

# rexahn

PHARMACEUTICALS



## Rexahn Pharmaceuticals Overview

May 2018

# Safe Harbor Statement

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The statements that follow (including projections and business trends) are forward-looking statements. Rexahn's actual results may differ materially from anticipated results and expectations expressed in these forward-looking statements, including as a result of certain risks and uncertainties, such as Rexahn's lack of profitability, the need for additional capital to operate its business to develop its product candidates; the risk that Rexahn's development efforts relating to its product candidates may not be successful; the possibility of being unable to obtain regulatory approval of Rexahn's product candidates; the risk that the results of clinical trials may not be completed on time or support Rexahn's claims; demand for and market acceptance of Rexahn's drug candidates; Rexahn's reliance on third party researchers and manufacturers to develop its product candidates; Rexahn's ability to develop and obtain protection of its intellectual property; and other risk factors set forth from time to time in our filings with the Securities and Exchange Commission. Rexahn assumes no obligation to update these forward-looking statements.

# Rexahn Pharmaceuticals

## The Company

## The Pipeline

## The Future



# Investment Highlights

- Clinical stage biopharmaceutical company developing novel targeted cancer therapeutics
  - Unique drug targets localized in cancer cells
  - Inhibit growth/proliferation of cancer cells but spare normal, healthy cells
- Attractive pipeline of novel agents:
  - RX-3117 has generated strong safety and encouraging efficacy data and is currently in Phase 2 clinical trials in metastatic pancreatic cancer and advanced bladder cancer
  - RX-5902 (Supinoxin™) a first-in-class compound with a novel mechanism of action, currently being evaluated in a Phase 2 clinical trial in triple negative breast cancer
  - Validating partnership to develop RX-0201 (Archexin®) for hepatocellular carcinoma
- Meaningful data expected in 2018/2019
- Strong leadership team with significant drug development and strategy experience



# Experienced Leadership Team

## Peter D. Suzdak Ph.D., Chief Executive Officer

- 25+ years biopharma experience; broad experience spanning pre-clinical, development, commercialization; 18 IND filings, 3 NDA submissions



## Douglas Swirsky CPA, CFA, President and Chief Financial Officer

- 10+ years experience; corporate financing, corporate strategy, financing, licensing, and M&A
- 9+ years experience in investment banking



## Ely Benaim M.D., Chief Medical Officer

- 25+ years biopharma experience; extensive clinical research experience in academia, government, and industry



## Lisa Nolan Ph.D., Chief Business Officer





- 25+ years experience in corporate strategy, M&A, funding partnerships, out-licensing, and in-licensing transactions



# Differentiated Therapeutic Approach

	Traditional Chemotherapy	PD1 / CAR T-Cell Therapies	Rexahn Therapies
Selectively targets cancer cells	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
Reduced adverse events	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
Convenient oral dosing (RX 3117 and RX-5902)	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
Active against toughest cancers	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input checked="" type="checkbox"/>
Synergistic with existing therapies	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input checked="" type="checkbox"/>
Broad spectrum of anti-cancer activity	<input checked="" type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>

# A Diversified Portfolio of Targeted Cancer Therapeutics

Drug Development Pipeline					
Drug Candidate	Cancer Indication	Market Opportunity*	Preclinical	Phase I	Phase 2
RX-3117	<u>Metastatic Pancreatic</u>	>\$3B			
	<u>Advanced Bladder</u>	>\$1B			
RX-5902 (Supinoxin™)	<u>Triple Negative Breast</u>	>\$6B			
RX-0201 (Archexin®)	<u>Hepatocellular Carcinoma</u>	>\$2B			



# Rexahn Pharmaceuticals

## The Company

## The Pipeline

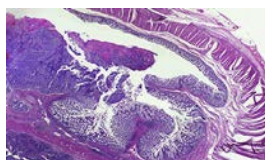
## The Future





# Advancing Our Product Pipeline

DRUG CANDIDATE	POTENTIAL INDICATION	STATUS	MARKET OPPORTUNITY <sup>(1)</sup>
RX-3117	Metastatic Pancreatic Cancer	Phase 2	>\$3B
	Advanced Bladder Cancer	Phase 2	>\$1B
RX-5902	Triple Negative Breast Cancer	Phase 2	>\$6B
RX-0201	Hepatocellular Carcinoma	Pre-clinical	>\$2B



# RX-3117: A Next Generation Nucleoside Compound

## The Candidate

- Small molecule nucleoside analogue similar to gemcitabine but cancer cell specific
- Prodrug activated by UCK2 which is predominantly present in cancer cells
- Active following oral administration
- Broad spectrum anti-cancer activity and active against gemcitabine resistant cancers

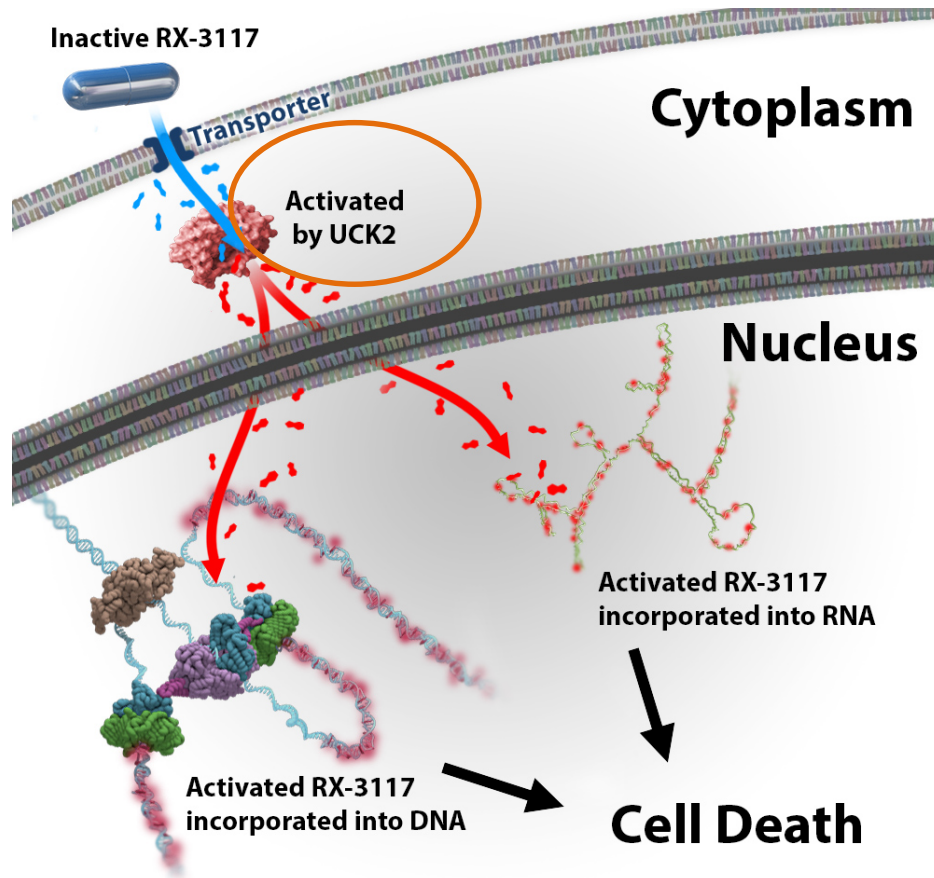
## Significant Unmet Medical Need

- Metastatic pancreatic and advanced bladder cancer
- Orphan drug designation for pancreatic cancer

