

Corporate Presentation

JANUARY 2021 | NASDAQ: BYSI



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BeyondSpring: Key Highlights



Mission

Committed to raising the standard of care for cancer patients in the largest global markets with first-in-class treatments that improve lives and clinical outcomes for millions of patients in need

2 near-NDA Assets Global Market Opportunities

PLINABULIN: Raising SOC in CIN & NSCLC

- First-in-Class immune agent
- New Chemical Entity
- ✓ IP through 2036 in 36 jurisdictions

CIN: Combo with G-CSF

- Final Ph 3 topline data Nov 2020
- NDA submission 1Q 2021
- Market: \$4.5B (US)
- Breakthrough Designation (US, China)

NSCLC: Combo with docetaxel

- Final Ph 3 data 1H2021
- Early 2022 NDA submission
- ✓ \$30B+ global market

Broad Pipeline

PLINABULIN: A pipeline in a drug

- Triple combo w/IO agents and radiation/chemo
- Expansion to additional solid tumors

Targeted Protein Degradation Platform

- Seed Therapeutics (Subsidiary)
- Collaboration with Eli Lilly

Three Pre-Clinical IO Agents

Global Capabilities Continuous Innovation

Strong clinical development

- Enrolled 1,000+ patients to final filing stage for CIN and NSCLC
- Dual U.S. and China development strategy
- ✓ Strong clinical investigator network

Deep Regulatory Expertise

Commercialization Planning Underway

Experienced leadership team with 50+ product launches



Strong network of advisors



Douglas Blayney, M.D.

Principal Investigator, CIN Study 105 & Study 106

- Founding member and former Board Member of the NCCN Guidelines for Neutropenia Management in U.S.
- Former president of ASCO

Management in China

Academy of Medical Sciences

• Former member of FDA's Oncologic Drugs Advisory Committee

Director of Oncology Department at Cancer Hospital Chinese

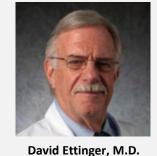
Medical Director of Stanford Cancer Institute

Principal Investigator, CIN Study 105 & 106 China

Chairman of the NCCN Guidelines for Neutropenia



Jeffrey Crawford, M.D.



SAB Member, NSCLC Study 103

U.S.

DSMB Chairman, CIN Study 105 & Study 106

Professor of Medicine at Duke University

• Chairman of NCCN Guideline for NSCLC and Board of Directors of NCCN Guideline

Chairman of NCCN Guidelines for Neutropenia Management in

• Lead investigator of the U.S. multicenter, randomized trial of Filgrastim (G-CSF, Neupogen), leading to FDA approval

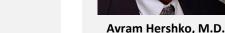
• Alex Grass Professor of Oncology, Sidney Kimmel Comprehensive Cancer Center at Johns Hopkins University

Yuankai Shi, M.D.



Principal Investigator, NSCLC Study 103 China

- Chairman of the NCCN Guidelines for NSCLC in China
- Co-founder of the Steering Committee of the Chinese Society of Clinical Oncology (CSCO)
- Director of GCP Center at Cancer Hospital of Chinese Academy of Medical Sciences



SAB Member, Ubiquitination Platform

- Nearly 50 years of research leadership in ubiquitination pathway
- 2004 Nobel Prize in Chemistry for discovery of ubiquitinmediated protein degradation
- Distinguished Professor at Rappaport Faculty of Medicine at Technion in Haifa

Yan Sun, M.D.



