



**Defeating Cancer:  
The Challenge.  
Our Mission.**

*May 2019*



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# Addressing Unmet Needs for Patients



**Three  
Clinical-Stage  
Programs**

**Two Randomized  
Phase 3 Trials Ongoing**

**Commercial  
Preparations  
Underway to Support  
First Potential  
Approval**

**Proprietary Kinase Switch Control Inhibitor Platform**

# Strong Clinical Stage Oncology Pipeline of Novel Kinase Inhibitors

	Pre Clinical	Phase 1	Phase 1b/2	Phase 3	Global Rights
<b>Ripretinib<sup>(1)</sup>: Broad Spectrum Inhibitor of KIT &amp; PDGFRα</b>					
INVICTUS (≥4L GIST <sup>(2)</sup> )					decīphera
INTRIGUE (2L GIST)					
GIST (2L, 3L, ≥4L)					
Other Solid Tumors					
<b>Rebastinib: Selective Inhibitor of TIE2</b>					
Solid Tumors in Combination with paclitaxel (includes breast, ovarian & endometrial cancers)					decīphera
Solid Tumors in Combination with carboplatin (includes mesothelioma, ovarian & breast cancers)					
<b>DCC-3014: Selective Inhibitor of CSF1R</b>					
Tenosynovial Giant Cell Tumors					decīphera
Solid Tumors					
<b>Additional Programs</b>					
Cancer Metabolism (undisclosed kinase)					decīphera
Immunokinase (undisclosed kinase)					decīphera



# Ripretinib: Designed to Address Relevant Mutations in GIST

## Highly potent small molecule KIT and PDGFR $\alpha$ inhibitor

- Designed to inhibit the full spectrum of known KIT and PDGFR $\alpha$  mutations

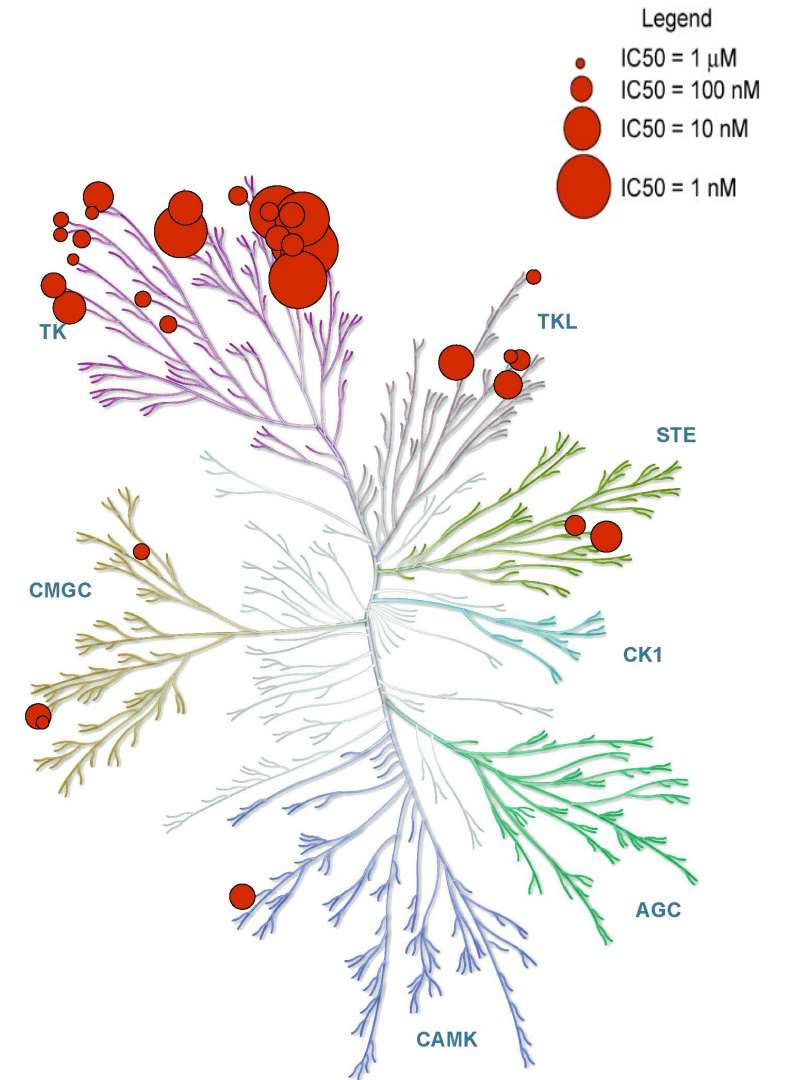
## Two Randomized Phase 3 Clinical Trials Ongoing

- INVICTUS: Placebo-controlled, pivotal trial in  $\geq 4^{\text{th}}$  line GIST
  - PFS primary endpoint
  - Top-line data read-out expected mid-2019
- INTRIGUE: Compared to sunitinib, pivotal trial in 2<sup>nd</sup> line GIST
  - PFS primary endpoint
  - Initiated December 2018
- Phase 1 expansion study ongoing