## Clinical Development – Status

- Completed Phase 2a clinical proof-of-concept trial for metastatic pancreatic cancer (3<sup>rd</sup> line)
- Initiated Phase 2a combination study with Abraxane® in pancreatic cancer (1<sup>st</sup> line)
- Completing Phase 2a clinical proof-of-concept trial in advanced bladder cancer (3<sup>rd</sup> line)

# RX-3117: A Selectively Activated Nucleoside Analogue



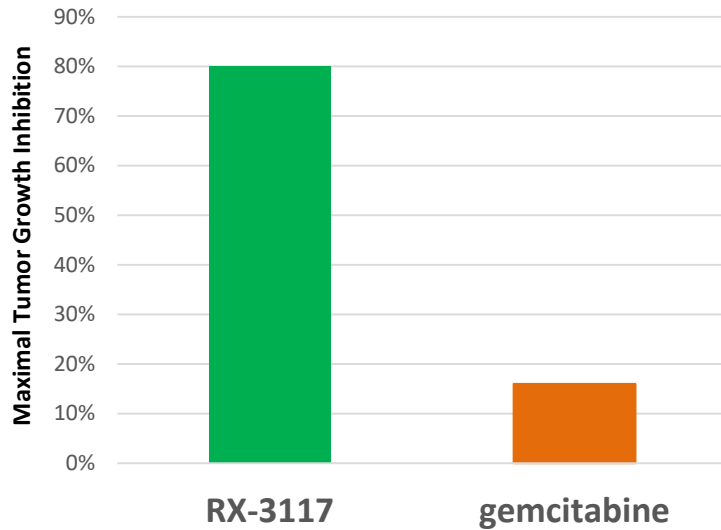
- Activation enzyme (UCK2) found only in cancer cells
- Targeted approach enhances safety and efficacy
- RX-3117 is active against cancers that have developed resistance to gemcitabine
- Proven mechanism of action reduces development risk

**RX-3117 MOA supports a biomarker strategy for patient selection**

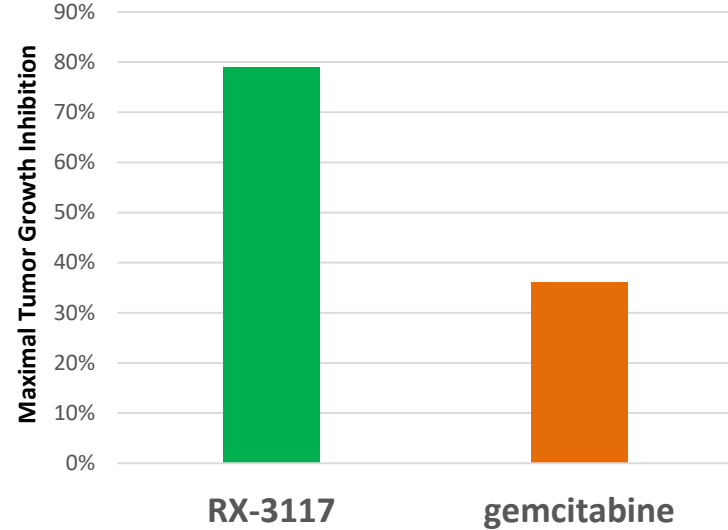
# Strong Preclinical Data Supports Lead Indications

RX-3117 is more effective than gemcitabine in preclinical models of pancreatic and bladder cancer

Tumor Growth Inhibition in a Patient-derived Bladder Cancer Model

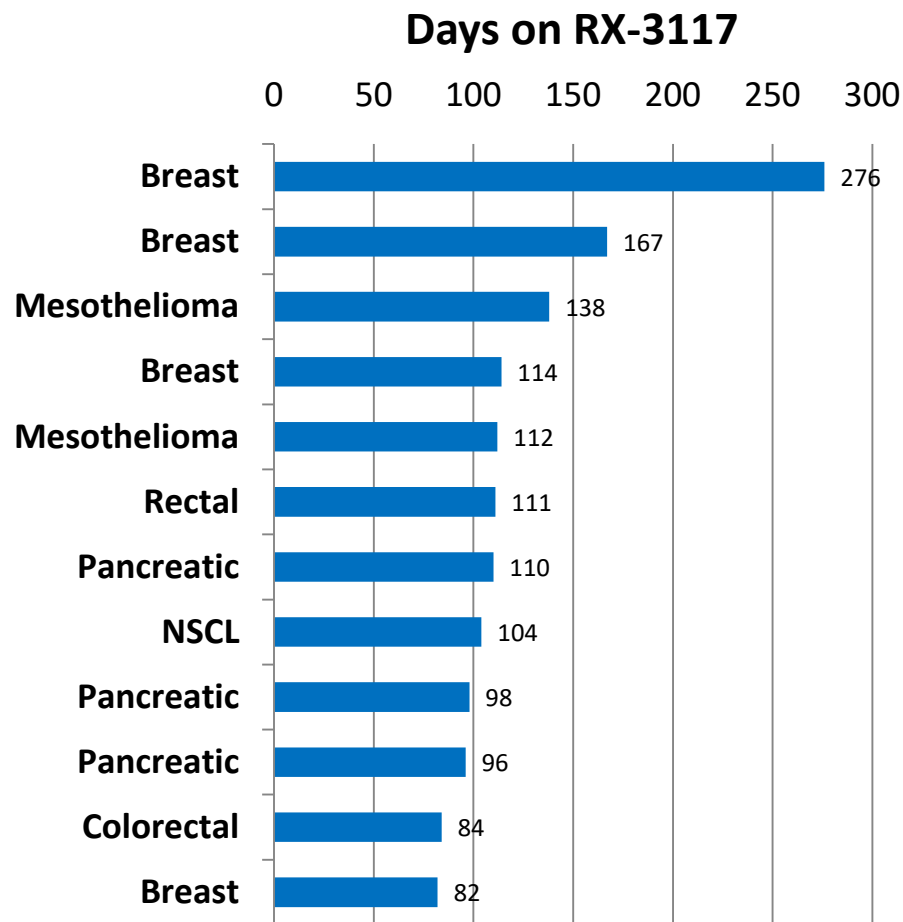


Tumor Growth Inhibition in a Patient-derived Pancreatic Cancer Model



# RX-3117 Completed Phase Ib Trial

- 74% of patients had received 3 or more prior cancer therapies
- Evidence of single-agent activity
- Stable disease achieved in 12 patients persisting up to 276 days
- Tumor burden reduction in 3 patients; Pancreatic, Breast (15% reduction in a liver lesion), Mesothelioma (9% in tumor volume)
- RX-3117 was safe and well tolerated at recommended Phase 2a dose with only mild/moderate adverse events