Two Near-term NDAs & robust drug development pipeline

	Indication	Program	Trial name / collaborator	Preclinical	Phase 1	Phase 2	Phase 3	Commercial rights	Status/Ne>	t Milestone
Late stage	CIN (All cancer, all chemo)	Plinabulin + pegfilgrastim	PROTECTIVE-1 (Study 105)	Phase 3 primary endpoint met at interim analysis			Global ¹	China NDA Submission Q1	U.S. NDA Submission	
			PROTECTIVE-2 (Study 106)	Phase 3 primary e	ndpoint met at interim	analysis			2021 ¹	Q1 2021
	NSCLC (2 nd /3 rd line)	Plinabulin + docetaxel	DUBLIN-3 (Study 103)	Phase 3 second int	erim analysis complete	ed		Global ¹		Global Final Ph3 data H1 2021
Investigator-initiated IO	NSCLC (2 nd /3 rd line)	Plinabulin + nivolumab	Fred Hutch/Univ. Washington/UCSD					Global ¹	Finished phase 1	
	SCLC	Plinabulin + nivolumab + ipilimumab	Rutgers University					Global ¹	Ongoing	
	Multi-cancer (2 nd /3 rd line)	Plinabulin + PD-1/PD-L1 + radiation/chemo	MD Anderson						Initiate phase 1 in 7 cancers Q1 2021	
Investigator-initiated IO	Oral T cell co-stimulator	BPI-002						Global		
	IKK inhibitor	BPI-003						Global		
	Oral neo-antigen generator	BPI-004						Global		
Subsi diary	1 st target KRAS	Targeted Protein degradation (TPD, molecular glue)	Seed Therapeutics ³					Global ¹		

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Note: ¹ Rolling submission basis in China. ² Global rights to Plinabulin ex-China. 58% ownership of Chinese subsidiary, Dalian Wanchunbulin Pharmaceuticals Ltd., which owns Chinese rights to Plinabulin. ³ Seed Therapeutics is a subsidiary of BeyondSpring Therapeutics.

Plinabulin differentiated tubulin binding and its effect in DC maturation and GEF-H1 release published in Chem and Cell Reports

Chem

CellPress

Giuseppina La Sala, Natacha

José Fernando Díaz, Michel O.

Plinabulin is a phase 3 anticancer

Olieric, Ashwani Sharma, ...

Steinmetz, Andrea Cavalli

michel.steinmetz@psi.ch (M.O.S.)

and antineutropenia drug

Plinabulin binding to tubulin

We report crystal structures of

We performed thermodynamic

selectivity and mechanism of

and kinetic studies on plinabulin

plinabulin in complex with BII- and

differentiates it from other

andrea.cavalli@iit.it (A.C.)

HIGHLIGHTS

candidate

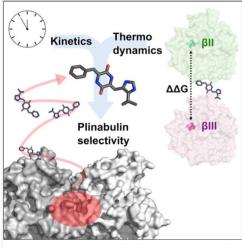
compounds

action

βIII-tubulin isotypes

Article

Structure, Thermodynamics, and Kinetics of Plinabulin Binding to Two Tubulin Isotypes



Plinabulin is a novel tubulin-binding agent that is currently in phase 3 clinical trials for cancer treatment and prevention of chemotherapy-induced neutropenia. Plinabulin binds within a distinct tubulin pocket, which differentiates it from other tubulin binders. Aimed at disclosing structural and energetic details of plinabulin binding to tubulin, we combine X-ray crystallography and computational modeling. We compare the plinabulin residence time with that of colchicine and combretastatin-A4. Our study helps understand potential mechanisms underlying differential effects of this family of anti-tubulin drugs.



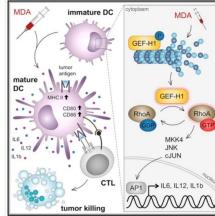
La Sala et al., Chem 5, 1–18 November 14, 2019 © 2019 Elsevier Inc. https://doi.org/10.1016/j.chempr.2019.08.022

Cell Reports

GEF-H1 Signaling upon Microtubule Destabilization Is Required for Dendritic Cell Activation and Specific Anti-tumor Responses

Graphical Abstract

Highlights



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Article

In Brief

Certain chemotherapeutics elicit potent anti-tumor immunity. Kashyap et al. demonstrate that microtubuledestabilizing chemotherapeutics induce maturation of dendritic cells through activation of microtubule-associated protein GEF-H1. This leads to effective priming of CD8 T cells against tumor antigens. GEF-H1 is critical for anti-tumor immunity of microtubule-targeting

chemotherapy.

