patients
eligible for treatment⁽⁶⁾

Avapritinib is approved in the U.S. for GIST patients with PDGFRα exon 18 mutations only, which are harbored by ~6% of patients with newly diagnosed GIST.^(9,10)

Notes: HR=hazard ratio; mOS=median overall survival; mPFS=median progression free survival; ORR=objective response rate.

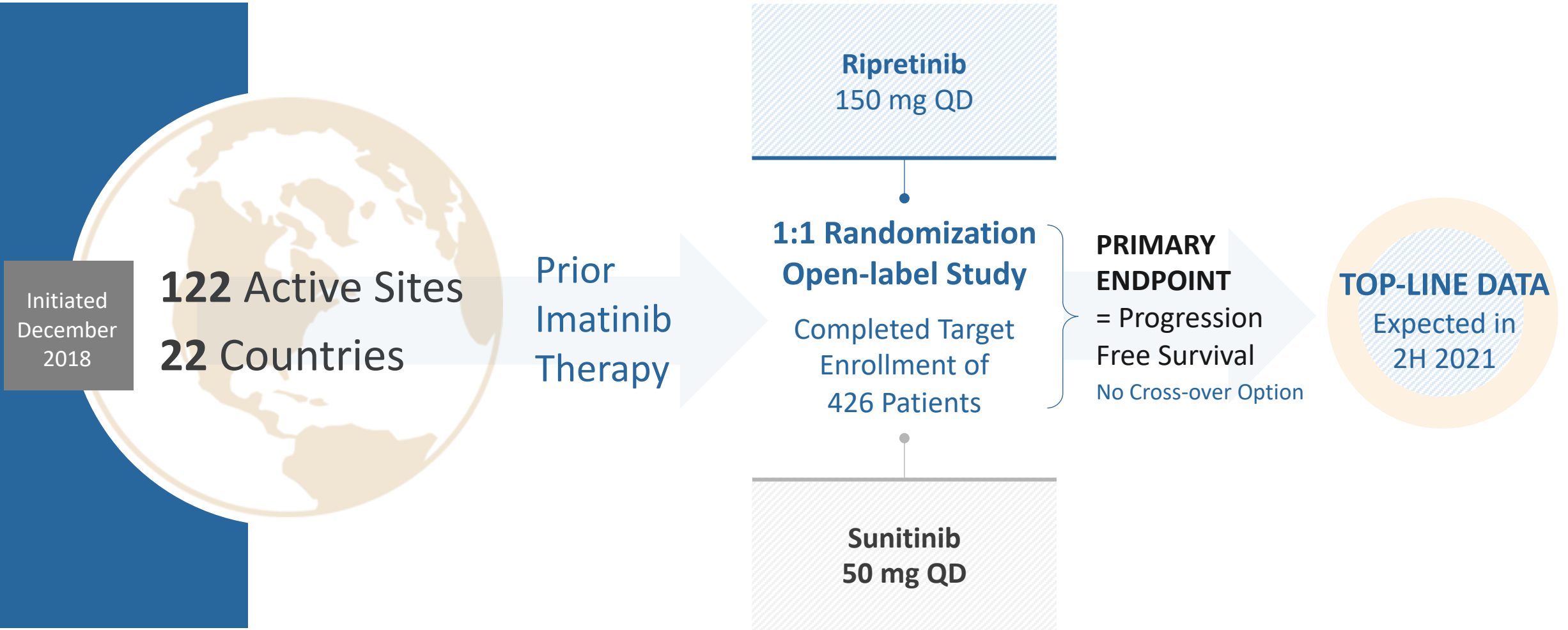
(1) American Cancer Society, Key Statistics for Gastrointestinal Stromal Tumors, Accessed December 13, 2019; (2) Gleevec [package insert]. East Hanover, NJ: Novartis Pharmaceuticals Corp; 2020; (3) Gastrointestinal Stromal Tumor Meta-Analysis Group (MetaGIST). *J Clin Oncol*. 2010; 28:1247-1253; (4) Sutent [package insert]. New York, NY: Pfizer; 2020, mPFS and mOS converted from weeks to months; (5) Garrett CR, et al. Poster presented at: Connective Tissue Oncology Society: November 13-15, 2008; London, UK. Abstract 35049; (6) Internal Deciphera estimates of annual new treatment-eligible patients are based on analyses of U.S. claims data; eligible patients for 3rd and 4th lines exclude the estimated proportion of patients across lines that die, discontinue oncology treatment, or enter clinical trial and, therefore, are not eligible for treatment. Estimates are inherently uncertain; (7) Stivarga [package insert]. Germany: Bayer Healthcare; 2020; (8) QINLOCK [package insert]. Waltham, MA: Deciphera Pharmaceuticals; 2020; (9) Ayvakit [package insert]. Cambridge, MA: Blueprint Medicines Corp; 2020; (10) Lopes LF, Bacchi CE. *J Cell Mol Med*. 2010;14:42-50.