Data presented at ASCO 2016

# RX-3117 Development in Pancreatic and Bladder Cancer



## HIGH UNMET MEDICAL NEED

- Advanced Pancreatic Cancer – poor prognosis; short survival
- Advanced Bladder Cancer – limited treatment choices



## COMMERCIAL POTENTIAL

- **Pancreatic Cancer** – U.S. 49,000 new cases/40,000 deaths (2015)
- **Bladder Cancer** – U.S. 74,000 new cases/16,000 deaths (2015)



## REGULATORY / CLINICAL PATH TO MARKET

- Accelerated pathway for high unmet medical need indications
- Well defined development pathway



## PRECLINICAL / CLINICAL DATA

- Robust *in vitro* activity and *in vivo* data demonstrating efficacy
- Initial evidence of single agent activity of RX 3117 - stable disease and tumor response in ongoing Phase 2a clinical trials



# RX-3117 Pancreatic Cancer Clinical Development Plan

## Phase 2a MONOTHERAPY THIRD LINE

**Monotherapy**  
**Stage 1 n=10**

**Stage 2**  
**Expansion n=40**

Smallest market segment  
Completed Stage 2  
Data presented at 2018 ASCO-GI  
Facilitates progressing into first line  
therapy in combination with Abraxane

## Phase 2a COMBINATION WITH ABRAXANE FIRST LINE

Largest market segment  
Initiated Q4 2017  
**>\$3B Market Opportunity**

**Maximum Tolerated Dose  
of combination**  
**Stage 1 n=10**

**Combination**  
**Expansion n=25**



# Pancreatic Cancer Phase 2a Study Design

<b>Patient Population</b>	<ul style="list-style-type: none"><li>Relapsed and refractory metastatic pancreatic cancer</li></ul>
<b>Study Design</b>	<ul style="list-style-type: none"><li>Multicenter, open-label, single agent, 2-stage design</li><li>Part 1: 10 patients</li><li>Part 2: Option to enroll 40 additional patients if predefined primary efficacy criteria are achieved in part 1</li></ul>
<b>Dosing Schedule</b>	<ul style="list-style-type: none"><li>700 mg; 5x weekly dosing; 3 weeks on/1 week off</li></ul>
<b>Efficacy Criteria</b>	<ul style="list-style-type: none"><li><math>\geq 4</math> month increase in PFS in 20% of patients; or</li><li><math>\geq 1</math> PR or CR; or</li><li>For pancreatic cancer: tumor biomarker CA 19-9 (50% reduction), and improvement of overall clinical benefit in 20% of patients</li></ul>
<b>Endpoints</b>	<ul style="list-style-type: none"><li>Time to disease progression; overall response rate; duration of response</li><li>Pharmacokinetics and safety</li></ul>

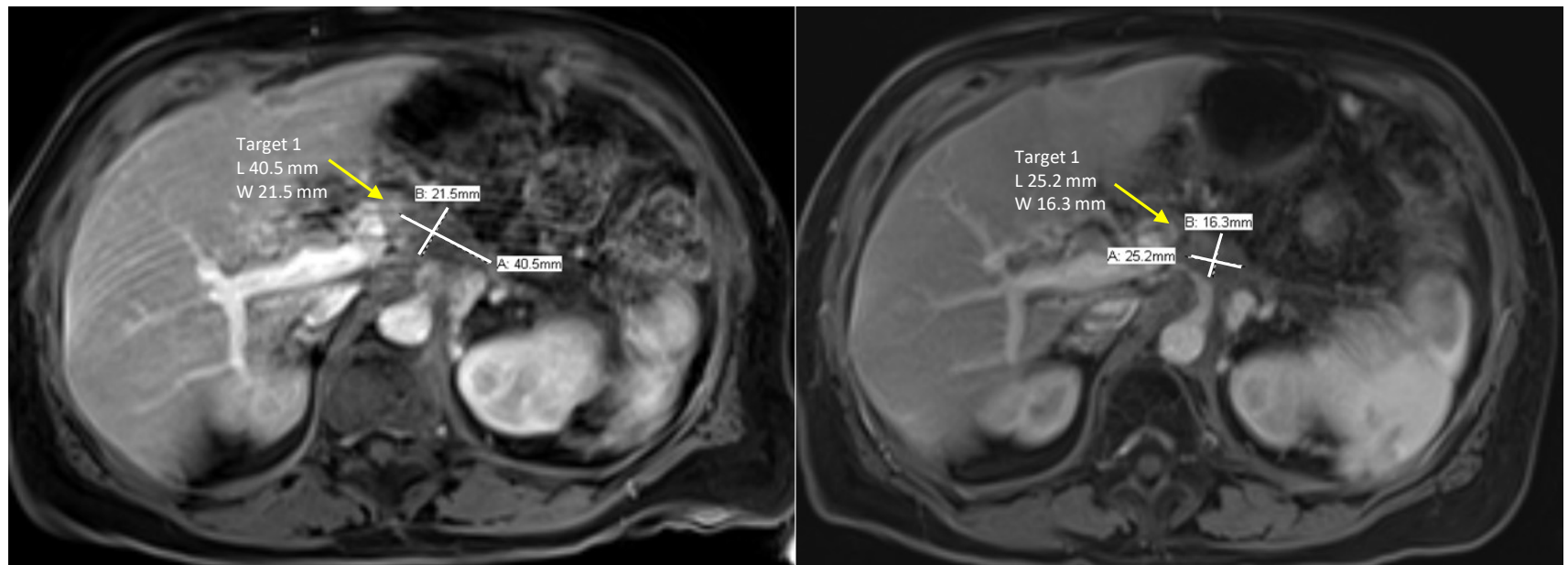
# RX-3117 Phase 2a Pancreatic Monotherapy Clinical Trial