## Clinical proof-of-concept demonstrated in 178 GIST patients in Phase 1

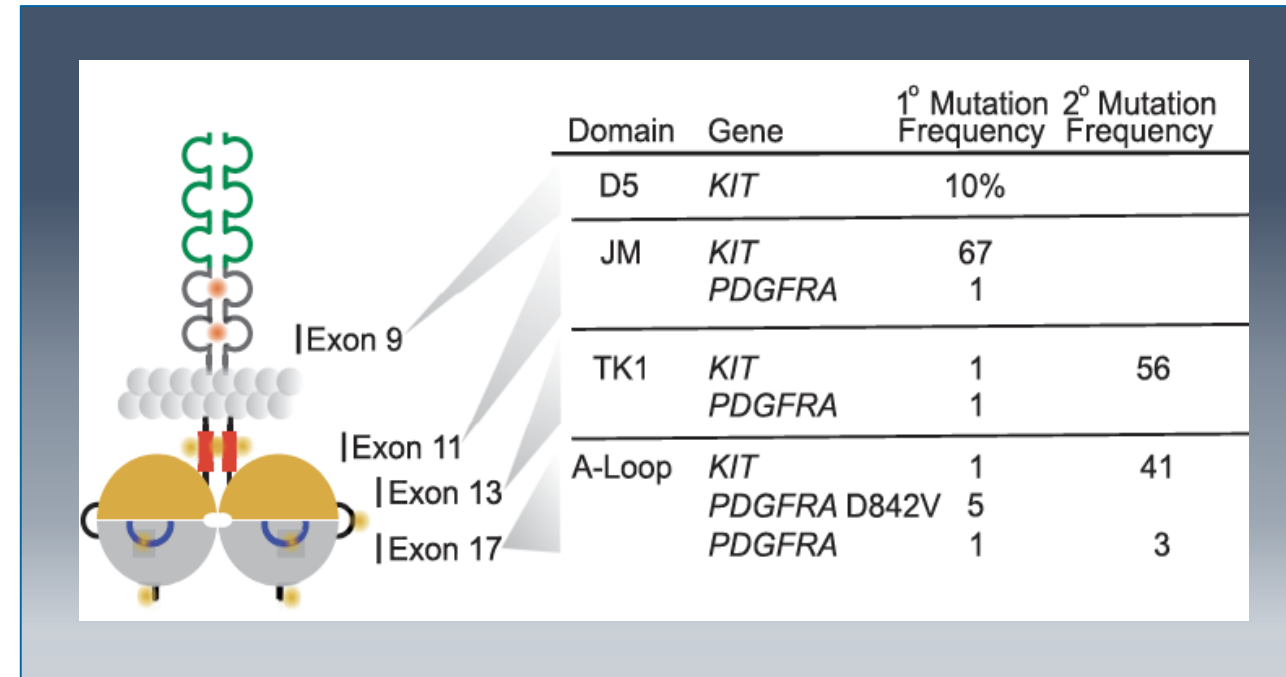
## Favorable tolerability profile

## IP: Composition and method of use (2032)



# Mutations in KIT Drive ~80% of GIST

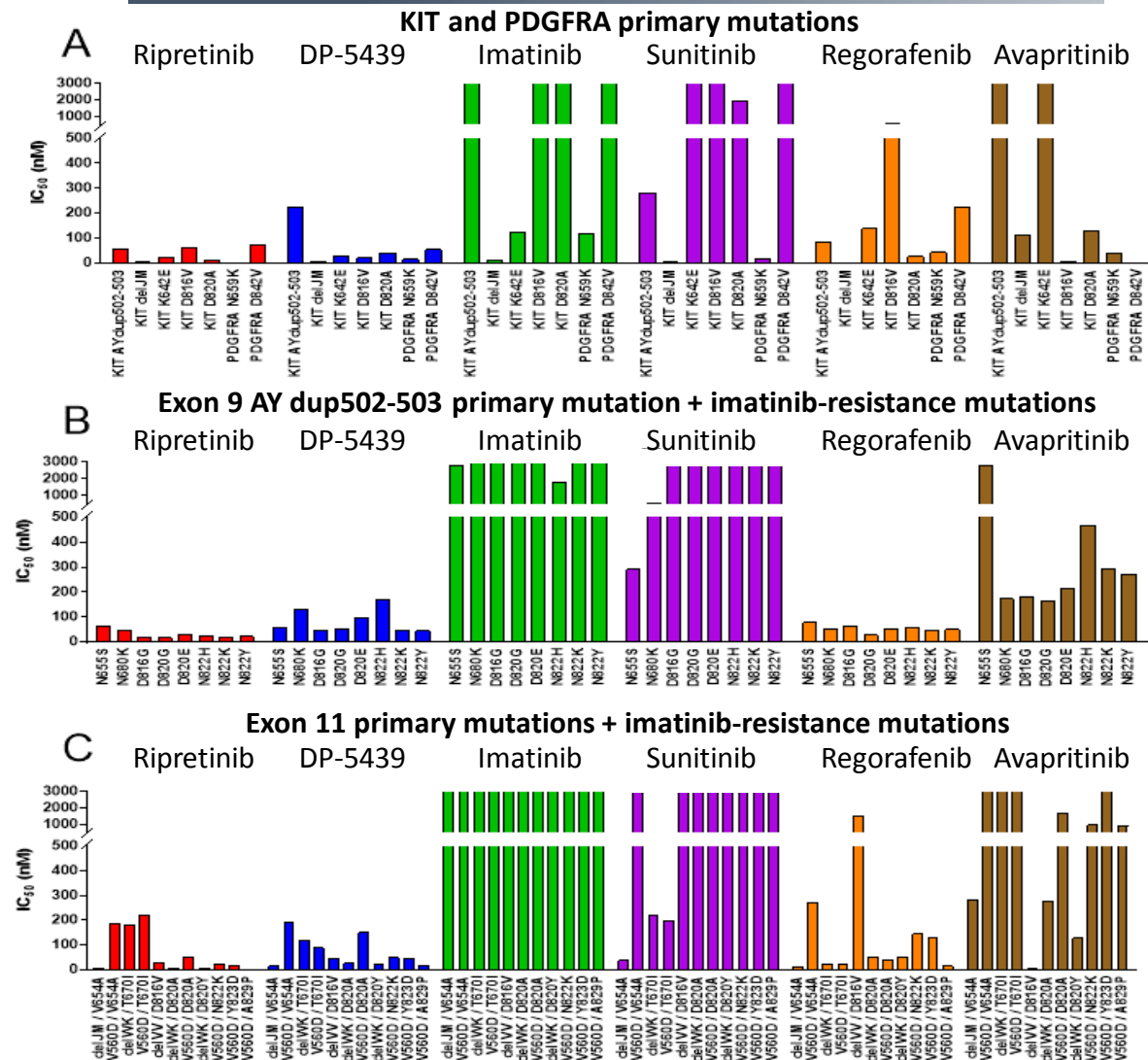
- Majority of patients with KIT primary mutations respond to 1<sup>st</sup> line imatinib
  - Resistance develops most commonly due to secondary mutations in KIT
- Approved 2<sup>nd</sup> and 3<sup>rd</sup> line agents (sunitinib and regorafenib) confer modest clinical benefit compared to imatinib
  - Multiple drug-resistant mutations often arise in individual tumors
- Unmet medical need for agents that can address relevant primary/secondary KIT mutations across all lines of therapy



# Ripretinib: Broad Mutational Coverage in KIT and PDGFRα (In Vitro Data)

- Broadly inhibits relevant GIST mutations
  - KIT mutations: Exons 9, 11, 13, 14, 17, 18
  - PDGFRA mutation: Exon 18
- Type I inhibitors exhibit weak activity in relevant GIST mutations
  - Primary KIT mutations: Exon 9, exon 11 V560D and exon 13 K642E
  - Secondary KIT mutations: Exon 13 and exon 14

## Ripretinib Inhibits Phosphorylation of KIT and PDGFRα in Cellular Assays



Notes: (A) GIST primary mutations; or imatinib-resistant KIT mutations with (B) exon 9 or (C) exon 11 primary mutations.

# Approved Therapies for GIST: Clinical Goal is Disease Control

	First Line	Second Line	Third Line
	imatinib (n=147) (Blanke et al. 2008)	sunitinib (n=243) (Demetri et al. 2012)	regorafenib (n=133) (Demetri et al. 2013)
Progression Free Survival <sup>(1)</sup> (weeks)	104.0	26.6	20.8
Objective Response Rate (%)	68.1%	7.0%	4.5%
Stable Disease (%)	15.6%	53.0%	48.1%
<b>Disease Control Rate (“DCR”) (%)</b>	<b>83.7%<sup>(2)</sup></b>	<b>60.0%<sup>(2)</sup></b>	<b>52.6%<sup>(3)</sup></b>

**No approved therapy for 4<sup>th</sup> line patients**



# Ripretinib: Phase 1 Trial Summary

## Part 1: Dose Escalation

- Key Objectives: MTD, recommended Phase 2 dose, safety, tolerability, pharmacokinetics and anti-tumor activity
- Design: 3+3 design with enrichment of targeted patients
- Dose Levels: 20, 30, 50, 100, 150, and 200 mg BID; and 100, 150 and 250 mg QD
- MTD: not determined

Advanced Malignancies  
(n=68)