QINLOCK®: Phase 1 GIST Cohorts

Positive Updated Results Across All Lines of Treatment

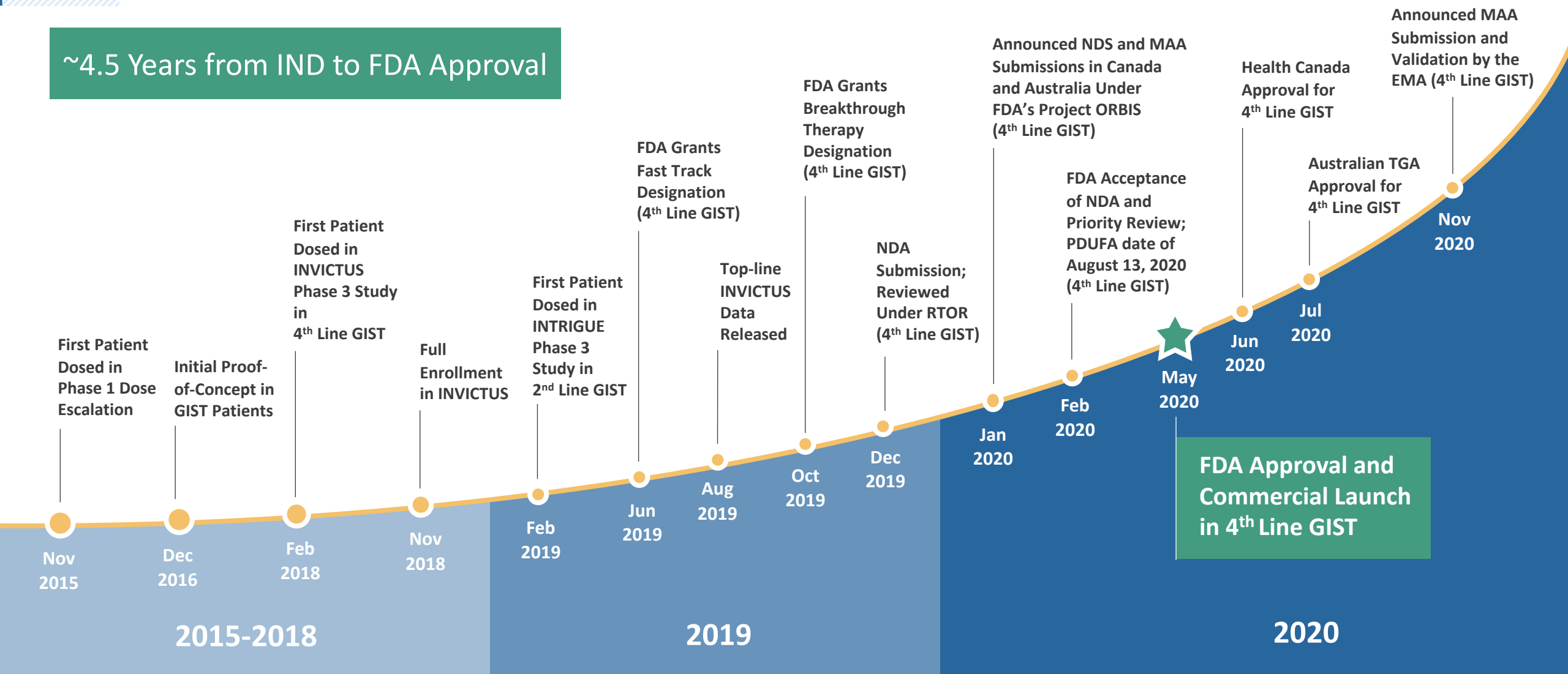
Line of Therapy ⁽¹⁾	2 nd Line (n=31)	3 rd Line (n=28)	≥4 th Line (n=83)
Median Progression Free Survival	10.7 months	8.3 months	5.5 months
Objective Response Rate	19.4%	14.3%	7.2%
Median Duration of Response	18.4 months	NE	17.5 months
Mean Treatment Duration ^(2,3)	13.2 months	13.4 months	10.5 months

Ripretinib 150 mg QD (n=142)



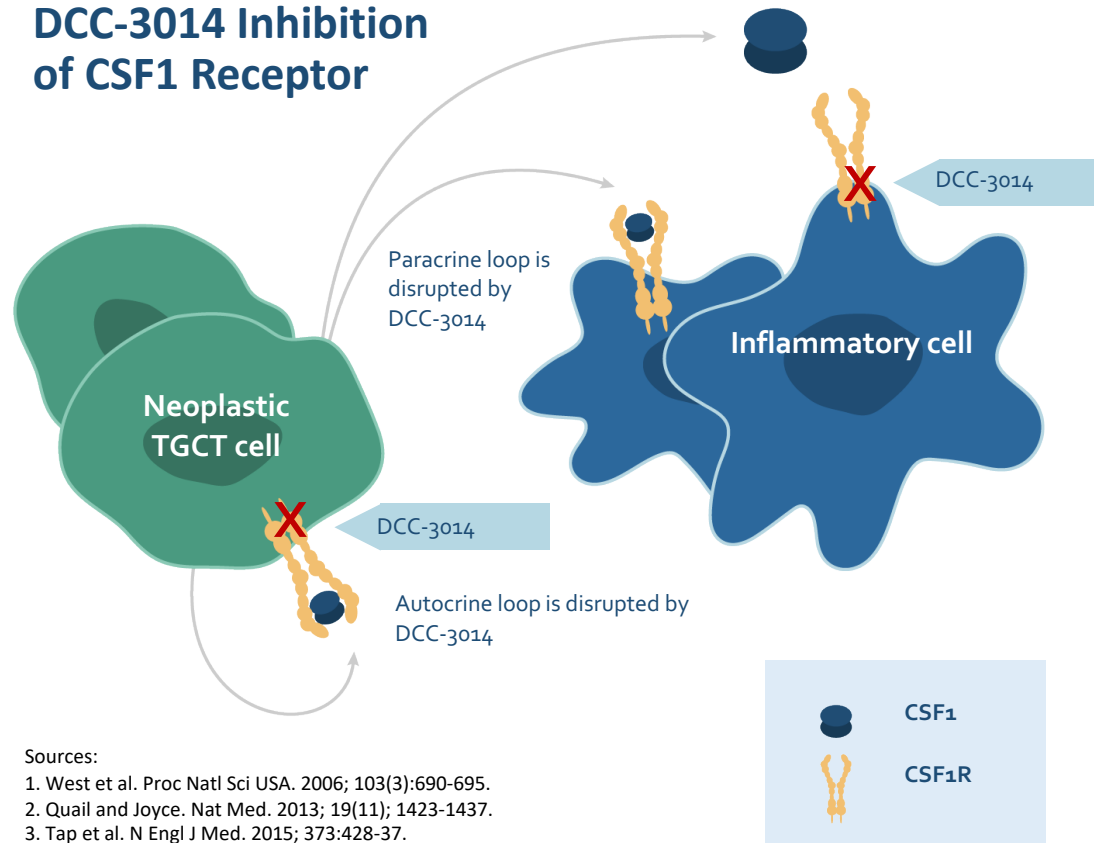
QINLOCK® (ripretinib): Rapid Clinical Development to Approval