- 46 patients, 43 response evaluable
- 78% percent of the patients had received two or more prior cytotoxic therapies
  - Expected median survival of 1.5 months
  - 93% of the patients had progressed after receiving gemcitabine therapy
- 31% of patients had disease stabilization for two months or more
  - Compares favorably to published data in second line patients where gemcitabine produced disease stabilization in 17% of the patients<sup>1</sup>
- One patient had a partial response (32.7% reduction in total tumor volume)
- Confirms that RX-3117 is safe and very well tolerated

1. Fernandes BM et al. J Clin Oncol 2017;35:4 suppl. 489

# RX-3117 Phase 2a Pancreatic Monotherapy Clinical Trial

- Prior Therapies:
  - Gemcitabine and Abraxane
  - FOLFIRI (folinic acid, 5-fluorouracil, irinotecan)
  - Rt salpingo-oophorectomy, pelvic mass, partial omentectomy, tumor debulking
  - Palliative radiotherapy to pancreatic primary mass
- 32.7% reduction in total tumor volume after 1 cycles of treatment)



CT scan with contrast: markings represent borders of tumor

A: Baseline image, B: Image following 1 cycle of treatment

# RX-3117: Phase 2a Combination Study with Abraxane® in Newly Diagnosed Pancreatic Cancer Patients

- Study initiated in Q4 2017
- Stage 1: Determination of maximum tolerated dose (MTD) of combination
  - Determination of doses for Stage 2 expected Q2 2018
- Stage 2: Expansion cohort (n=25) at MTD
- Preliminary efficacy read out Q4 2018

# RX-3117 Bladder Cancer Development Plan

## Phase 2a MONOTHERAPY THIRD LINE

**Monotherapy**  
Stage 1 n=10

**Stage 2**  
Expansion n=20

Monotherapy study in third line patients necessary to support studies in first line patients. Enrollment expected to complete Q1 2018

## Phase 2a COMBINATION WITH OTHER AGENTS IN FIRST LINE

Largest market segment  
Target indication is first line use  
Evaluating clinical protocols to advance RX-3117 into first line patients  
**>\$1B Market Opportunity**

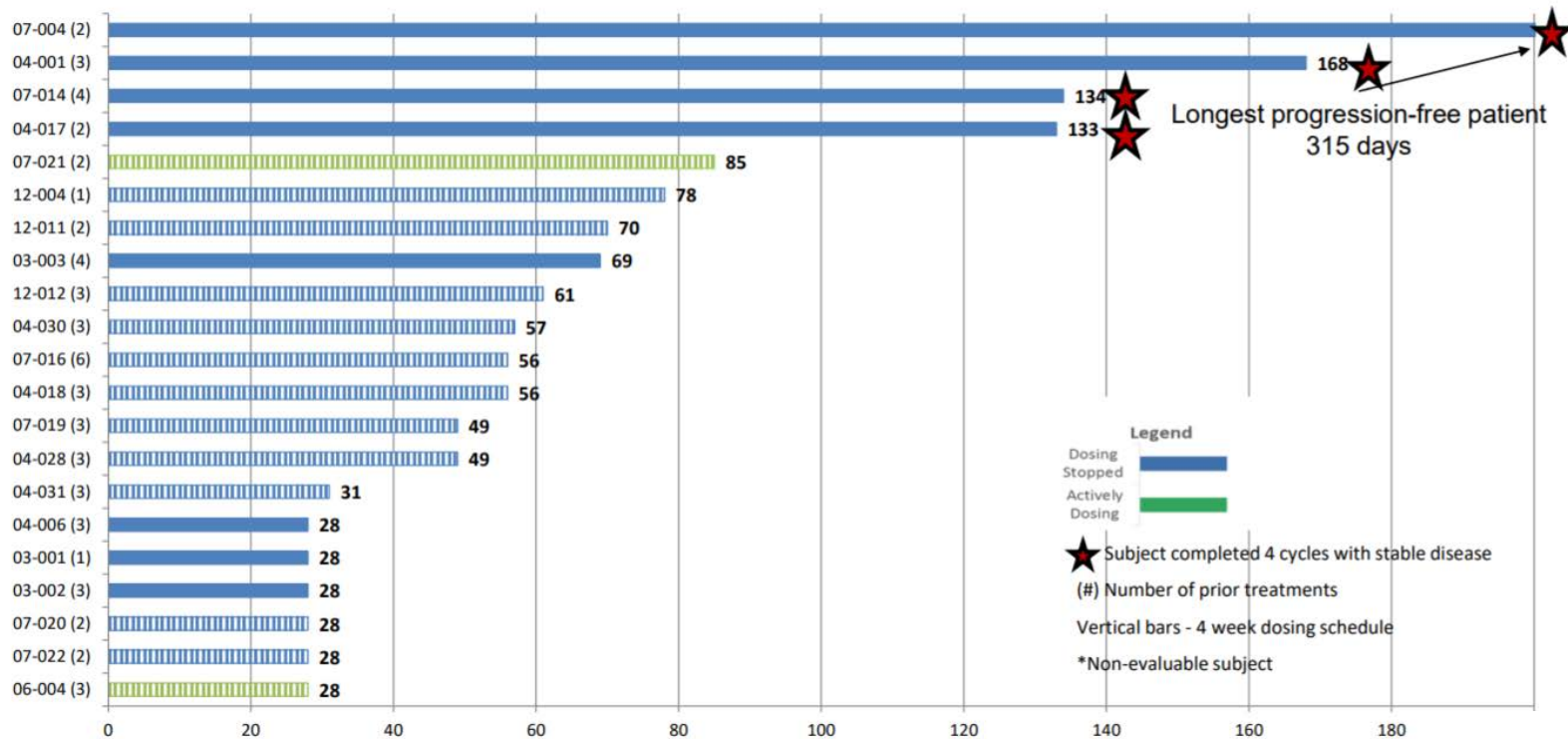
**Maximum Tolerated Dose**  
of combination  
Stage 1 n=10 -15

**Combination**  
Expansion n=50

# RX-3117 Phase 2a Bladder Cancer Clinical Trial Design

<b>Patient Population</b>	<ul style="list-style-type: none"><li>▪ Metastatic bladder cancer</li></ul>
<b>Study Design</b>	<ul style="list-style-type: none"><li>▪ Multicenter, open-label, single agent, 2-stage design</li><li>▪ Part 1: 10 patients</li><li>▪ Part 2: Option to enroll 20 additional patients if predefined primary efficacy criteria are achieved in part 1</li></ul>
<b>Dosing Schedule</b>	<ul style="list-style-type: none"><li>▪ 700 mg; 5x weekly dosing; 3 weeks on/1 week off</li></ul>
<b>Efficacy Criteria</b>	<ul style="list-style-type: none"><li>▪ <math>\geq 4</math> month increase in PFS in 20% of patients; or</li><li>▪ <math>\geq 1</math> PR or CR</li></ul>
<b>Endpoints</b>	<ul style="list-style-type: none"><li>▪ Time to disease progression; overall response rate; duration of response</li><li>▪ Pharmacokinetics and safety</li></ul>