 GEF-H1 is released from microtubules, leading to its activation

· Microtubule destabilization in dendritic cells drives DC

maturation and T cell activation

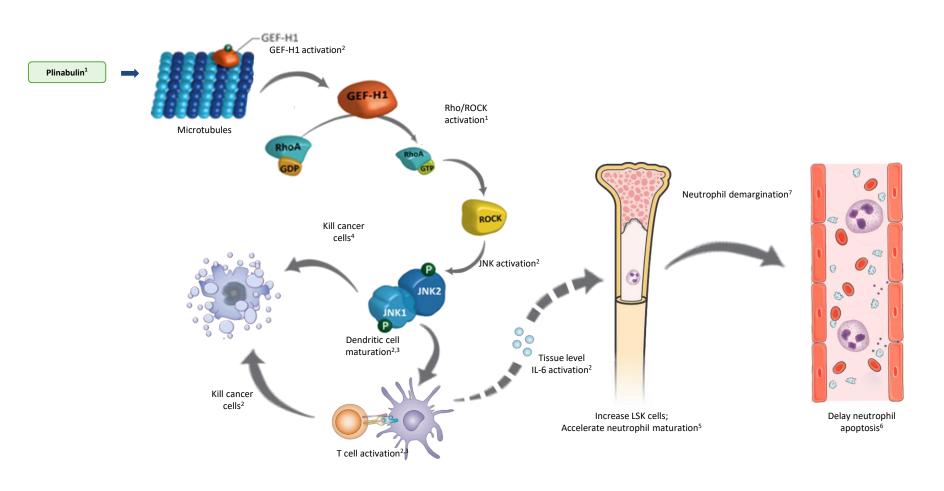
- GEF-H1 release triggers the RhoA-JNK-c-Jun signaling axis and AP-1 transcriptional response
- GEF-H1 is critical for DC maturation, antigen crosspresentation, and anti-tumor immunity

Kashyap et al., 2019, Cell Reports 28, 3367-3380 September 24, 2019 © 2019 The Author(s). https://doi.org/10.1016/j.celrep.2019.08.057





Plinabulin: first-in-class agent, stimulating innate and adaptive immune system (Proven Target: Immune Defense Protein GEF-H1)



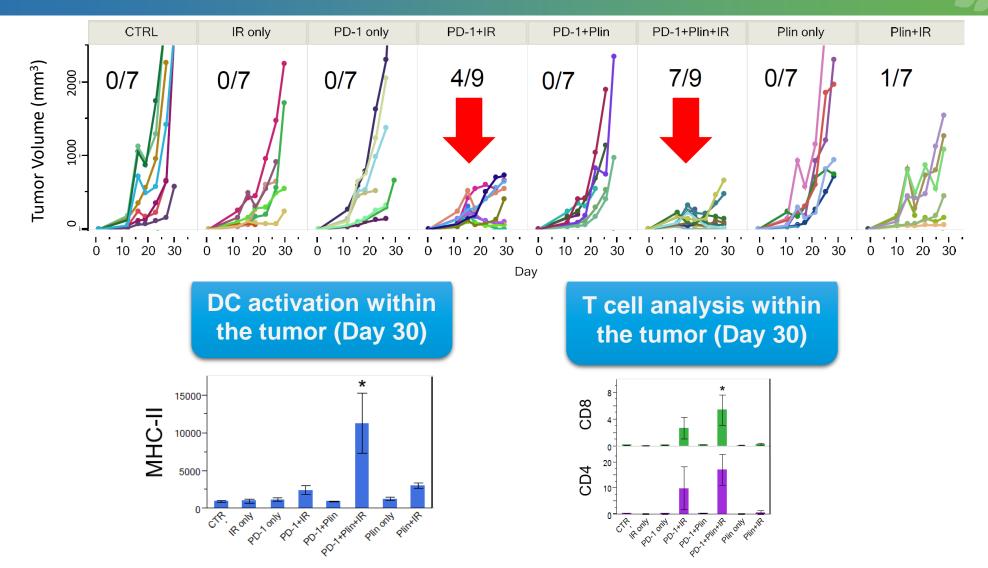
Plinabulin's immune mechanism designed to enable its effects in multiple cancer indications:

- Chemotherapy Induced Neutropenia (CIN): Designed to protect progenitor cells from chemo assault in bone marrow with week 1 benefit, which compliments G-CSF week 2 benefit for improved benefit potential
- NSCLC: Chemo (e.g. docetaxel) introduces real time tumor antigen, Plinabulin is designed to mature DC, leading to T cell activation, and durable anti-cancer benefit
- Multiple Cancer Indications: Triple combo combines "tumor antigen generation" from chemo/radiation, plinabulin "adding T cell gas", and PD-1/PD-L1 "release the brake" for potential maximum durable anticancer benefit



Note: ¹ La Sala et al., 2019 Chem. ² Kashyap et al., 2019 Cell Reports. ³ Zhang et al., 2005 Mol Cell Biol. ⁴ Singh et al., 2011 Blood. ⁵ Suwa et al., 2000 Am J Physiol Heart Circ Physiol; Ghosh et al., 2018 ACR Annual Conference; Blayney et al., Society of Leukocyte Biology. ⁶ Asensi et al., 2004 Infection and Immunity.