~4.5 Years from IND to FDA Approval



DCC-3014: A Highly Selective and Potent CSF1R Inhibitor

DCC-3014 Inhibition of CSF1 Receptor

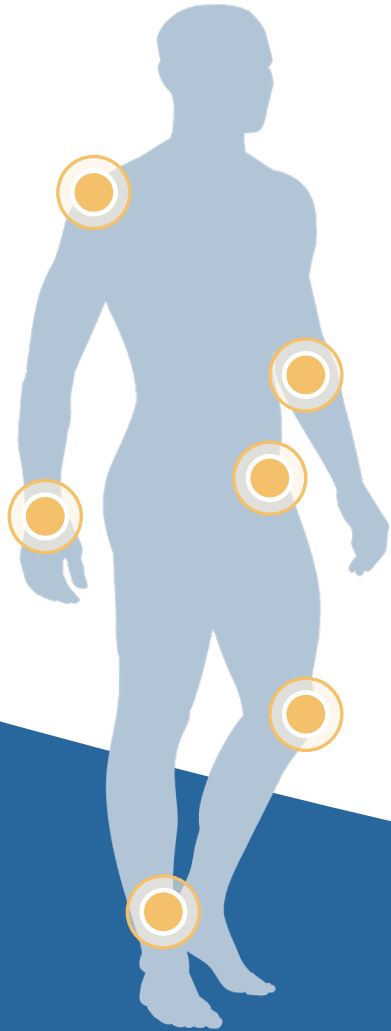


Sources:

1. West et al. Proc Natl Sci USA. 2006; 103(3):690-695.
2. Quail and Joyce. Nat Med. 2013; 19(11): 1423-1437.
3. Tap et al. N Engl J Med. 2015; 373:428-37.
4. Cannarile et al. J Immunother Cancer. 2017; 5(1): 53.

- Phase 1/2 study of DCC-3014 is ongoing
- Preliminary results showed highly encouraging signs of antitumor activity in TGCT patients (n=22)
 - 41% of patients (9 of 22) across all TGCT cohorts achieved an objective response, including one complete response
 - 78% of responders (7 of 9 patients) had a partial response at their first restaging scan evaluation (week 9)
- Treatment was generally well-tolerated with treatment emergent adverse events mostly grade 1/2
- The recommended Phase 2 dose for DCC-3014 in TGCT patients was determined to be 30 mg twice weekly (no loading dose)

Unmet Medical Need in **Tenosynovial Giant Cell Tumor (TGCT)**



Symptoms

- Rare, locally aggressive tumors
- Genetic translocation causes overproduction of CSF-1, triggering migration of inflammatory cells to tumor sites

Two Types of TGCT

1. Localized TGCT

- Affects fingers, toes, knee, wrist and ankle
- Annual incidence of new cases in the U.S.: ~13,000⁽¹⁾

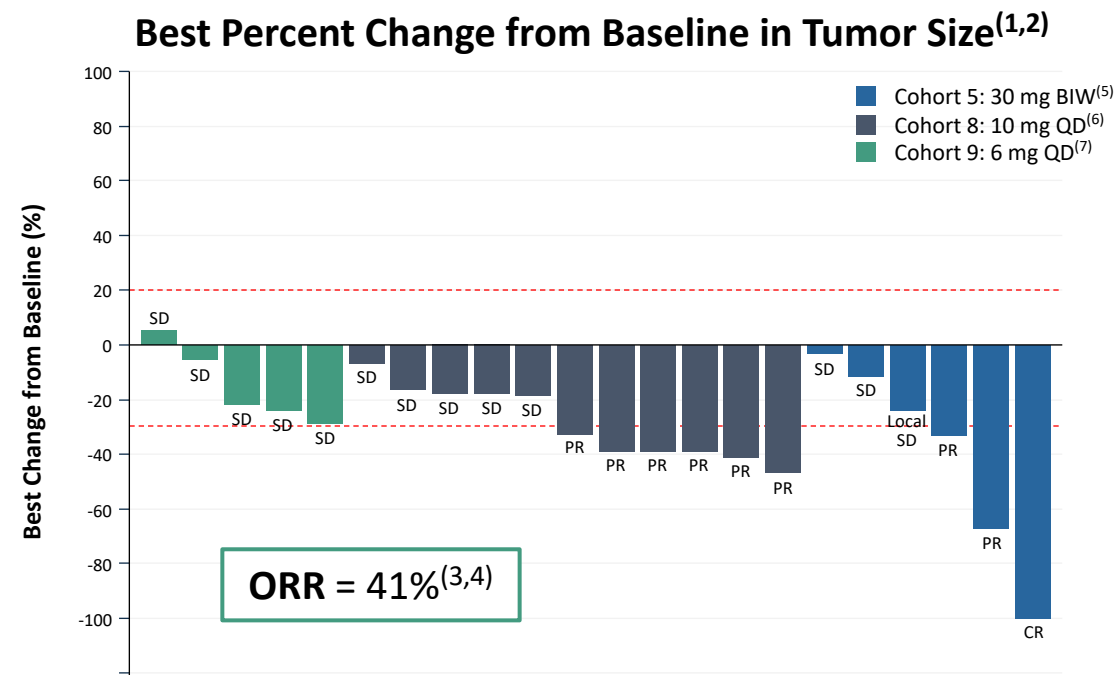
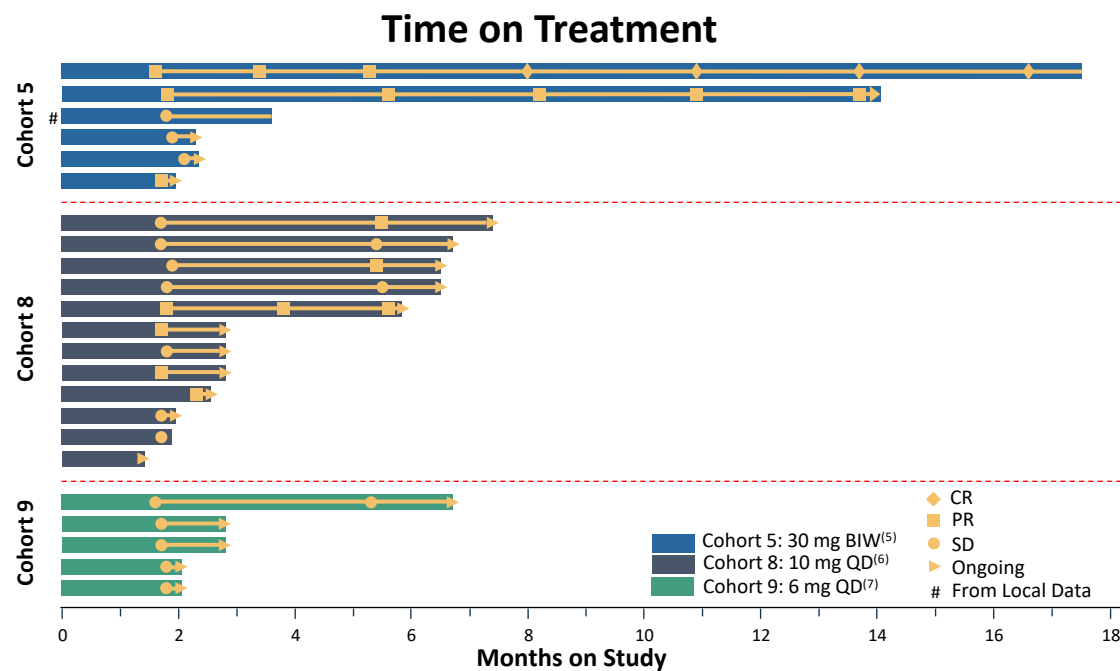
2. Diffuse TGCT