**Recommended  
Dose 150 mg  
QD<sup>(1)</sup>**

## Part 2: Dose Expansion

- **10 cohorts up to 270 pts; 4 cohorts fully enrolled**

### Enrollment Complete

2<sup>nd</sup> – 3<sup>rd</sup> Line  
GIST

n=55

4<sup>th</sup> Line  
GIST

n=40

>4<sup>th</sup> Line  
GIST

n=35

Other Solid  
Tumors

n=10

### Enrollment Continuing

Systemic  
Mastocytosis

Malignant  
Gliomas

### New Cohorts Initiated

NSCLC, Germ  
Cell & Penile

Melanomas

Soft Tissue  
Sarcomas

Renal  
Impairment<sup>(2)</sup>

# Ripretinib: Phase 1 Demography and Baseline Characteristics

ESMO 2018				
GIST Patients at >100 mg/d	2 <sup>nd</sup> Line (n=38)	3 <sup>rd</sup> Line (n=29)	≥ 4 <sup>th</sup> Line (n=111) <sup>(4)</sup>	Total (n=178)
<b>Age Median (min, max)</b>	60 (32, 80)	64 (48, 82)	60 (27, 87)	61 (27, 87)
<b>Primary Mutation<sup>(1)</sup> n (%)</b>				
KIT Exon 9	4 (11%)	8 (28%)	22 (20%)	34 (19%)
KIT Exon 11	31 (82%)	20 (69%)	71 (64%)	122 (69%)
Other KIT <sup>(2)</sup>	0 (0%)	1 (3%)	12 (11%) <sup>(3)</sup>	13 (7%) <sup>(3)</sup>
PDGFRα	3 (8%)	0 (%)	6 (5%)	9 (5%)
<b>Pts at RP2D<sup>(5)</sup> (150 mg QD)</b>	32 (84%)	27 (93%)	83 (75%)	142 (80%)

Notes: (1) Primary mutation per local assessment; (2) KIT exon 13 (4), KIT exon 17 (5), not done (3); (3) Includes one SDH deficient patient; (4) Mean # of prior therapies is 4.63 (range 4-7); (5) RP2D = Recommended Phase 2 Dose.

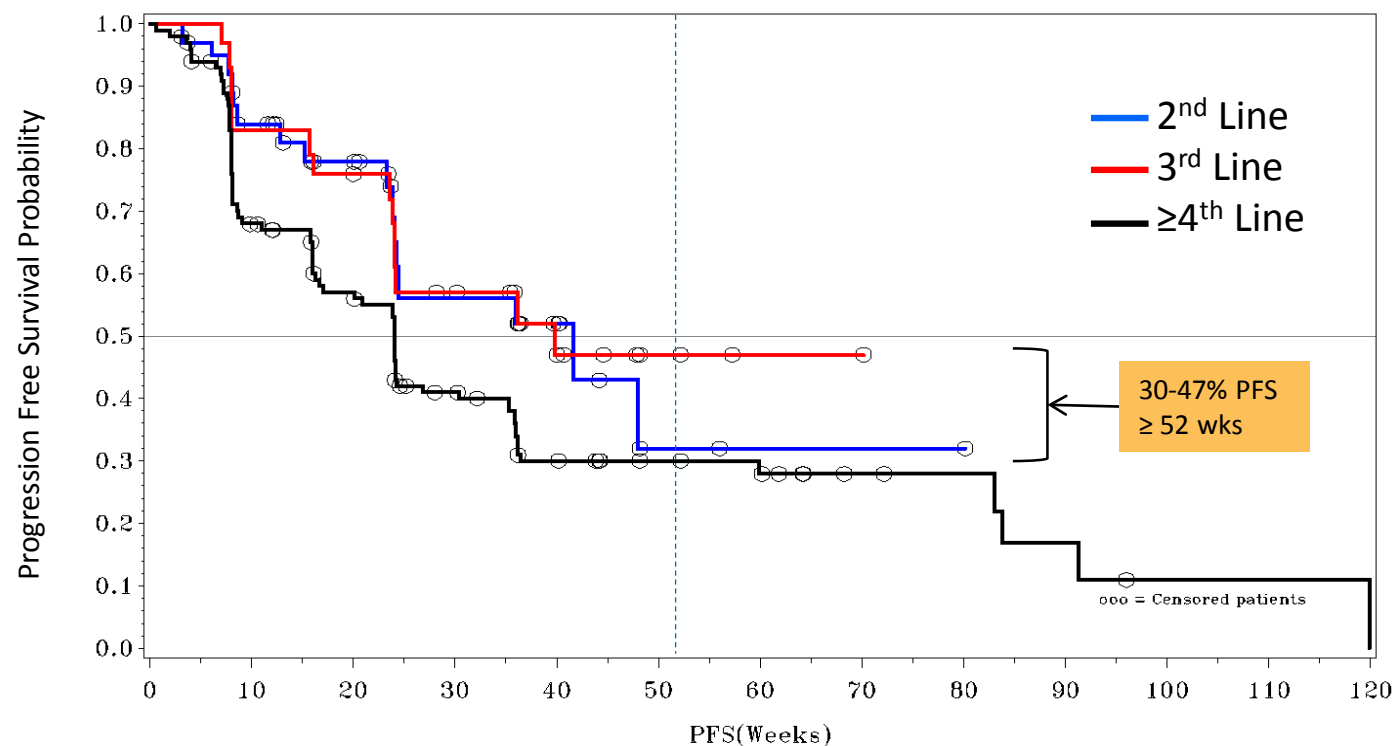
# Ripretinib: Preliminary Phase 1 Results Provided Encouraging Efficacy Measures Across All Lines of Treatment >100 mg/d (n=178)

Line of Therapy	Objective Response Rate <sup>(1)(2)</sup>	Disease Control Rate @ 3 Months <sup>(1)</sup>	Median Progression Free Survival (mPFS) <sup>(1)</sup>	Censored Patients for mPFS <sup>(1)</sup>	Median Treatment Duration <sup>(5)</sup>
2 <sup>nd</sup> Line (n=38)	18% <sup>(3)</sup>	79%	42 weeks	58%	48 weeks
3 <sup>rd</sup> Line (n=29)	24%	83%	40 weeks	52%	NR
≥4 <sup>th</sup> Line (n=111)	9% <sup>(4)</sup>	66%	24 weeks	35%	28 weeks
2 <sup>nd</sup> & 3 <sup>rd</sup> Line (n=67)	21% <sup>(3)</sup>	81%	40 weeks	55%	52 weeks

Notes: (1) Based on cut off date of August 31, 2018; RECIST data per investigator assessment; (2) ORR by Best Response includes nine unconfirmed responses in 2<sup>nd</sup> line (n=1), 3<sup>rd</sup> line (n=3) and ≥4<sup>th</sup> line (n=5); (3) Does not reflect one PR reported after cut off date; (4) Excludes five patients due to missing data at the time of data cut off (n=2), lack of first tumor assessment (n=1), withdrawal of consent prior to first assessment (n=1) and unrelated death at C1D4 prior to first assessment; (5) Includes 14 patients who elected for intra-patient dose escalation.

# Ripretinib Phase 1 Data Demonstrated Prolonged Progression Free Survival in a Meaningful Subset of Patients Across All Lines of Treatment