Triple Combo: Plinabulin + PD-1 + Radiation the Best Tumor Response in PD-1 non-responsive tumor model (MD Anderson)







Breakthrough Therapy Designation (US & China FDA) - NDA submission Q1 2021

Plinabulin + G-CSF

in Chemotherapy-Induced Neutropenia (CIN)

Plinabulin will add value to a large and growing CIN market - PD-1 + chemo approved, so chemo will not go away

Plinabulin + G-CSF in each cycle of chemo in non-myeloid cancers prevented or reduced the severity of neutropenia



U.S. Sales -- \$4.5 Billion²

As a combination therapy Plinabulin's base of business is G-CSF units

G-CSF cycles/year:

- U.S.: 1.3 million²
- Global: 4 million³

Unit growth (U.S.):²

- MAT Aug '19: 6.8%
- MAT Aug '20: 1.1%*

50%+ growth expected in use of first-line chemotherapy by 2040 worldwide⁴

Note: ¹ Centers for Disease Control and Prevention. Information for Health Care Providers. Available at: <u>www.cdc.gov/cancer/preventinfections/providers.htm</u>. Accessed February 21, 2020; ² NSP IQVIA July '20, ³G-CSF market size based on IQVIA data (MIDAS for ex-U.S. and DDM MD for U.S.; Q3 '16 to Q2 '18. Standardized G-CSF units. 4. Wilson B, Jacob S, Yap ML, et al. Estimates of global chemotherapy demands and corresponding physician workforce requirements for 2018 and 2040: a population-based study. Lancet Oncology 2019; 20(6): 769-780.

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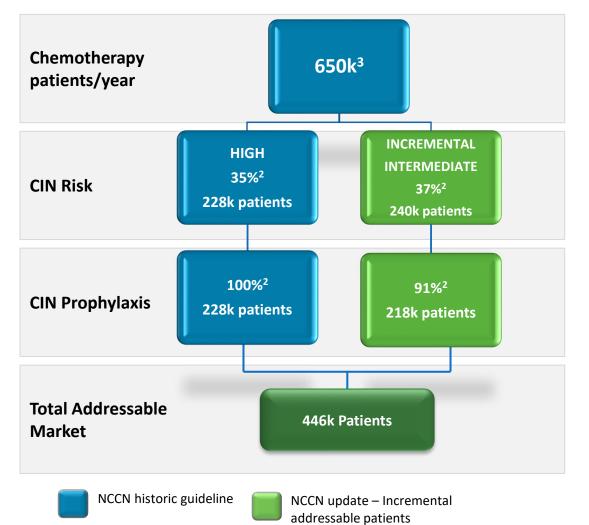
* Growth despite a 20% decline in chemotherapy cycles nationwide from March – June '20 due to the pandemic.

New CIN guidelines double the Addressable US Market

• CIN guidelines modified in early 2020:

- COVID-19 recognized as a universal risk factor
- Prophylaxis now recommended for both high and intermediate risk patients
- The addressable population increased by 100%:
 - 2019: 30% of intermediate risk patients received prophylaxis for CIN¹
 - 2020: 90% dramatic jump in approach to preventing CIN²

CIN Prophylaxis Market dynamics post-guideline update



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1 BYSI qual market research Sept 2019, 2 BYSI qual Market research Aug 2020, 3 https://www.cdc.gov/cancer/preventinfections/providers.html

High unmet medical need even with SOC G-CSF

CIN is a dangerous decrease in a patient's white blood cell count. If Grade 4 neutropenia (ANC < 0.5x109 cells/L) is not treated, patients could die in first cycle of chemotherapy

Short-term Outcome Benefit

G-CSF monotherapy is suboptimal and leaves a significant clinical gap

CIN #1 reason for FN, hospitalization, sepsis, mortality and chemotherapy disruption¹