- Most commonly affects the knee, as well as hip, ankle, elbow and shoulder
- Annual incidence of new cases in the U.S.: ~1,300⁽¹⁾

Unmet Medical Need

- Surgical resection is standard treatment
- High rate of recurrence in diffuse TGCT
- CSF1R inhibition has demonstrated clinical benefit in diffuse TGCT
- Pexidartinib received FDA approval in August 2019
 - REMS and intensive monitoring required due to hepatotoxicity concerns (off-target)
 - CHMP adopted a negative opinion on the MAA in June 2020
- Unmet medical need for effective CSF1R inhibitor with favorable safety/tolerability profile for TGCT patients

DCC-3014: Dose Escalation in Phase 1 Shows Encouraging Anti-Tumor Activity in TGCT Patients



Encouraging Preliminary Anti-Tumor Activity^(3,4)

- 9 patients (41%) across all TGCT cohorts achieved an objective response (1 CR, 8 PR)
- 7 of the 9 responders (78%) had a partial response at their first restaging scan evaluation (week 9)

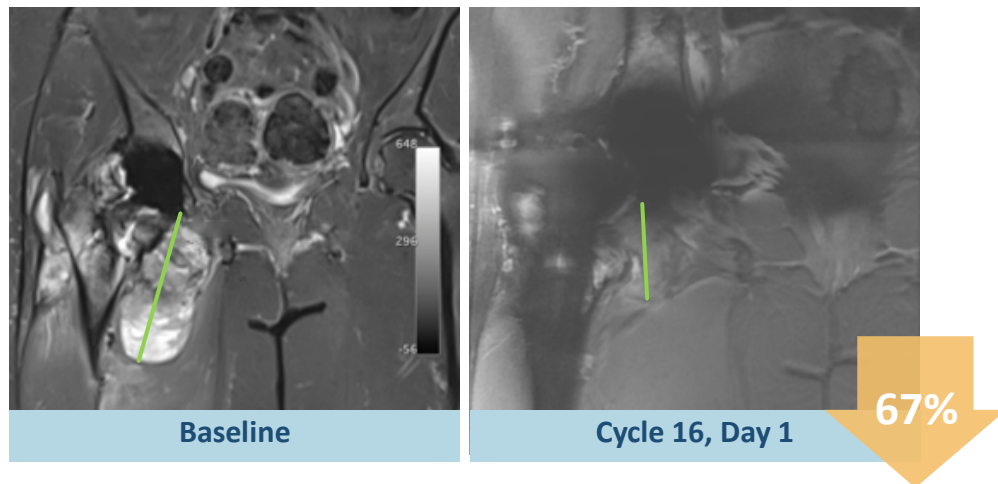
Preliminary Safety Data Shows DCC-3014 as Well Tolerated In TGCT Patients

- TEAEs occurring in $\geq 25\%$ of patients regardless of relatedness were blood CPK increased (52%), AST increased (44%), periorbital edema (44%), fatigue (40%), lipase increased (32%), and ALT increased (28%). No SAEs related to DCC-3014 were reported
- All bilirubin levels were within the normal limit and observed transaminase and pancreatic enzyme elevations were asymptomatic and not clinically significant
- One patient (4%) discontinued treatment due to an adverse event (Grade 3 AST elevation from Grade 1 at baseline)

Notes: Data presented at CTOS Annual Meeting 2020; results are reported for patients with TGCT with data cutoff for safety as of September 23, 2020 and efficacy as of October 5, 2020; safety population n=25, modified intent-to-treat population n=22; PR=partial response; ORR=objective response rate; SD=stable disease; PD=progressive disease; TEAE=treatment-emergent adverse events; SAE=serious adverse events; BIW=twice weekly; QD=daily; CPK=creatinine phosphokinase; AST=aspartate amino transferase; ALT=alanine aminotransferase; (1) Waterfall plot excludes 3 patients yet to reach the study's first efficacy assessment timepoint; (2) Dotted lines denote 30% decrease and 20% increase in tumor size cutoffs for partial response and progressive disease, respectively; (3) Assessed by independent central review unless otherwise noted (RECIST v1.1) (4) Includes 1 complete response (confirmed) and 8 partial responses (2 confirmed and 6 to be confirmed at future follow up); (5) After 5-day 30 mg QD loading dose; (6) After 3-day 30 mg QD loading dose; (7) After 3-day 20 mg QD loading dose.