# Phase 2a Data: RX-3117 Monotherapy in Advanced Bladder Cancer

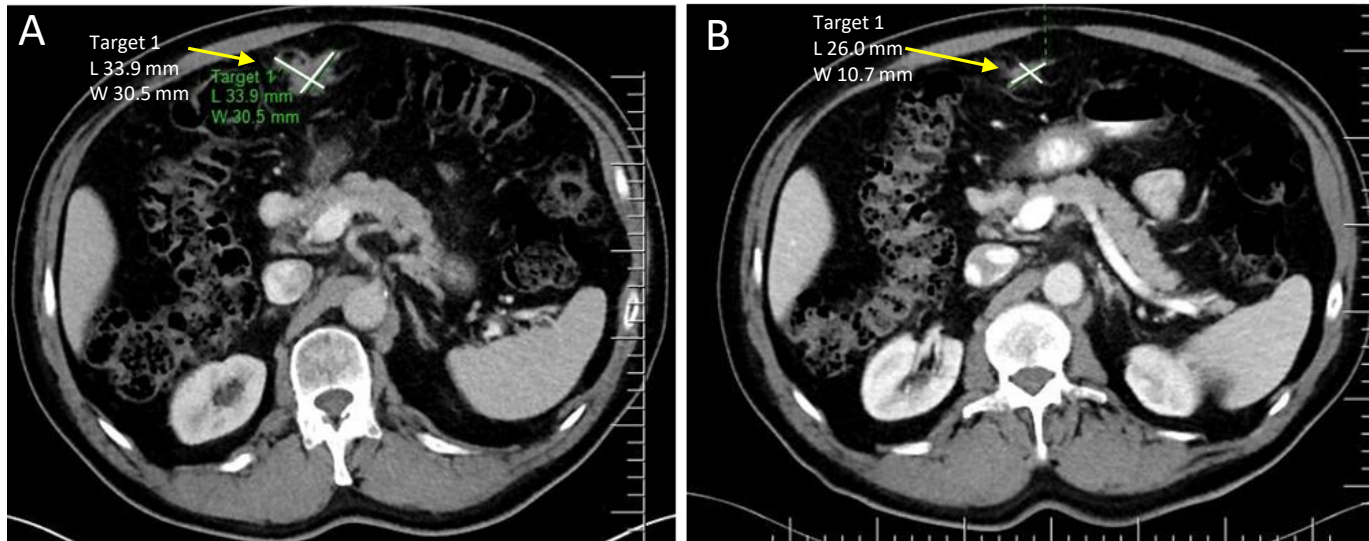


- 60% of patients had three or more prior cancer therapies, including gemcitabine in combination with cisplatin, carboplatin and/or immunotherapy
- Disease control at 8 weeks was 35%
- Four patients on treatment for >5 months without disease progression
  - One patient at 9 months without disease progression
- Tumor reduction seen in 5 patients



# RX-3117: Patient 06-004

- Prior Therapies
  - MVAC - 4 months; completed therapy
  - Herceptin® and pertuzumab for 25 months; discontinued due to disease progression
  - Oncolytic virus and pembrolizumab for 3 months; discontinued due to disease progression
- Following one cycle of treatment with RX-3117 there was a 23% reduction in tumor size



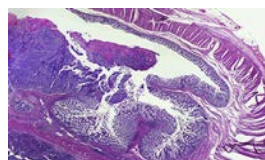
CT scan with contrast: markings represent borders of tumor

A: Baseline image, B: Image following 1 cycle of treatment

Data from ASCO GU January 2018

# Advancing Our Product Pipeline

DRUG CANDIDATE	POTENTIAL INDICATION	STATUS	MARKET OPPORTUNITY <sup>(1)</sup>
RX-3117	Metastatic Pancreatic Cancer	Phase 2	>\$3B
	Advanced Bladder Cancer	Phase 2	>\$1B
RX-5902	Triple Negative Breast Cancer	Phase 2	>\$6B
RX-0201	Hepatocellular Carcinoma	Pre-clinical	>\$2B



# RX-5902 is a Potential First-in-Class Inhibitor of a Unique Cancer Protein

## The Candidate

- Orally active, highly potent small molecule inhibitor of phosphorylated p68 (p-p68)
  - Modulates  $\beta$ -catenin/Wnt pathway
  - Increases immunogenicity of cancer cells

## Significant Unmet Medical Need

- Demonstrated activity in >100 human cancer cell lines including: triple-negative breast, ovarian, pancreas, non small cell lung cancer, colon, melanoma and renal cancer