Long-term Outcome Benefit

Chemotherapy's anti-cancer effectiveness is linear to its dose

Slight Changes in Dosing or Delivery Can Have A Devastating Impact on Survival²

Reduction in Relative Dose Intensity

15%

Reduction in Overall Survival

50%

More than 75% of negative clinical consequences occur in Week 1 after chemo; G-CSF cannot prevent week 1 Plinabulin + G-CSF has the potential to address this important unmet clinical need³



Source: ¹ Lalami Y, Klastersky J.. Crit. Rev Oncol Hematol. 2017; 120:163-179.² Bonadonna G, Valagussa P, Moliterni A et al. Adjuvant cyclophosphamide, methotrexate, and fluorouracil in node-positive breast cancer. N Engl J Med 1995;332:901-906.³ Cancer, Volume: 98, Issue: 11, Pages: 2402-2409, First published: 17 November 2003, DOI: (10.1002/cncr.11827)

Grade 4 neutropenia leads to development of fever and infection; and to chemo dose reduction and less survival

Grade 4 neutropenia was associated with fever

Grade 4 neutropenia leads to dose reduction to (p = 0.04), documented infection (p < 0.0001), <85% of optimum dose \rightarrow lower OS² and bacteremia $(p = 0.002)^1$ Median years (95% CI) Dose reduction ≥15% (n=408) Percent without infection 2.31 (1.87-2.87) Dose reduction <15% (n=411) Percent without feve 100 3.67 (3.15-4.19) Log-rank P value 0.0195 75-75 80 50-50-**BREAST CANCER** 25 25-Overall Survival, 60 0 10 15 20 25 30 35 0 5 10 15 20 25 30 5 **Days Elapsed Days Elapsed** 40 Percent without bacteremia 20 75-**Baseline Absolute** Neutrophil Count 50-0 --- <500 cells/ul → ≥500 cells/µL 25 0 2 5 8 Time 15 20 25 30 35 10 0 5 DR ≥15% 297 209 170 145 89 38 10 0 **Days Elapsed** 323 264 220 175 96 39 6 0 DR <15% 411

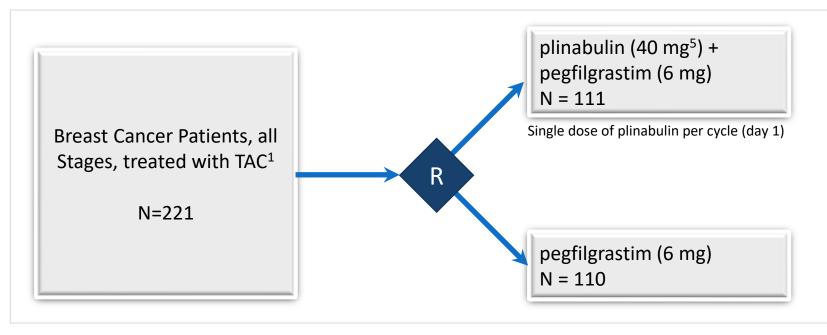
Clinical Breast Cancer, October 2018

Note: ¹Buckley SA et al., "Prediction of adverse events during intensive induction chemotherapy for acute myeloid leukemia or high-grade myelodysplastic syndromes." American J. Hematology 2014; 89(4): 423-28. 2. Denduluri N et al., Clinical Breast Cancer 18(5): 380-386 (2018); Lalami et al. Critical Reviews in Oncology / Hematology 120: 163-179 (2017).



Plinabulin trials designed to maximize broad potential: Plinabulin + G-CSF for all chemo in non-myeloid cancers

Protective-2 Phase 3 Design (4 cycles of chemo treatment)



Double blinded, active controlled, global trial (CRO & central lab: Covance)



¹TAC=Docetaxel, doxorubicin and cyclophosphamide. ²Duration of Severe (Grade 4) Neutropenia ³Absolute Neutrophil Count ⁴Relative Dose Intensity ⁵Fixed dose, equivalent to 20 mg/m²