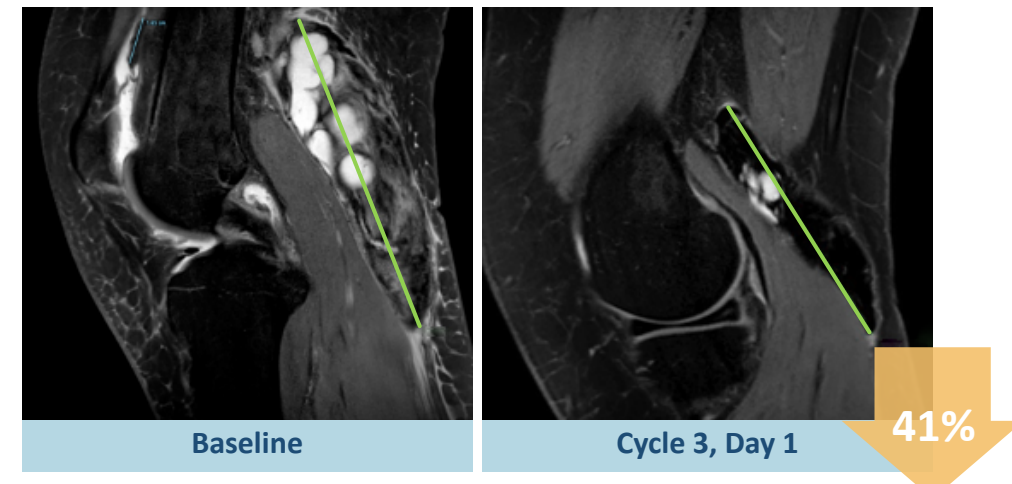
DCC-3014: TGCT Case Studies from Phase 1

Case Study 1



- 57-year-old female diagnosed with TGCT (hip) in 2014
- Prior surgeries: 2 resections, 2 synovectomies, 1 total hip replacement, and 1 cryoablation (2014-2019)
- No prior systemic therapy
- Enrolled in July 2019 (cohort 5 – DCC-3014 dose: 30 mg twice weekly⁽¹⁾)
 - Dose reduced to 20 mg twice weekly in cycle 6 due to grade 3 urticaria, re-escalated in cycle 10
- Partial response after 2 cycles (33% decrease from baseline)
- Treatment ongoing in cycle 16 (67% decrease at cycle 16, day 1)

Case Study 2

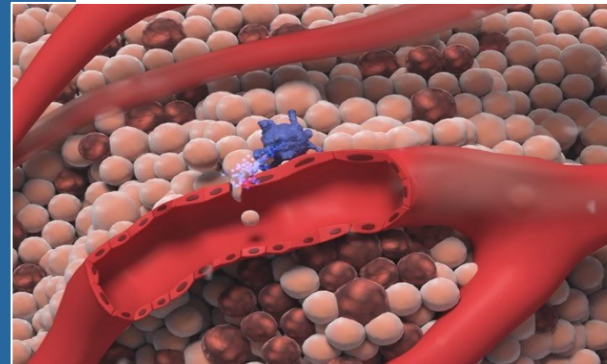


- 39-year-old female diagnosed with TGCT (knee) in 2020
- No prior systemic therapy or surgery
- Enrolled in June 2020 (cohort 8 – DCC-3014 dose: 10 mg daily⁽²⁾)
- Partial response after 2 cycles (41% decrease from baseline)
- Treatment ongoing in cycle 4

Rebastinib: A Highly Potent and Selective TIE2 Inhibitor

Potential Benefits in Combination with Chemotherapy

- Chemotherapy leads to recruitment of pro-tumoral M2 macrophages from the bone marrow
- M2 macrophages serve as pumps for tumor cell intravasation and metastasis
- Rebastinib is designed to inhibit chemotherapy-induced recruitment of M2 macrophages to tumors
- Rebastinib is designed to block rebound angiogenesis and inhibit M2 macrophages



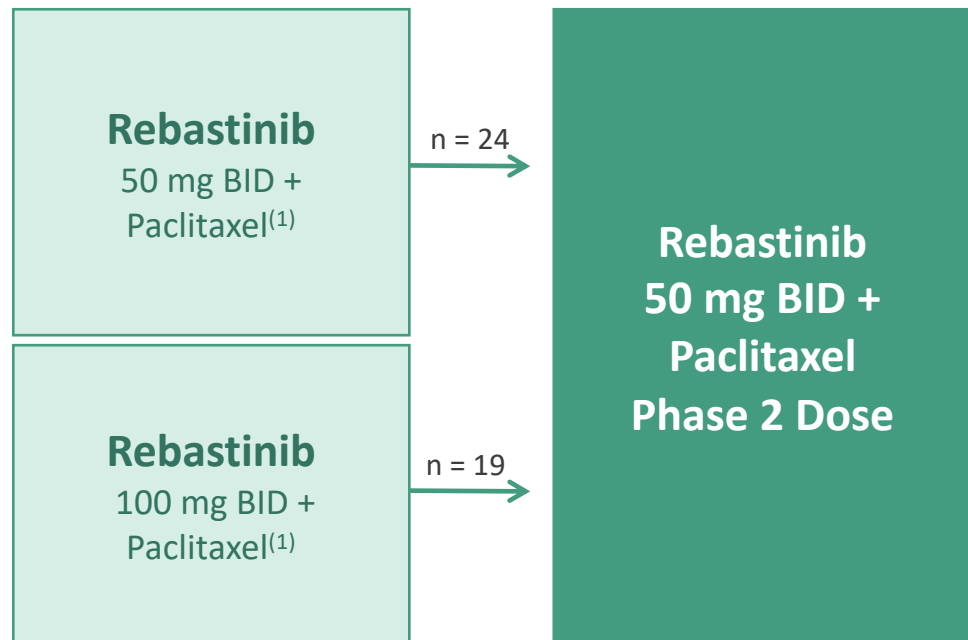
- Potent, small molecule inhibitor of TIE2
- Targets TIE2 expressing macrophages (TEMs) and endothelial cells in tumor environment
- Combination of rebastinib with chemotherapy may increase tumor killing through multiple mechanisms
 - Tumor vascularization, dissemination, metastasis, immunotolerance