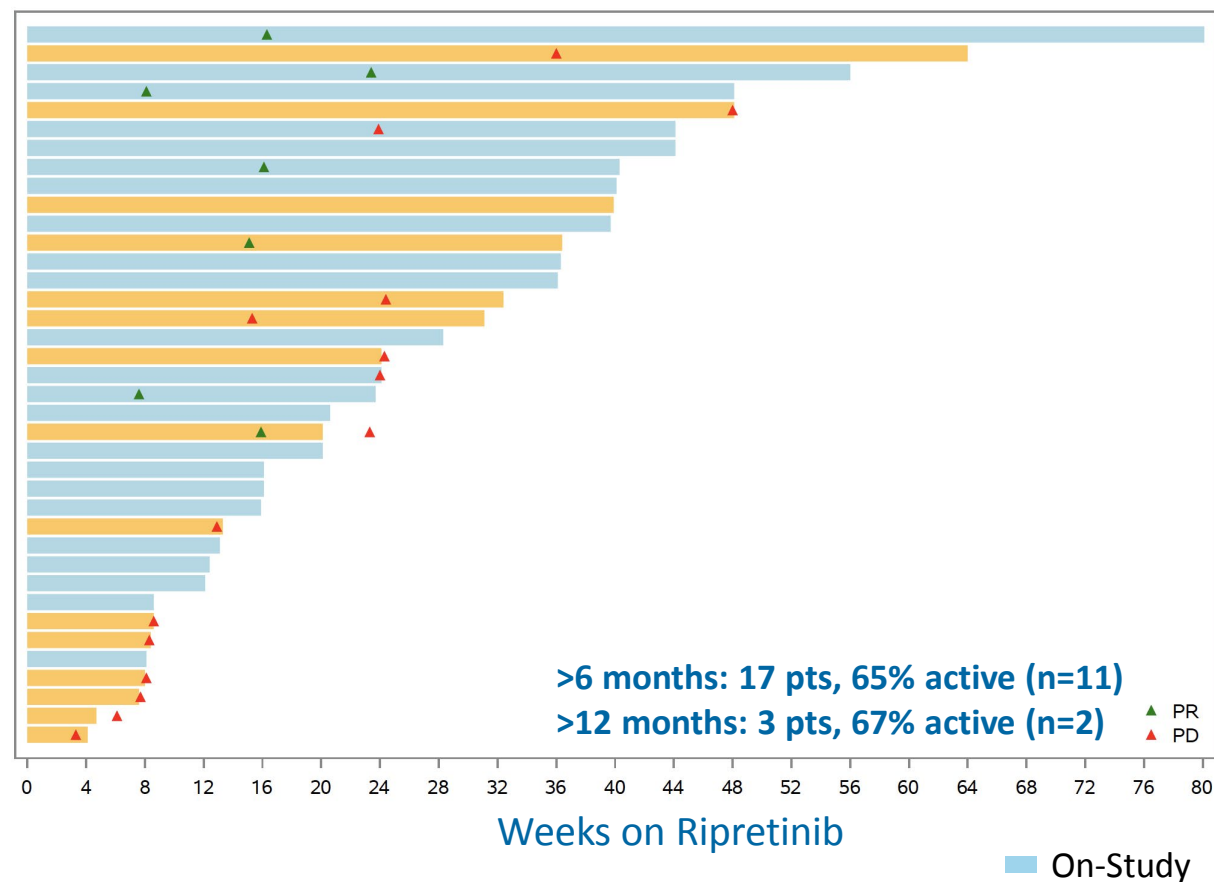
**Tumor Control per RECIST<sup>(1)</sup>  
KIT & PDGFRα @ ≥ 100 mg/d (n=178)**



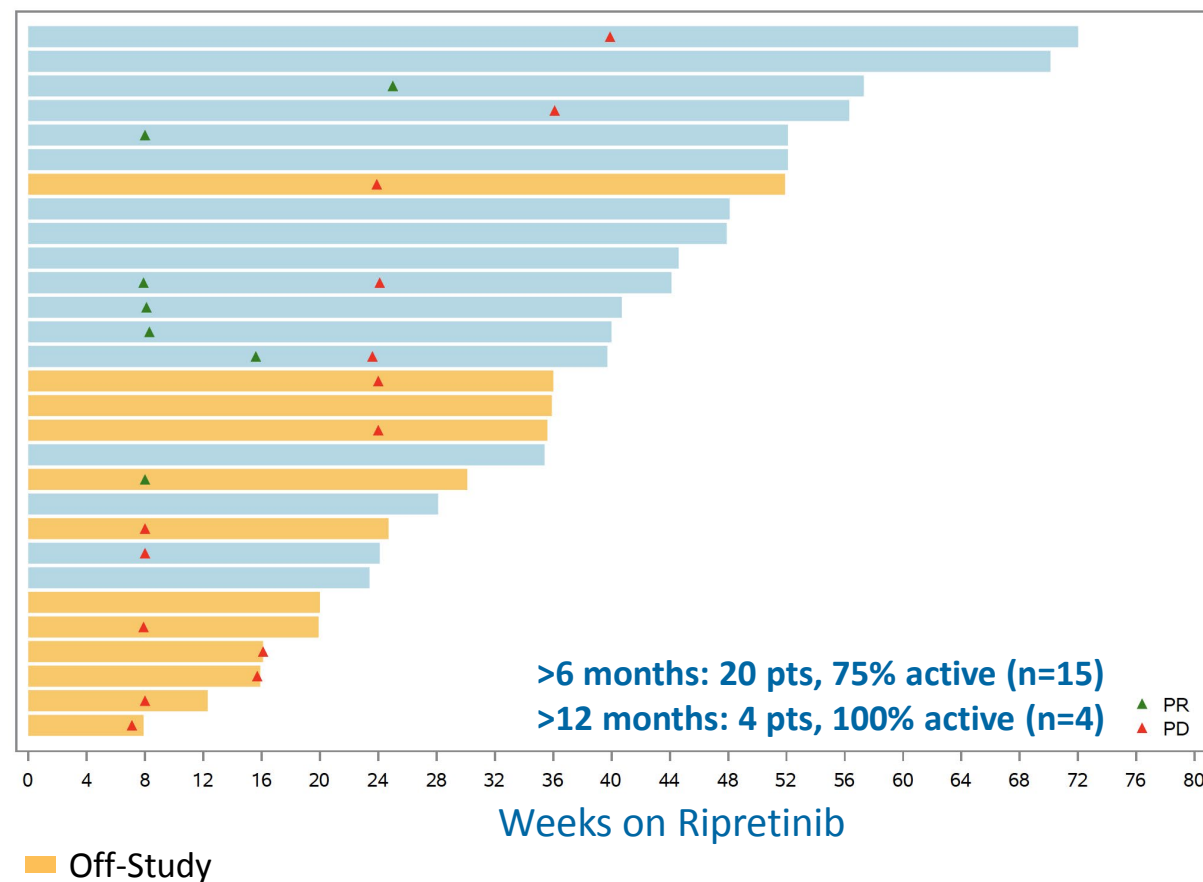
2 <sup>nd</sup> Line	38	30	21	13	8	2	1	1	1	0	0	0	0
3 <sup>rd</sup> Line	29	24	21	14	8	4	1	1	0	0	0	0	0
≥4 <sup>th</sup> Line	111	71	53	32	20	14	12	6	5	3	1	1	0

# Ripretinib: Generally Well-tolerated in Phase 1 Data; Allowed for Prolonged Treatment Duration in 2<sup>nd</sup> & 3<sup>rd</sup> Line GIST Patients at ≥100 mg/d (n=67)