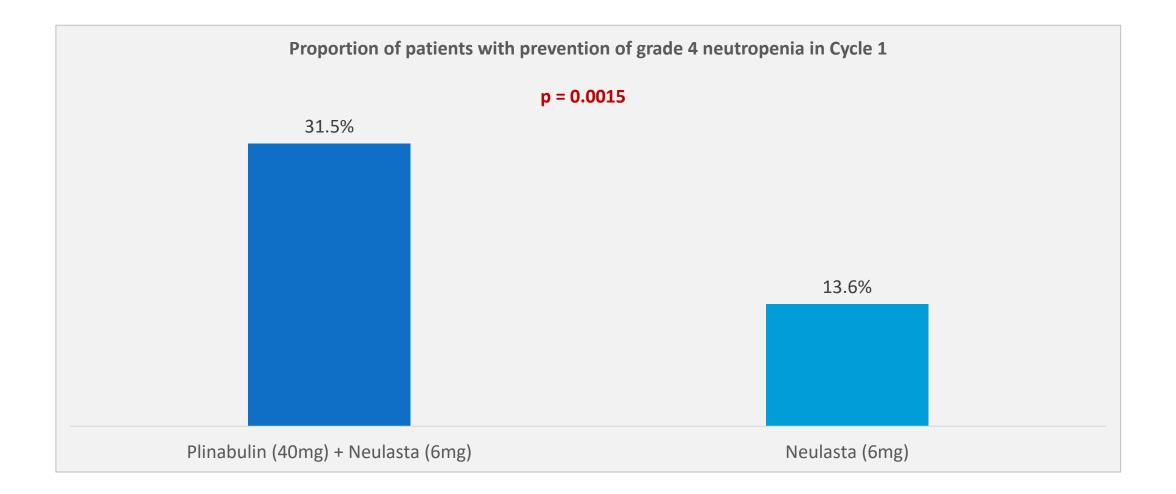
Primary Endpoint:

 % prevent Grade 4 neutropenia (Cycle 1)

Secondary Endpoints:

- Mean DSN² (Cycle 1, Day 1-8)
- Mean ANC³ nadir (Cycle 1)
- % of prevention of grade 3 and 4 neutropenia (Cycle 1)
- DSN (Cycle 1)
- % of bone pain (Cycle 1)
- Composite risk
- % of RDI⁴ < 85%

Plinabulin – G-CSF combination demonstrated over 100% better prevention of grade 4 neutropenia in Cycle 1



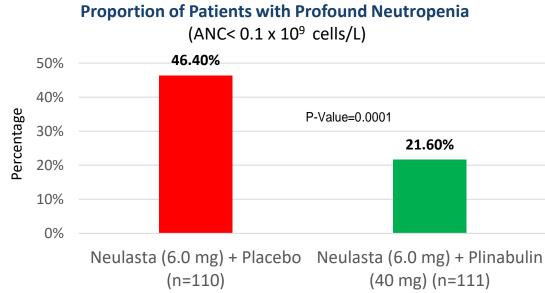


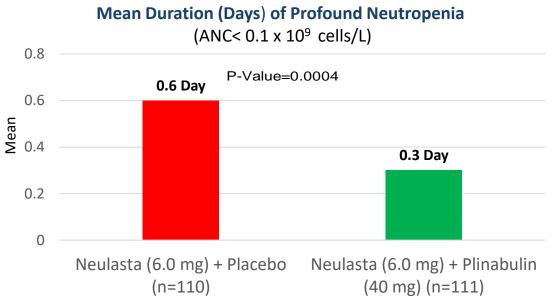
PROTECTIVE-2 Phase 3 data: positive topline results with statistical significance favoring the combination

Key Efficacy Endpoints	Results (combo n=111, pegfilgrastim n=110)		
Primary endpoint:			
Proportion of patients with prevention of grade 4 neutropenia in Cycle 1	 31.5% vs. 13.6%, p=0.0015 >100% better prevention rate in combination of plinabulin + G-CSF 	Better safety profile in the combination vs. SoC	
Key secondary endpoints (based on ANC):		 >20% less grade 4 AEs in the combination (58.6%), 	
Mean DSN in Cycle 1, Day 1-8	 p = 0.0065 Plinabulin's MoA of early onset in Week 1 	compared to pegfilgrastim alone (80.0%)	
Mean DSN in Cycle 1 (severe neutropenia: ANC < 0.5 x 10 ⁹ cells/L)	 p = 0.0324 Combination is better in CIN benefit vs. G-CSF in cycle 1 		
Mean ANC Nadir (x 10 ⁹ cells/L)	 0.538 vs. 0.308, p = 0.0002 The combination helps to lift patients away from grade 4 danger zone 	Profound Neutropenia leads to 80% death in first week of infection ¹ , 48% FN and 50% Infection ² .	
Mean Duration of Profound Neutropenia in cycle 1 (Profound Neutropenia: ANC < 0.1 x 10 ⁹ cells/L)	 p = 0.0004 Combo better than G-CSF alone in CIN benefit 		