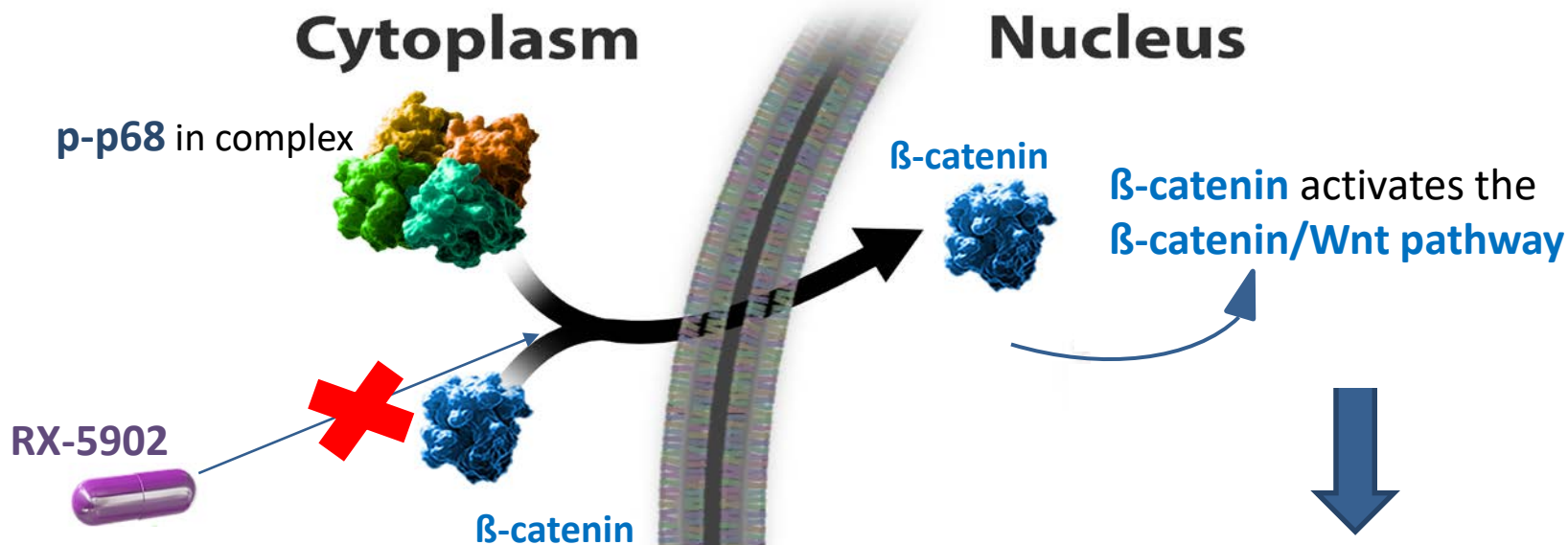
## Clinical Development – Status

- Completed Phase 1 clinical trial with RX-5902 in cancer patients and determined recommended Phase 2a dose
  - Final clinical data presented at ESMO (September 2017)
- Dosed first patient in a Phase 2a clinical proof-of-concept study in metastatic triple negative breast cancer
  - Initial data read out Q2 2018



# Novel Mechanism of Action on Key Cancer Pathway

p-p68 promotes cancer growth and metastases through  $\beta$ -catenin/Wnt pathway

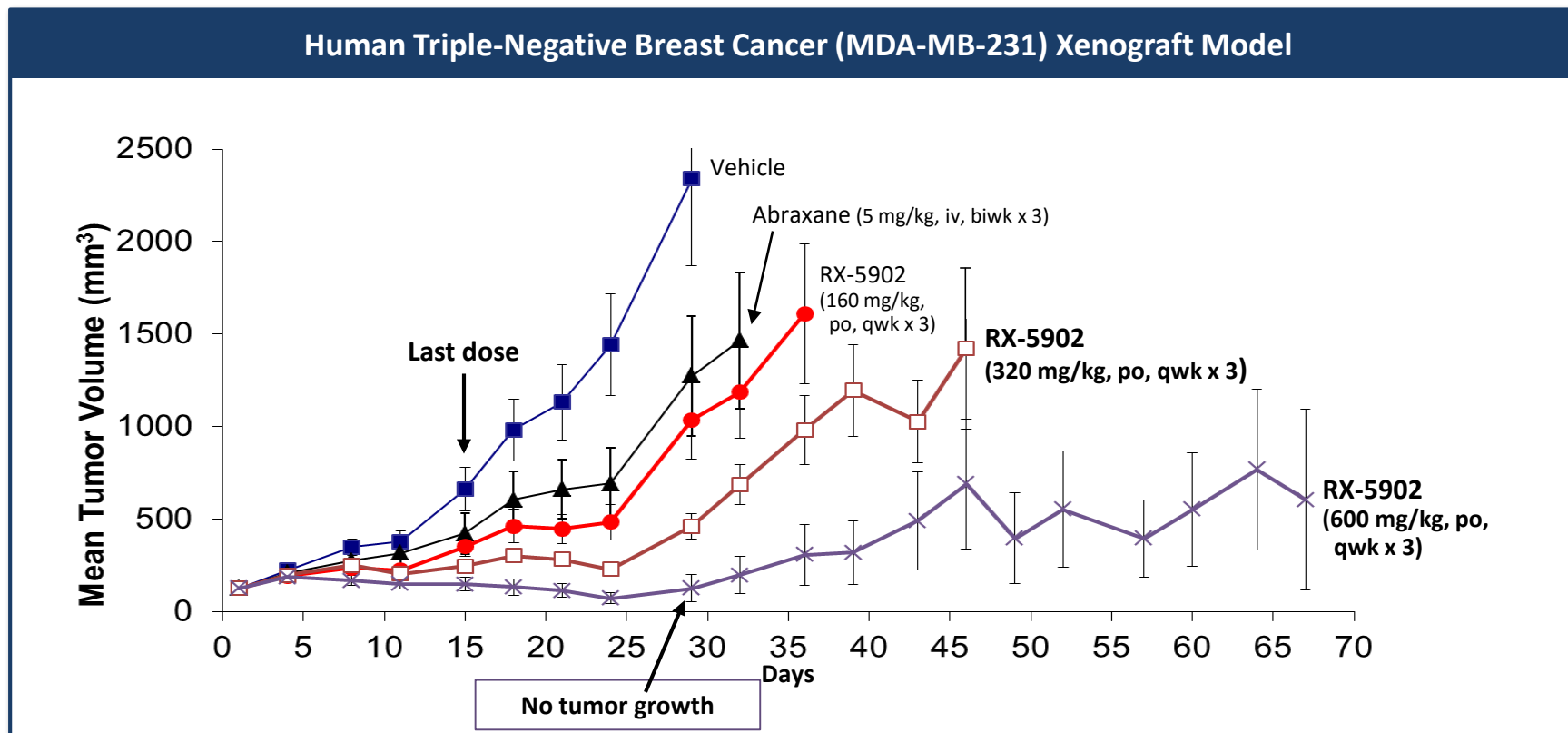


- p-p68 forms a complex with  $\beta$ -catenin and transports it into the nucleus
- RX-5902 binds to p-p68 and **inhibits**  $\beta$ -catenin transport into the nucleus
  - Decreased cancer cell growth
  - Increased cancer cell immunogenicity