## 2<sup>nd</sup> Line KIT and PDGFRα Patients (n=38)<sup>(1)(2)</sup>



## 3<sup>rd</sup> Line KIT and PDGFRα Patients (n=29)<sup>(1)(2)</sup>





# Ripretinib: Favorable Tolerability Profile @ ≥100mg Daily

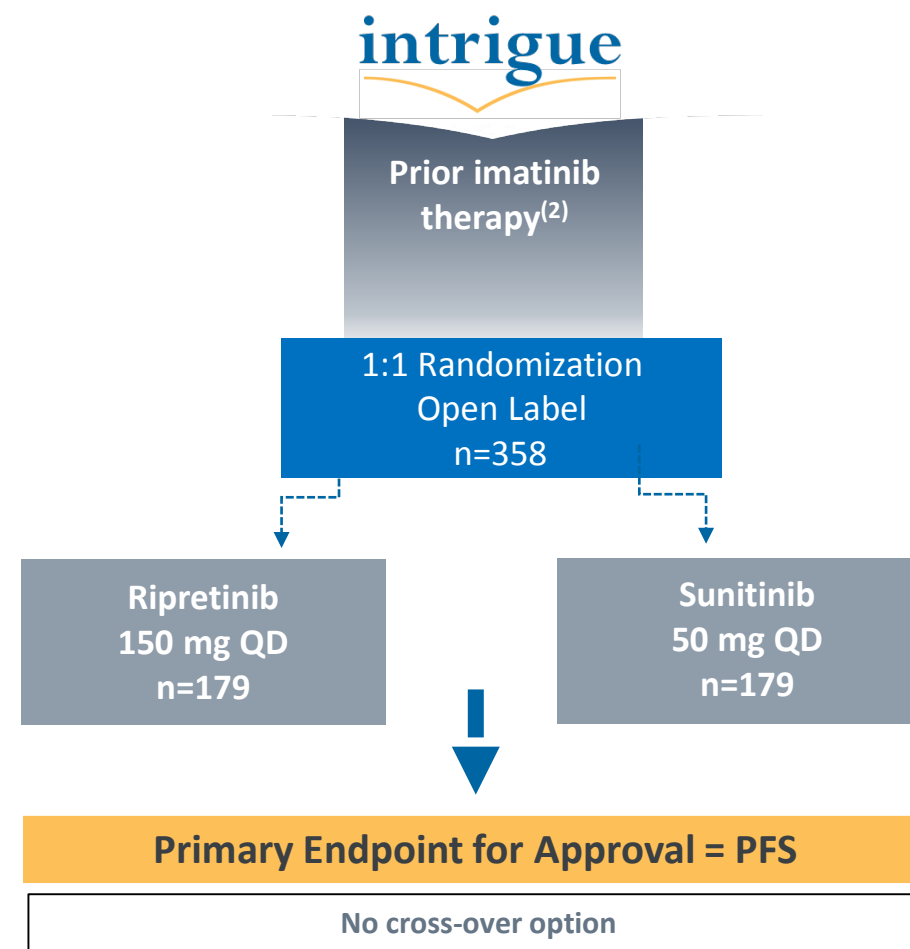
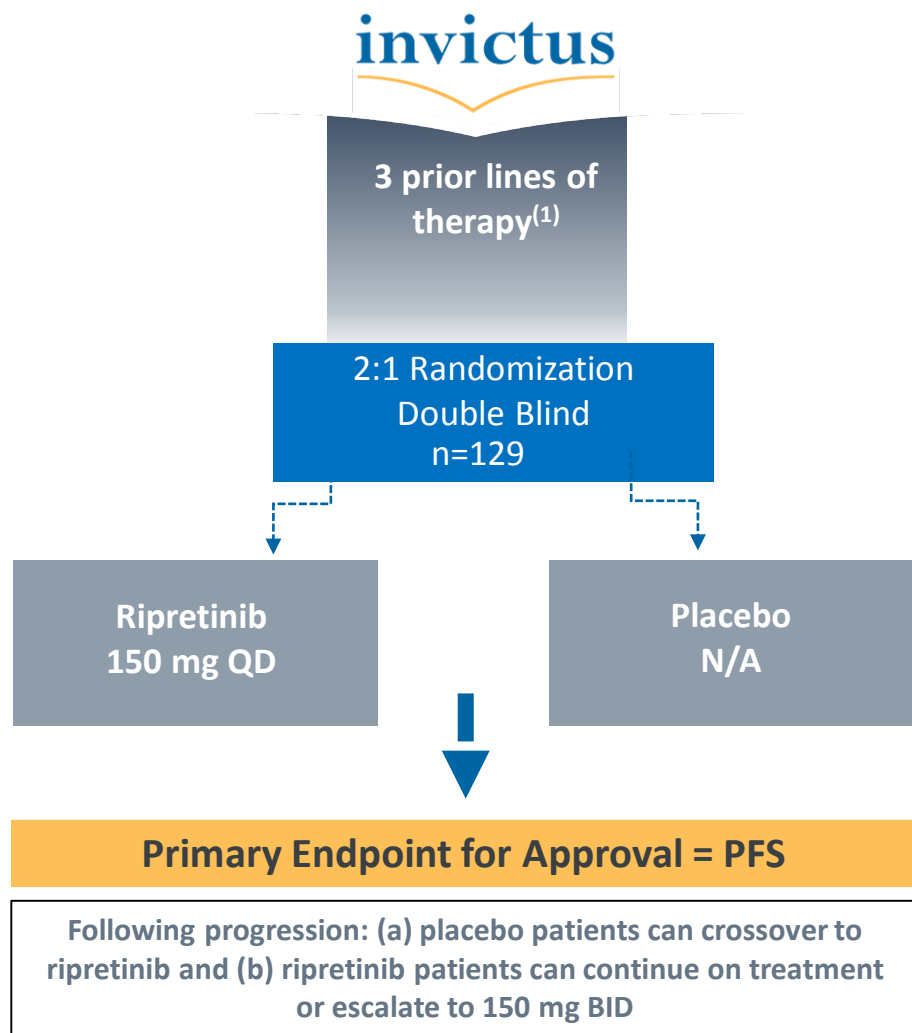
## Treatment-emergent Adverse Events (TEAE) in >10% GIST Patients (n=178) @ ≥100 mg Daily

- 14% (24 of 178) patients experienced dose reductions due to TEAEs
- 11% (19 of 178) patients experienced treatment discontinuations due to TEAEs
- Clinically asymptomatic lipase elevations most frequent G3 TEAE

GIST PATIENTS @ ≥100 MG DAILY			
ADVERSE EVENT	GRADE 1-2 (N=178)	GRADE 3-4 (N=178)	GRADE 1-4 TOTAL (N=178)
Alopecia	89 (50%)	0 (0%)	89 (50%)
Myalgia	79 (44%)	0 (0%)	79 (44%)
Fatigue	74 (42%)	2 (1%)	76 (43%)
Constipation	60 (34%)	0 (0%)	60 (34%)
Hand-Foot Skin Reaction <sup>1</sup>	56 (32%)	1 (1%)	57 (32%)
Nausea	53 (30%)	0 (0%)	53 (30%)
Decreased appetite	47 (26%)	2 (1%)	49 (28%)
Weight decreased	43 (24%)	0 (0%)	43 (24%)
Abdominal pain	33 (19%)	8 (5%)	41 (23%)
Diarrhea	38 (21%)	3 (2%)	41 (23%)
Lipase increased	21 (12%)	20 (11%)	41 (23%)
Vomiting	32 (18%)	1 (1%)	33 (19%)
Arthralgia	32 (18%)	0 (0%)	32 (18%)
Hypertension	22 (12%)	10 (6%)	32 (18%)
Dry skin	31 (17%)	0 (0%)	31 (17%)
Rash	31 (17%)	0 (0%)	31 (17%)
Muscle spasms	30 (17%)	0 (0%)	30 (17%)
Anemia	14 (8%)	13 (7%)	27 (15%)
Dyspnea	25 (14%)	2 (1%)	27 (15%)
Cough	26 (15%)	0 (0%)	26 (15%)
Headache	25 (14%)	0 (0%)	25 (14%)
Dizziness	23 (13%)	0 (0%)	23 (13%)
Back pain	20 (11%)	2 (1%)	22 (12%)
Blood bilirubin increased	15 (8%)	6 (3%)	21 (12%)
Pain in extremity	21 (12%)	0 (0%)	21 (12%)
Dysgeusia	18 (10%)	0 (0%)	18 (10%)
Hypomagnesaemia	18 (10%)	0 (0%)	18 (10%)
Pruritus	18 (10%)	0 (0%)	18 (10%)

# Ripretinib: INVICTUS Pivotal Phase 3 Top-line Data Expected Mid-2019

## Global Pivotal Phase 3 GIST Programs



Notes: (1) Phase 3 pivotal study in  $\geq 4^{\text{th}}$  line patients who previously received at least imatinib, sunitinib, and regorafenib; (2) Phase 3 pivotal study in 2<sup>nd</sup> line patients who previously received imatinib.