Development status

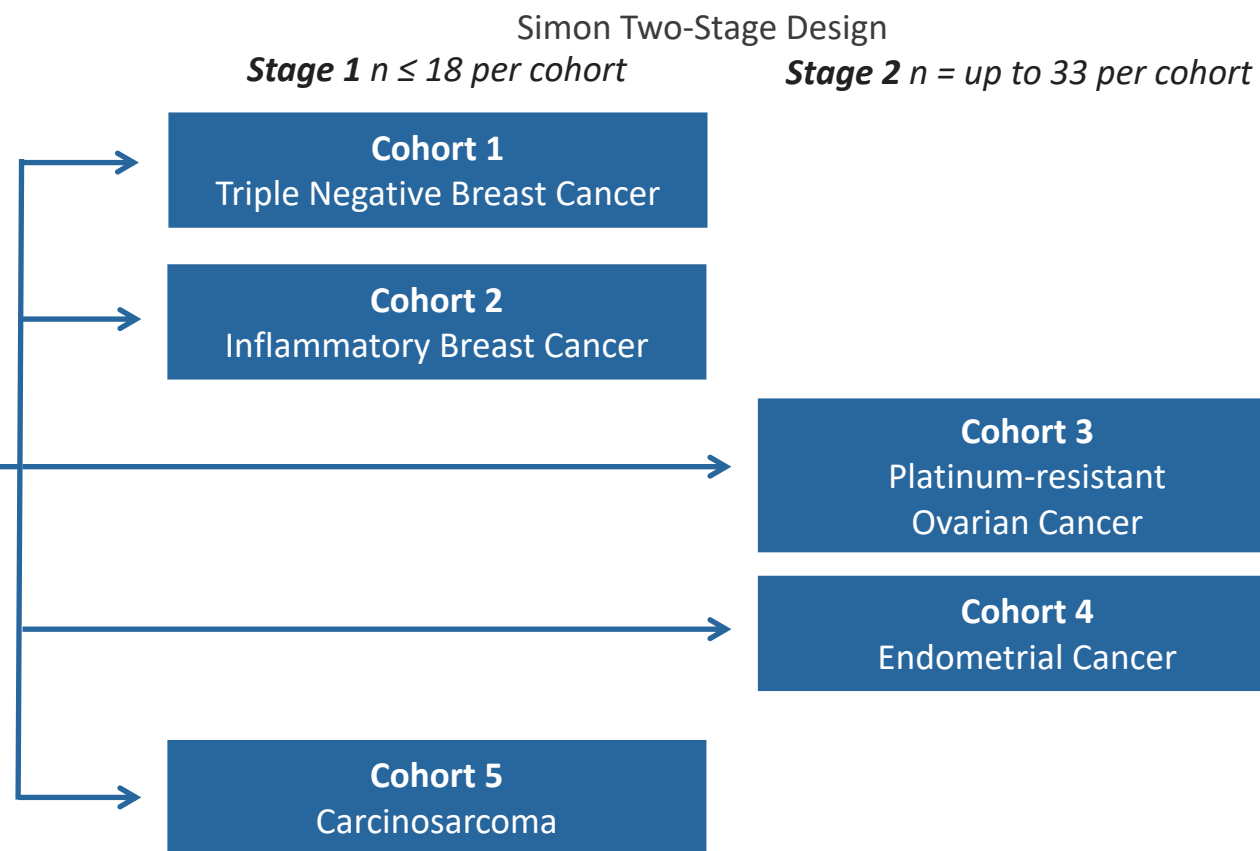
- Two ongoing studies in combination with paclitaxel or carboplatin
- Encouraging preliminary results from Part 2 of the Phase 1b/2 study in combination with paclitaxel in endometrial and platinum-resistant ovarian cancer
- Selected Phase 2 dose and activated Part 2 of the Phase 1b/2 study in combination with carboplatin in January 2020; data from Part 1 of study presented at the ESMO 2020

Rebastinib: Phase 1b/2 Study in Combination with Paclitaxel

Part 1

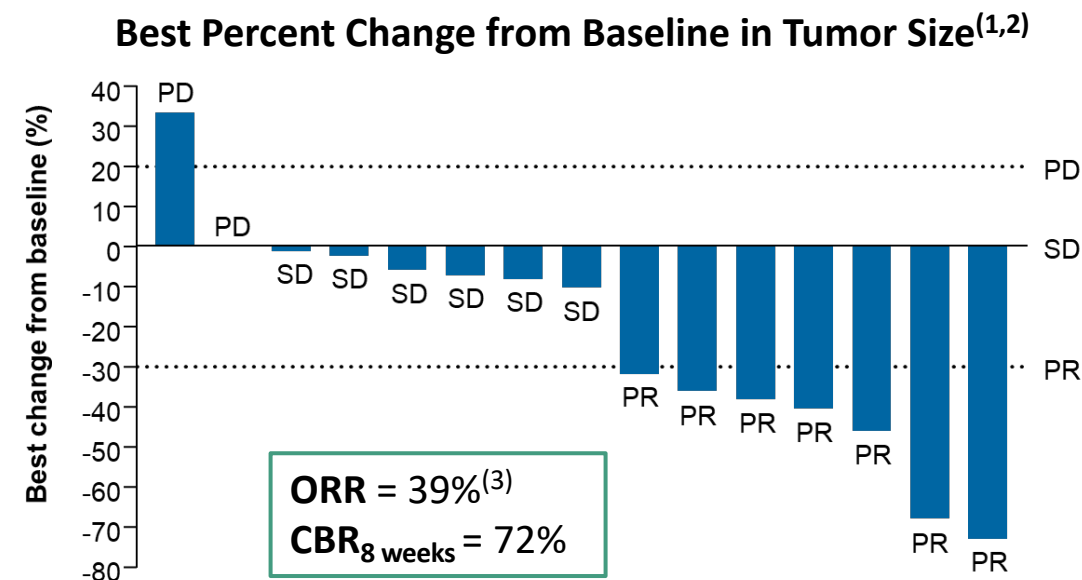
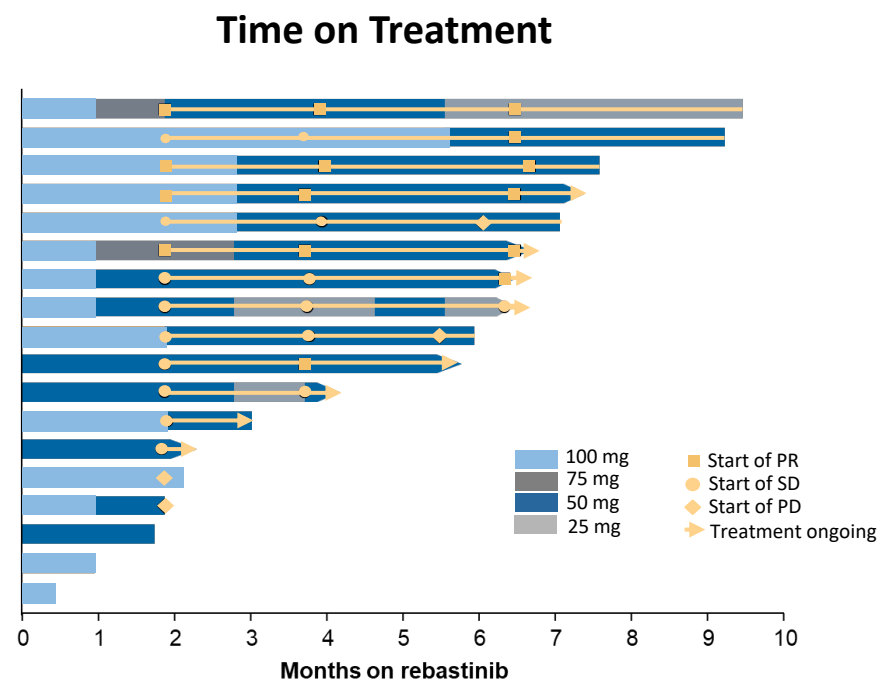