Decreased T-cell infiltration,  
Increased oncogene expression,  
cancer cell proliferation,  
migration and metastases

p-p68 is found only in tumor cells

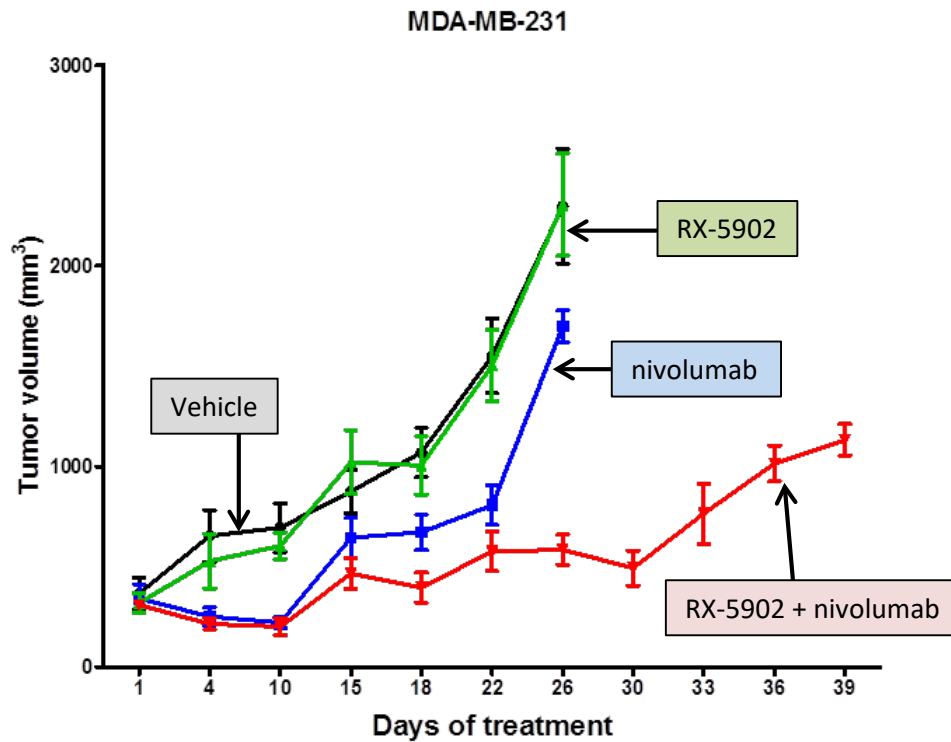
# RX-5902 inhibits the growth of human triple negative breast cancer cells in preclinical models



*Large opportunity: Triple negative breast cancer represents 20% of breast cancer diagnoses with limited treatment options; potential rapid path to market*

# RX-5902 Potentiates the Efficacy of a Checkpoint Inhibitor in Preclinical Models

- RX-5902 enhanced the efficacy of nivolumab (Opdivo®) in a humanized mouse xenograft model of TNBC
  - Modulation of the  $\beta$ -catenin/Wnt pathway by RX-5902 increases the effectiveness of nivolumab (Opdivo®)



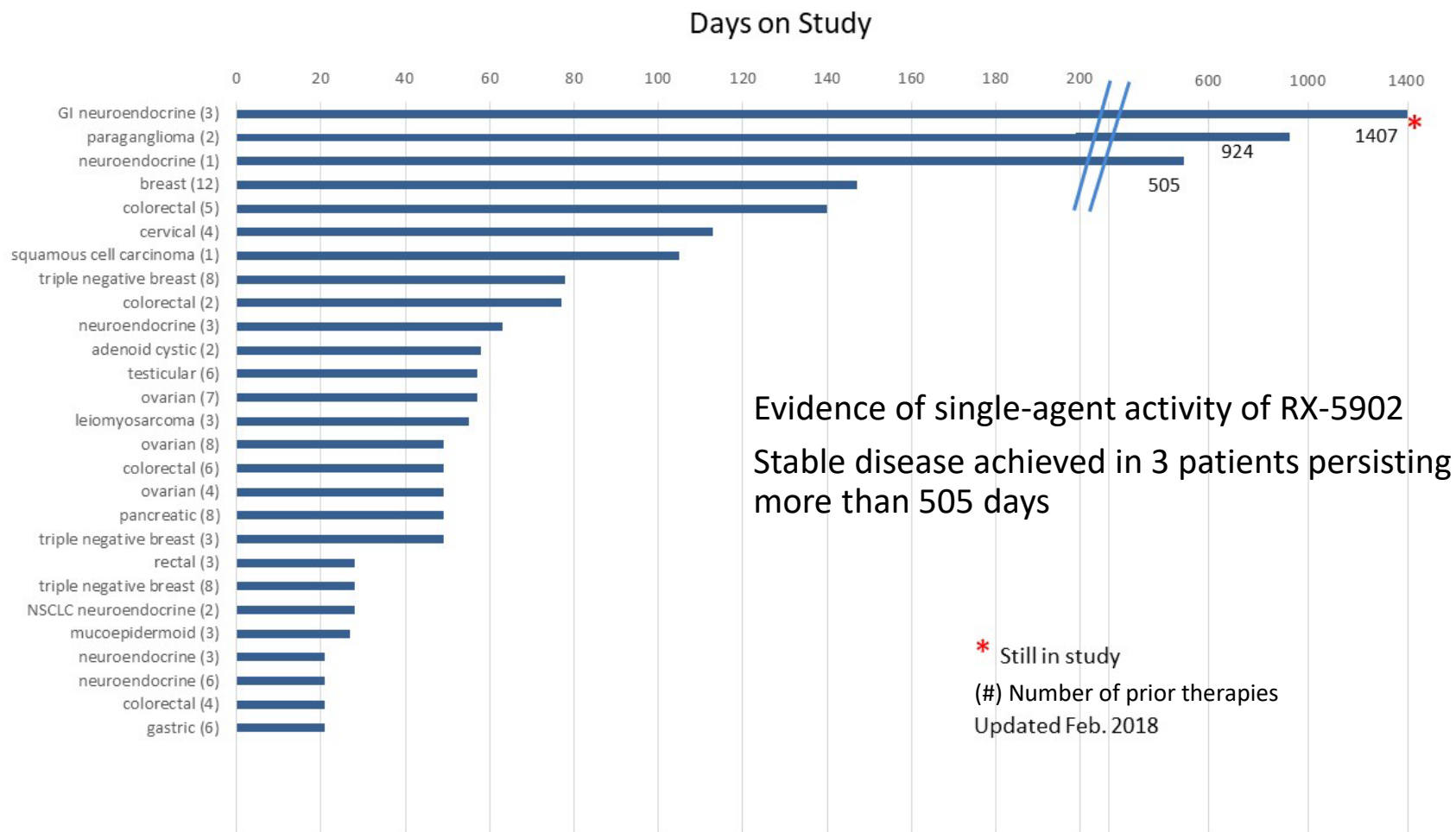
Treatment Group	Tumor Growth Inhibition*
Vehicle	-
RX-5902	0%
nivolumab (Opdivo®)	32%
<b>RX-5902 + nivolumab</b>	<b>85%</b>

\*Tumor growth inhibition at Day 26

\*Data presented at the San Antonio Breast Cancer meeting December, 2017

# Tumor Responses in Phase 1 Clinical Trial of RX-5902

- 64% of patients had received 4 or more, prior cancer therapies
- Safe and well tolerated with no Grade 3 or Grade 4 adverse events