# Ripretinib: Estimated GIST Market Opportunity: US, EU & Japan

	US 	EU & Japan  	Total (exc. ROW)
GIST KIT 4 <sup>th</sup> Line <sup>1&amp;2</sup>	~2,100	~4,100	~6,200
GIST KIT 2 <sup>nd</sup> Line <sup>1&amp;2</sup>	~2,600	~5,000	~7,600
GIST PDGFR $\alpha$ <sup>1&amp;2</sup>	~400	~760	~1,160

Estimated Annual Incidence of New Patients by Indication

Sources: Internal Deciphera estimates based on applying epidemiology data reported in the following publications to population estimates for US, EU (28) and Japan:

<sup>1</sup> Zhao *et al.* J Gastrointest Oncol 2012;3(3):189-208

<sup>2</sup> Metaxas Y, *et al.* ESMO Open 2016

# Expanding Clinical Stage Portfolio

## Rebastinib

- Two ongoing Phase 1b/2 trials
- Data from Part 1 of the Phase 1b/2 paclitaxel study expected 2H 2019

## DCC-3014

- Phase 1 trial ongoing
- TGCT<sup>(1)</sup> expansion cohort planned in 2Q 2019
- Phase 1 escalation data update expected 2H 2019

## Discovery

- Select clinical candidate and initiate IND-enabling studies
- First disclosure of cancer metabolism target

# Rebastinib: A Highly Potent and Selective TIE2 Inhibitor

## Potent, small molecule inhibitor of TIE2

## Preclinical anti-tumor activity

- Single agent and I/O or chemo combination

## TIE2 microenvironment mechanisms

- Tumor vascularization, dissemination, metastasis, immunotolerance

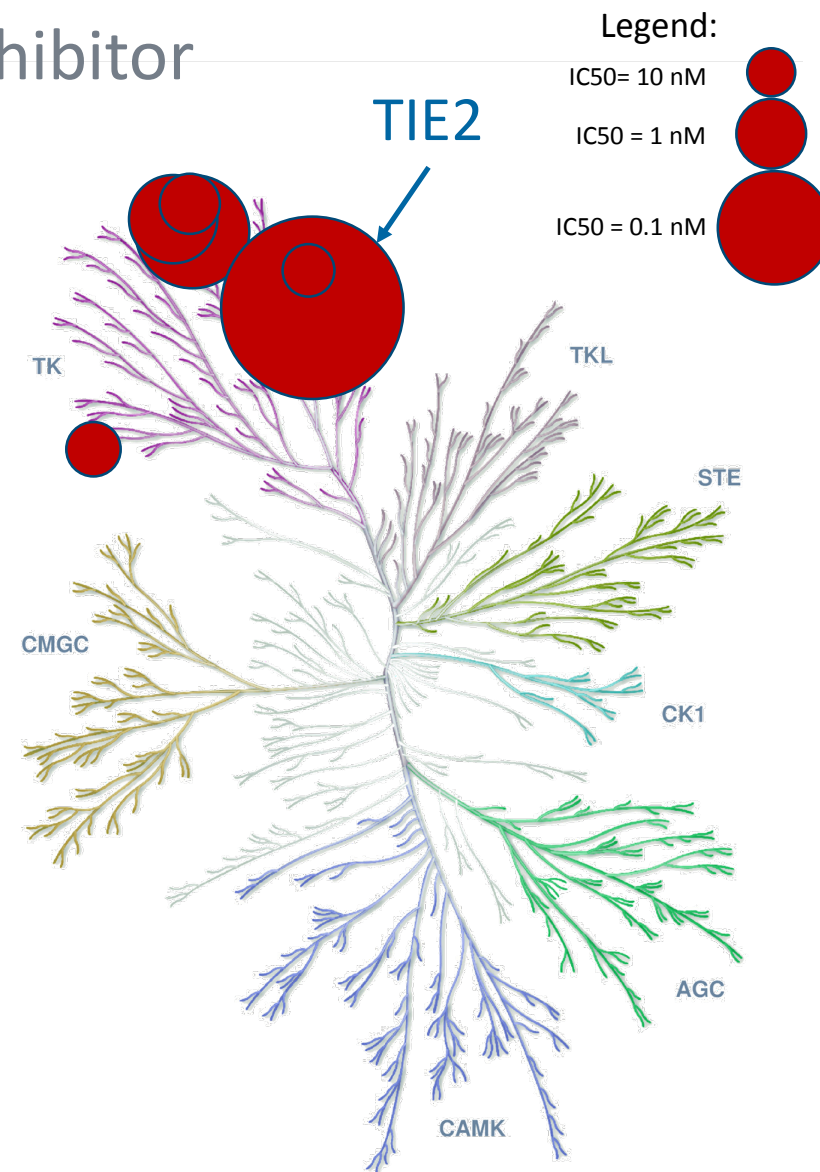
## Phase 1 study completed

- Identified 150 mg BID dose as maximum tolerated dose

## Development Status

- Two ongoing company-sponsored chemo combo trials with paclitaxel and carboplatin
- Data from Part 1 of the Phase 1b/2 study with paclitaxel expected 2H 2019

## IP: Composition (2027) and method of use (2034)

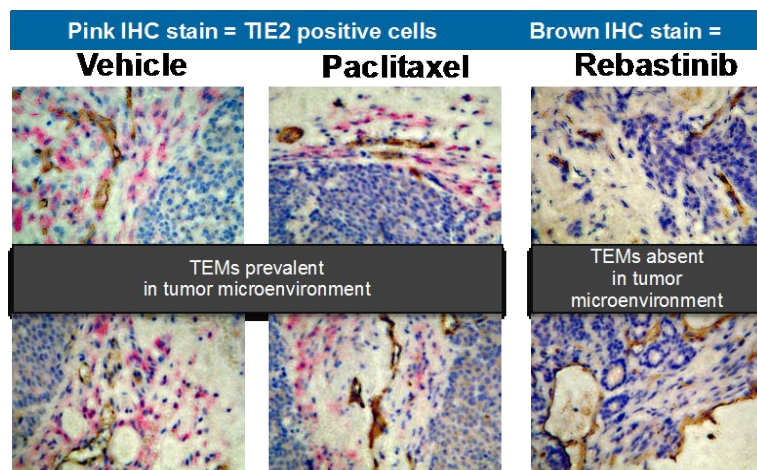




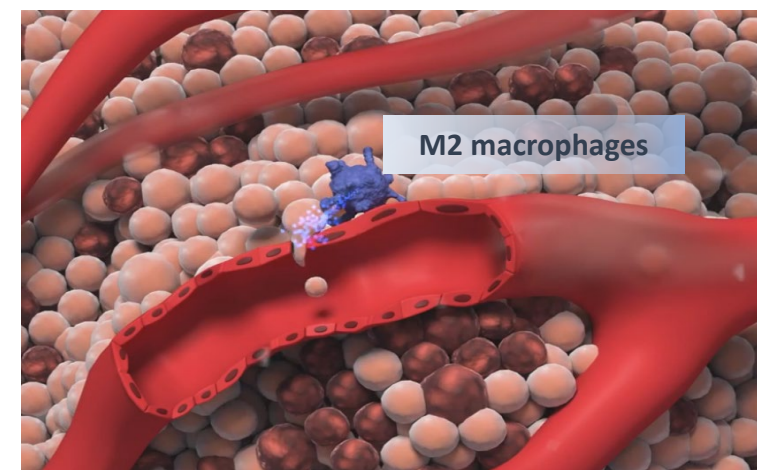
# Rebastinib: Potential Benefits in Combination with Chemotherapy