Part 2



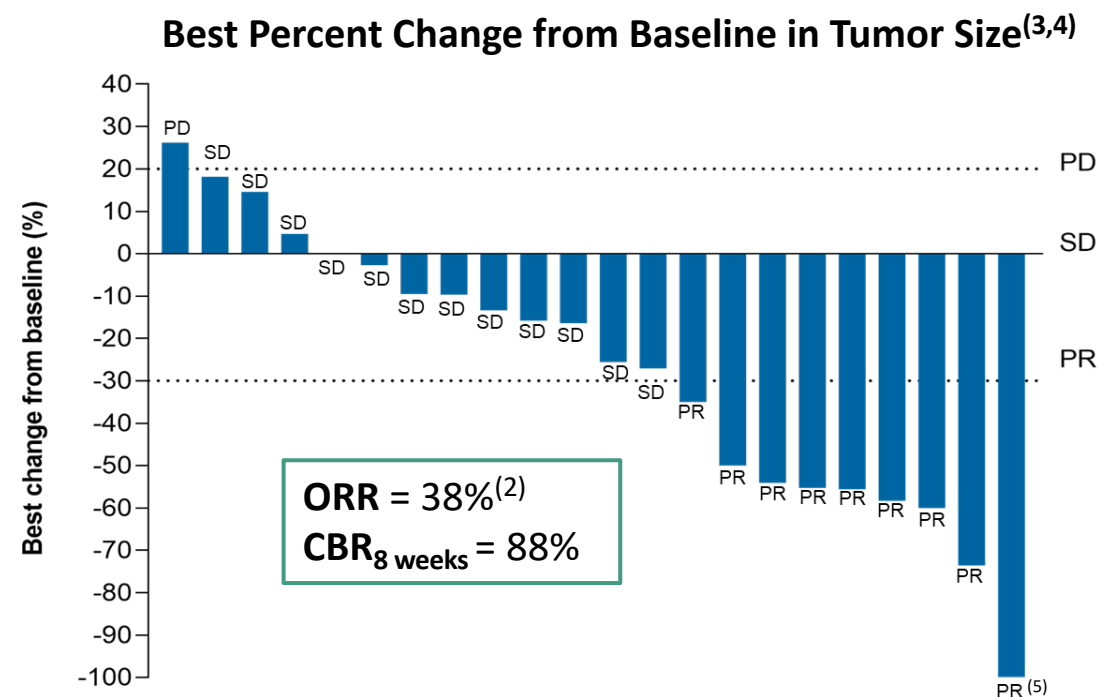
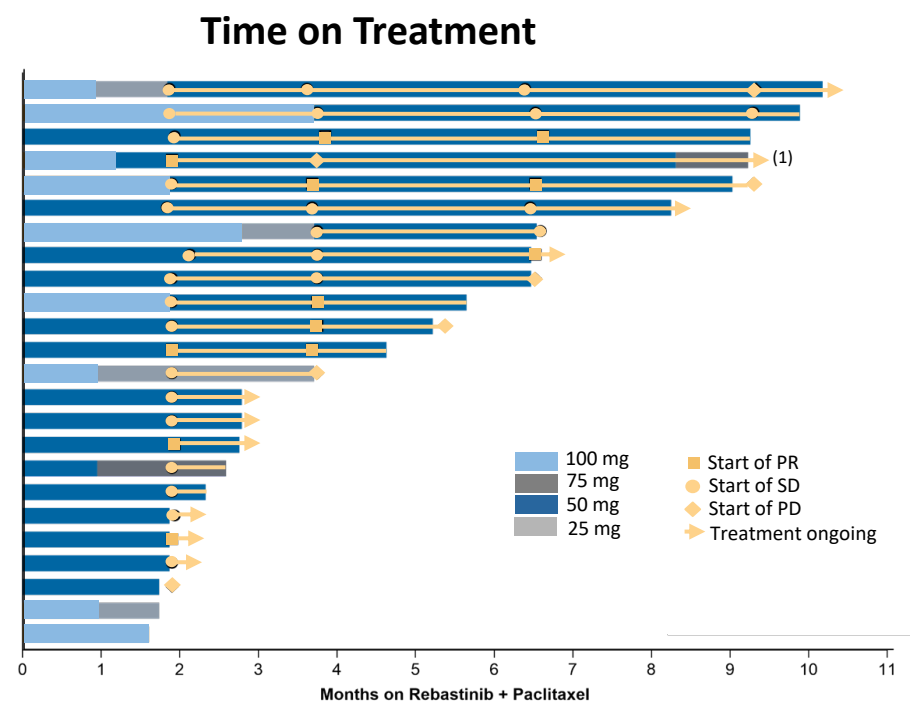
If ≥ 5 responses in Stage 1, enroll additional patients in Stage 2 (up to 33 patients per cohort)
If < 5 responses, discontinue the cohort

Rebastinib: Part 2 (Stage 1) of Phase 1b/2 Study in Combination with Paclitaxel Shows Encouraging Anti-tumor Activity in Endometrial Cancer