# Triple Negative Breast Cancer Selected for Phase 2a Clinical Proof-of-Concept Study



## HIGH UNMET MEDICAL NEED

- Triple Negative Breast Cancer (TNBC) – no approved therapy; poor prognosis



## COMMERCIAL POTENTIAL

- TNBC represents approximately 15-20% of breast cancer diagnoses



## REGULATORY / CLINICAL PATH TO MARKET

- Accelerated / breakthrough regulatory pathway for high unmet medical need indications



## CLINICAL / PRECLINICAL DATA

- Robust *in vitro* activity and *in vivo* data demonstrating efficacy
- P-p68/  $\beta$ -catenin pathway implicated in TNBC

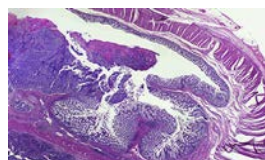


# RX-5902 Clinical Trial Status

- Phase 2a: Metastatic Triple Negative Breast cancer
  - Stage 1: enroll 10 metastatic triple negative breast cancer patients who have failed 2 (or more) prior therapies
    - Progression free survival (PFS) and tumor progression
    - Initial data read out Q2 2018
  - Stage 2: enroll additional patients, if warranted, based on the data readout from the initial cohort of patients
- Evaluating potential combination trials of RX-5902 with other anti-cancer agents in TNBC

# Advancing Our Product Pipeline

DRUG CANDIDATE	POTENTIAL INDICATION	STATUS	MARKET OPPORTUNITY <sup>(1)</sup>
RX-3117	Metastatic Pancreatic Cancer	Phase 2	>\$3B
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RX-5902	Triple Negative Breast Cancer	Phase 2	>\$6B
RX-0201	Hepatocellular Carcinoma	Pre-clinical	>\$2B



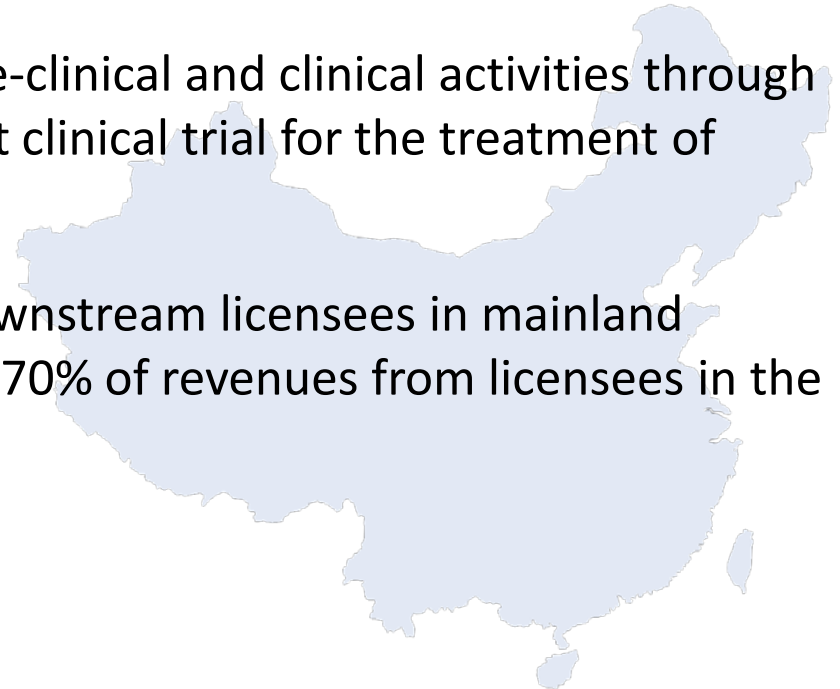
# RX-0201: Potential Best-in-Class AKT-1 Inhibitor

- Novel inhibitor of cancer cell signaling protein, AKT-1, increasing cancer cell death
  - Targets clinically validated cancer pathway
  - Inhibits drug resistance; synergistic with approved drugs
- PI3K/AKT-1/mTOR pathway involved in cancer cell growth and proliferation
  - AKT-1 Inhibition blocks the development of resistance to mTOR and Tyrosine Kinase inhibitors
  - Block the growth and proliferation of cancer cells
- Pre-clinical models of demonstrate potential of RX-0201
  - Decreases proliferation/growth of cancer cells
  - Decreases new blood vessel growth
  - Decreases drug resistance to existing therapies
- Re-focusing clinical development on hepatocellular carcinoma
  - Commercially viable market (>\$2B)
  - Advancing program through funded collaboration



# Zhejiang Haichang Biotechnology Collaboration

- Zhejiang Haichang Biotechnology Co., Ltd. is a privately owned specialized biotechnology company headquartered in Hangzhou, China, focused on the development and manufacture of complex intravenous pharmaceuticals
- Haichang will develop a nano-liposomal formulation of RX-0201 using its proprietary QTsome™ technology
- Haichang will fund the development of pre-clinical and clinical activities through completion of a Phase 2a proof-of-concept clinical trial for the treatment of hepatocellular carcinoma
- Rexahn retains 30% of economics from downstream licensees in mainland China, Hong Kong, Macau and Taiwan and 70% of revenues from licensees in the rest of the world



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# 2018 Key Milestones

Upcoming 2018 Clinical Milestones		Timing
RX-3117	Presentation of clinical data from <b>Phase 2a</b> clinical Proof-of-Concept trial in 3 <sup>rd</sup> line metastatic pancreatic cancer patients (monotherapy)	✓
	Determination of maximal tolerated dose (MTD) of combination of RX-3117/Abraxane in <b>Phase 2a</b> in newly diagnosed pancreatic cancer patients	2Q
	Interim clinical data from <b>Phase 2a</b> combination trial of RX-3117/Abraxane in newly diagnosed pancreatic cancer patients	4Q
	Completed data from <b>Phase 2a</b> trial of RX-3117 as 3 <sup>rd</sup> line treatment of metastatic bladder cancer patients	2Q
RX-5902	Interim clinical data from <b>Phase 2a</b> trial of RX-5902 in triple negative breast cancer patients	2Q
	Initiation of <b>Phase 2a</b> combination trial of RX-5902 in newly diagnosed metastatic triple negative breast cancer patients	4Q
RX-0201	Enter into collaboration to advance program using non-dilutive funding	✓



# Financial Summary

Exchange: Symbol	NYSE American: RNN
Common Shares Outstanding <sup>1</sup>	Outstanding: 31.7 million Fully Diluted: 42.7 million
Market Capitalization <sup>2</sup>	\$68 million
Resources <sup>3</sup>	Cash and Investments: \$19.3 million Estimated Quarterly Burn: \$5 million
Employees	17

<sup>1</sup>As of May 4, 2018; <sup>2</sup>Stock price as of May 7, 2018; <sup>3</sup>As of May 4, 2018

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