Rebastinib targets TIE2 expressing macrophages (TEMs) and endothelial cells in tumor environment

Combination of rebastinib with chemotherapy may increase tumor killing through multiple mechanisms

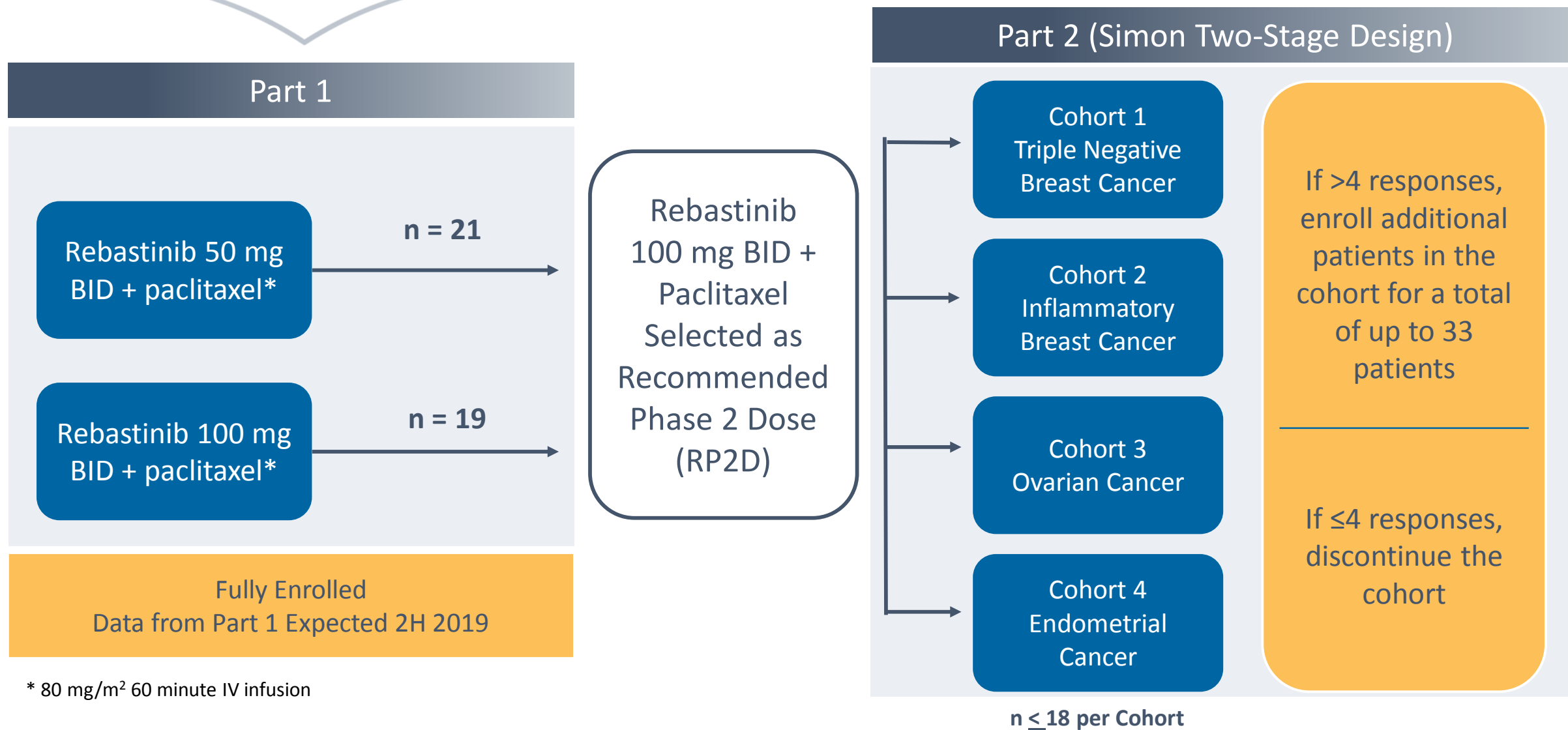


- Chemotherapy leads to recruitment of pro-tumoral M2 macrophages from the bone marrow
- Rebastinib is designed to inhibit chemotherapy-induced recruitment of M2 macrophages to tumors



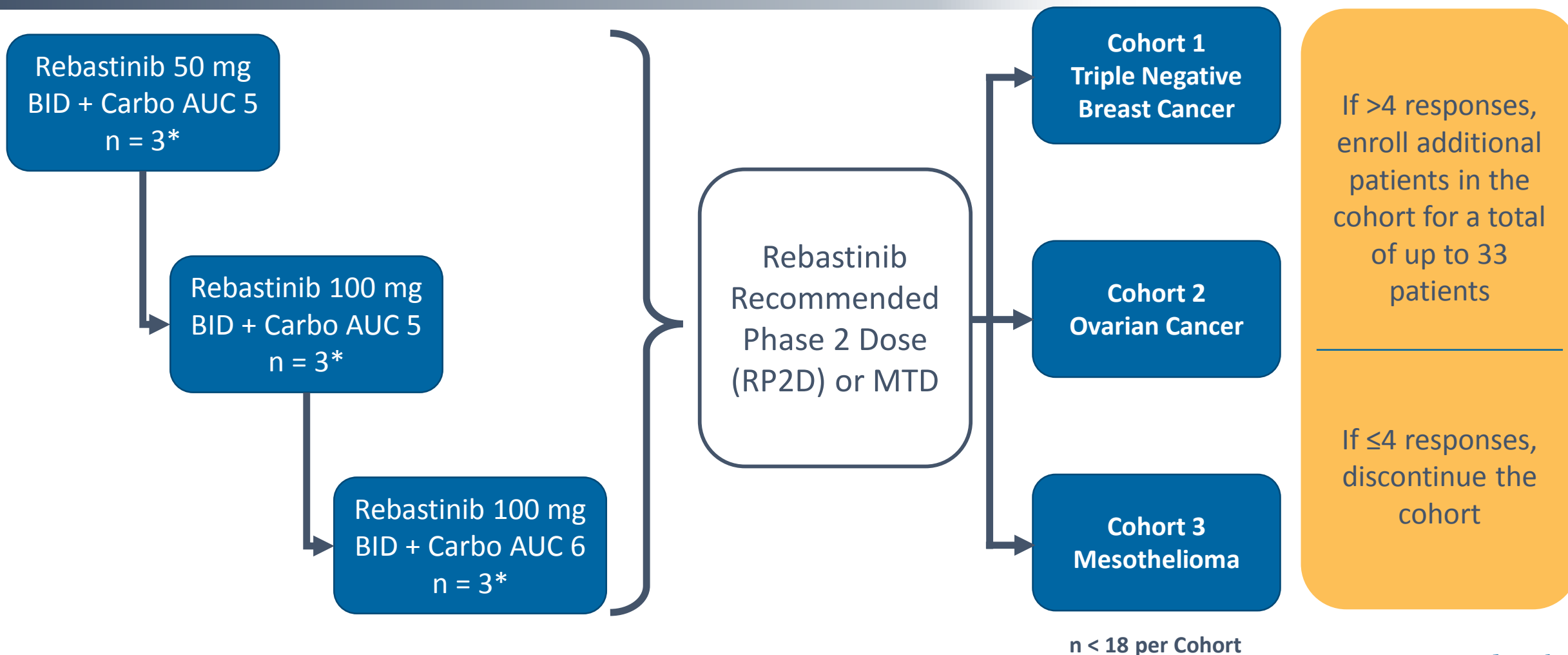
- M2 macrophages serve as pumps for tumor cell intravasation and metastasis
- Rebastinib is designed to block rebound angiogenesis and inhibit M2 macrophages