- All patients received ≥ 1 prior line of the combination of paclitaxel/carboplatin and 20 (95%) received ≥ 2 prior anti-cancer regimens
- Majority of the common ($\geq 15\%$) TEAEs regardless of causality were grade ≤ 2
- SAEs at least possibly related to rebastinib occurred only at 100 mg BID and resolved after dose reductions
- Nine patients experienced SAEs at least possibly related to rebastinib including muscular weakness (n=2), acute myocardial infarction (n=1), atrial flutter (n=1), dehydration (n=1), head discomfort (n=1), nausea (n=1), peripheral edema (n=1), and pneumonia (n=1)

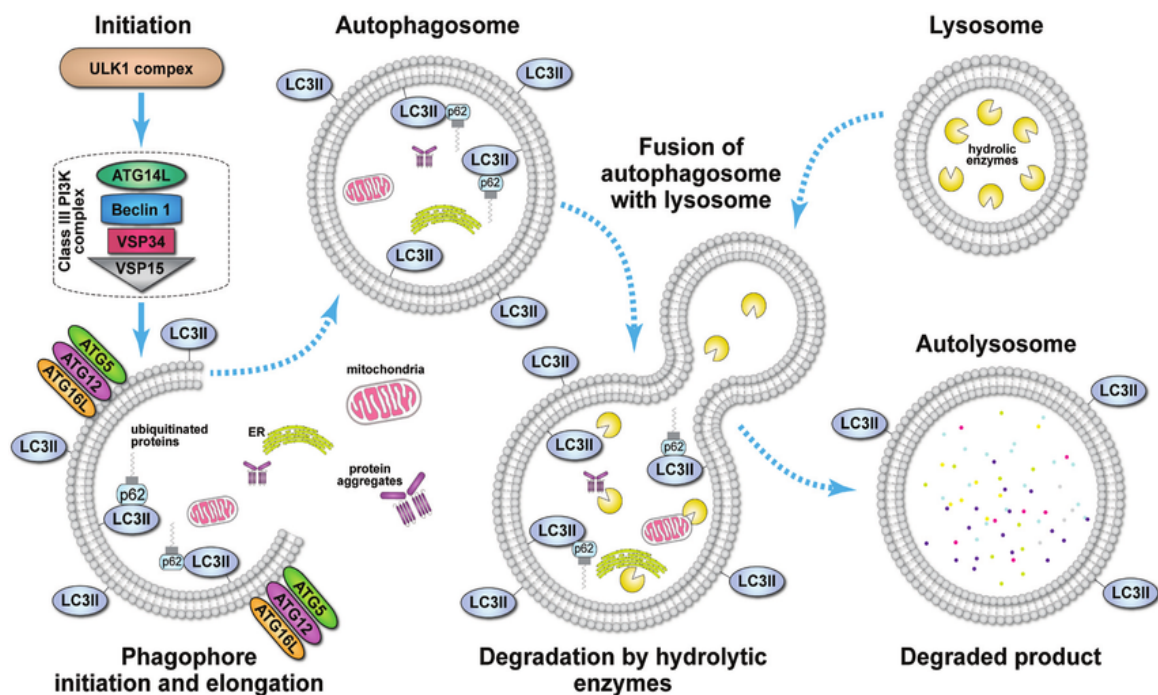
Rebastinib: Part 2 (Stage 1) of Phase 1b/2 Study in Combination with Paclitaxel Shows Encouraging Anti-tumor Activity in Platinum-resistant Ovarian Cancer



- All patients received ≥ 1 prior line of the combination of paclitaxel/carboplatin and 23 (79%) received ≥ 4 prior anti-cancer regimens
- TEAEs occurring in $\geq 25\%$ of patients regardless of causality were fatigue (41%), dry mouth (38%), nausea (34%), diarrhea (31%), stomatitis (31%), abdominal pain (28%), and peripheral sensory neuropathy (28%)
- 11 patients (38%) had a TEAE of Grade ≥ 3
- Two patients experienced SAEs at least possibly related to rebastinib: muscular weakness/fatigue (starting dose rebastinib 100 mg BID and resolved with drug interruption) and urinary tract infection (starting dose rebastinib 50 mg BID)
- A CA-125 response occurred in 10/17 patients (59%)

DCC-3116 is a Potent & Selective ULK Inhibitor Designed to Inhibit Autophagy

ULK: Initiating Factor for Autophagy



Adapted from: Ndoye A and Weeraratna AT. Autophagy- An emerging target for melanoma therapy [version 1]. F1000Research 2016, 5:1888.



First-in-class opportunity for new therapeutic in RAS mutant cancers

- RAS mutant cancers have high basal levels of autophagy
- DCC-3116 observed preclinically to durably and potently inhibit autophagy in RAS mutant cancer cell lines through the inhibition of ULK kinase
- Combination of DCC-3116 and MAPK pathway inhibitors have been observed to synergize to block RAS mutant cancers *in vivo*



Highly potent and selective (IC_{50} at 1 mM ATP)

- ULK1 4.7 nM and ULK2 35 nM
- No off-target kinases within 30-fold of ULK1/ 5 kinases within 100-fold of ULK1

Designed to avoid CNS exposure

IND submission expected in 4Q20

Financial Highlights

As of September 30, 2020

Shares
Outstanding


56.6 MM (basic)
63.5 MM (fully-diluted)

Cash Expected to Fund Operating
Expenses and CapEx into the
Second Half of 2022

Cash, Cash
Equivalents
& Marketable
Securities

\$584.3 MM

Significant Expected 2020 Milestones Across the Pipeline

	<ul style="list-style-type: none">✓ FDA approval and U.S. commercial launch in 4th line GIST (2Q20)✓ Present Phase 1 study expansion data (3Q20)✓ Submit EU Marketing Authorisation Application to EMA (3Q20)✓ Complete enrollment in the INTRIGUE Phase 3 study in 2nd line GIST (4Q20)
DCC-3014	<ul style="list-style-type: none">✓ Select Phase 2 dose for TGCT and initiate the expansion portion of study (4Q20)✓ Update Phase 1 data in TGCT patients (4Q20)
Rebastinib	<ul style="list-style-type: none">✓ Select Phase 2 dose and activate Part 2 of Phase 1b/2 study in combination with carboplatin (1Q20)✓ Present Phase 1b/2 data in combination with paclitaxel (2Q20 and 3Q20)✓ Present Phase 1b/2 data in combination with carboplatin (3Q20)
DCC-3116	<ul style="list-style-type: none">○ Submit IND application to FDA (4Q20)

THANK YOU