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



# Rebastinib Phase 1b/2 Study Combination with Carboplatin

## Simon Two-Stage Design Applied at MTD or RP2D



# DCC-3014: A Highly Selective and Potent CSF1R Inhibitor

**Phase 1 escalation trial ongoing, with data update in 2H 2019**

**Mechanistic Proof of Concept (mPoC)<sup>(1)</sup> Achieved**

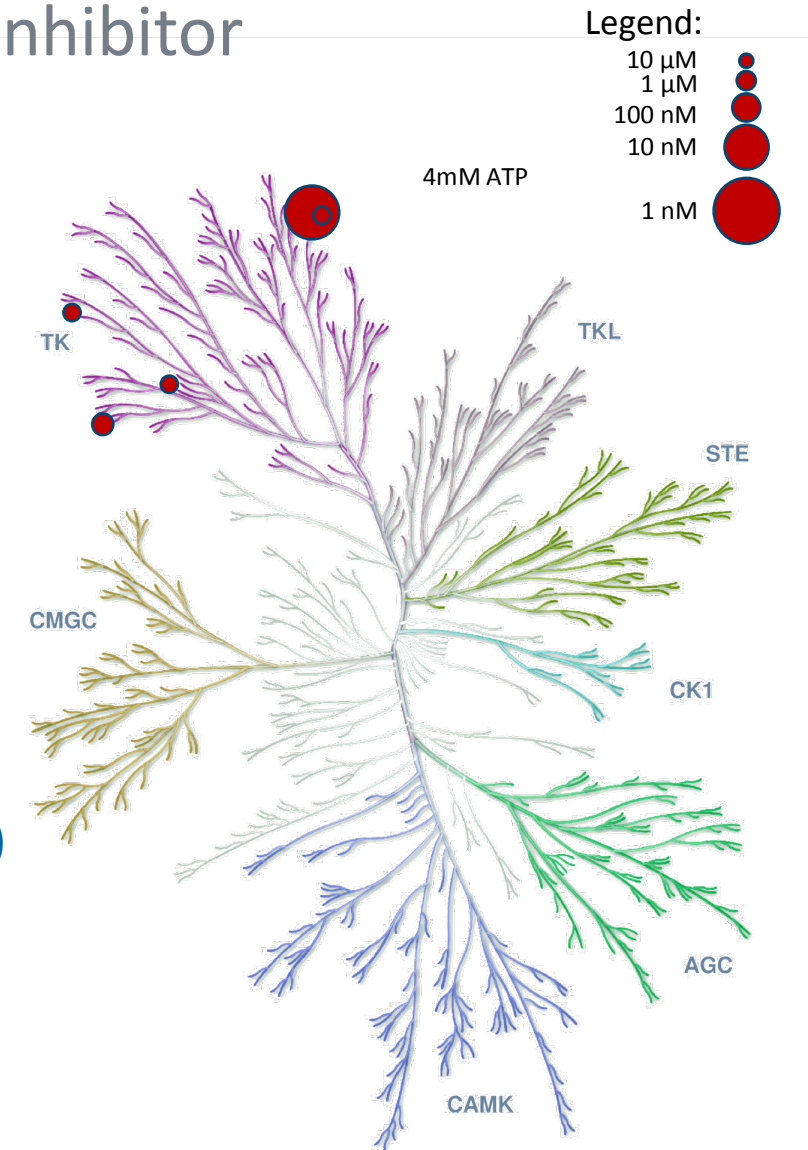
- Material reductions in circulating CSF1R+ macrophages

**Generally well-tolerated at doses of up to 30mg in patients receiving five-day loading, followed by twice-weekly maintenance regimen**

- No DLTs in loading and maintenance regimens
- No DCC-3014 related G3/4 TEAEs in  $\geq 10\%$  patients
- PK analysis demonstrated dose-proportional exposure for loading and maintenance regimen

**Study expanded to include patients with tenosynovial giant cell tumors (TGCT)**

**IP: Composition and method of use (2034)**



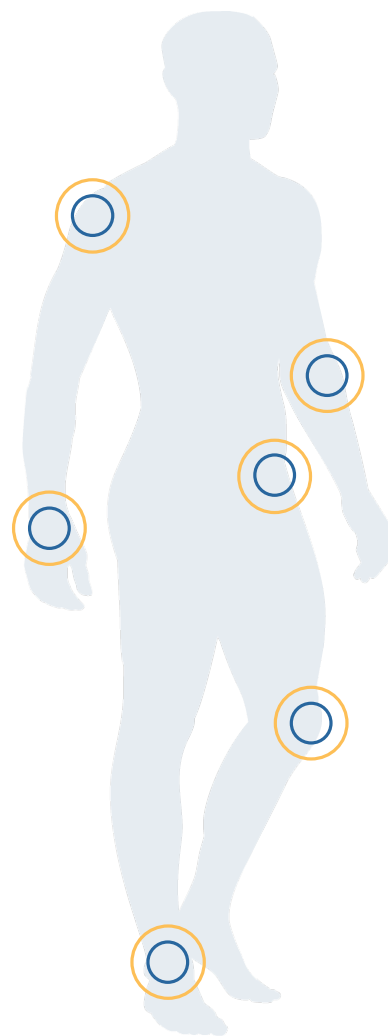
# Single Agent Opportunity in Tenosynovial Giant Cell Tumor (TGCT)

## Overview

- Rare, benign tumors involving the synovium, bursae and/or tendon sheath that damage surrounding tissues inducing pain, swelling, limitation of movement of the joint and cause severe disability
- Genetic translocation causes overproduction of CSF-1, the ligand for the CSF1R receptor, triggering migration of inflammatory cells to tumor sites

## Two Types of TGCT

1. **Localized TGCT**
  - Affects knee, wrist and ankle
2. **Diffuse TGCT (also known as PVNS)**
  - Mostly commonly affects the knee, as well as hip, ankle, elbow and shoulder



## Unmet Medical Need

- Surgical resection is standard treatment but with a high rate of recurrence in diffuse TGCT
- CSF1R inhibition has demonstrated clinical benefit in diffuse TGCT patients, but no currently approved systemic therapies
- Pexidartinib
  - 39% ORR VS. 0% for placebo in Phase 3 (n=120)
  - Hepatotoxicity concerns (off-target) may require a REMS and registry if approved
- Unmet medical need for effective CSF1R inhibitor with favorable safety/tolerability profile for diffuse TGCT patients



# Significant 2019 Milestones Across the Pipeline

## Ripretinib

INVICTUS ( $\geq 4^{\text{th}}$  Line GIST: Pivotal Phase 3 Results (Expected Mid-2019))

Phase 1 Expansion Data

## Rebastinib

✓ Phase 1b/2 Carboplatin Combination Initiated

✓ Part 1 of the Phase 1b/2 Paclitaxel Combination Completed Enrollment

Part 1 of the Phase 1b/2 Paclitaxel Combination Data

## DCC-3014

✓ Phase 1 Dose Escalation Presentation

Phase 1 Escalation Data Update

## Discovery Platform

Select Clinical Candidate & Initiate IND-enabling Studies

NASDAQ:DCPH

Shares Outstanding  
(as of 3/31/19)

38.2 MM (*basic*)  
45.0 MM (*fully-diluted*)

Cash, Cash Equivalents &  
Marketable Securities  
(as of 3/31/19)

\$262 MM

Cash expected to fund operating expenses and cap ex into 2H 2020



**Thank you to patients, caregivers  
and healthcare professionals who  
have participated in our clinical  
trials**

*May 2019*