BeyondSpring 1. Bodey et al. Ann Intern Med 64(2): 328 (1966); 2. Bodey et al. Cancer 41(4): 1610 (1978)

Protective-2 (Phase 3): Superior prevention of Profound Neutropenia with **Combination vs G-CSF alone**





The combination reduces the incidence of Profound Neutropenia by >50% Compared to G-CSF Alone, which correlates to >40% FN risk reduction in the combo vs. G-CSF.



Potential for a broad Prevention of CIN label: Plinabulin + G-CSF for all chemo in non-myeloid solid tumors

The Premise

- Chemo kills fast dividing cells, which includes cancer cells, and white blood cells in bone marrow
- CIN is a problem with bone marrow, and not a problem with a specific type of cancer
- All bio-similar filgrastim or pegfilgrastim approval was based on TAC and breast cancer, which is an example of high-risk chemotherapy.

Broad Label Potential for Plinabulin – G-CSF Combination

- G-CSF class provides a base protection from severe neutropenia in week 2 following chemotherapy use.
- Plinabulin has MoA of protecting neutrophil in week 1 after chemotherapy, which has been consistently shown in five (5) trials that includes various chemotherapies, and in various non-myeloid cancer trials.
- Thus, the combination provides increased protection for the complete cycle 1 following chemotherapy.

Proposed Label

Plinabulin when combined with G-CSF is indicated for concurrent administration with a myelosuppressive chemotherapeutic regimen in patients with non-myeloid solid tumor for the prevention of chemotherapeutic induced neutropenia (CIN).



Plinabulin+ G-CSF for CIN, NDA Submission in Q1 2021: Superior profile in a broad label: all chemo, non-myeloid solid tumor

Plinabulin shown to statistically reduce Grade 4 neutropenia in 6 clinical trials (1,200+ patients)

Registration Study

Supporting Study

Plinabulin vs. placebo

 Grade 4 reduction highly statistically significant (Study 101 and DUBLIN-3, p<0.0003 and p<0.0001 respectively) Plinabulin + G-CSF combo vs. G-CSF mono (Protective-2)

- Superior response in primary and key secondary endpoints with statistical significance
- Effect size correlate with clinical meaningful endpoints

MOA support from 5 studies: Plinabulin early onset in Week 1, G-CSF effect in Week 2

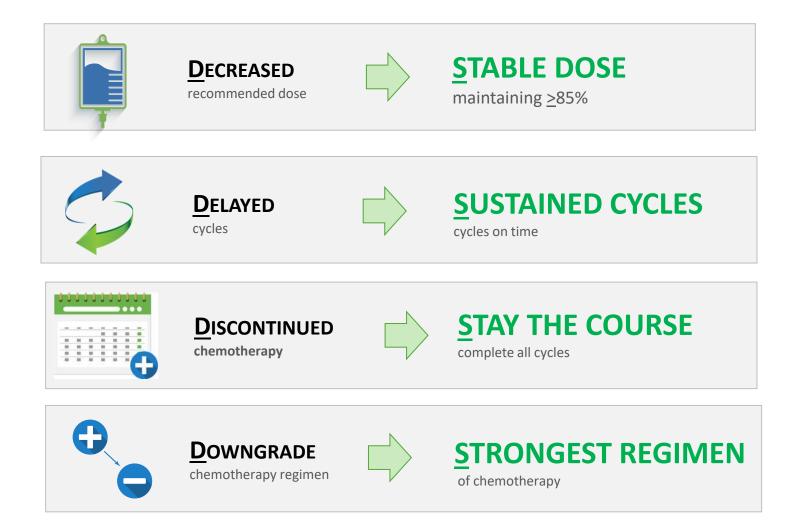
Supporting Study

Plinabulin vs. G-CSF (Protective-1)

- Non-inferior CIN activity
- Superior adverse event profile: limited bone pain, limited platelet reduction, and limited immune suppression¹

700+ cancer patients treated with Plinabulin (various doses)

Chemotherapy without compromise: Turning the 4 Ds into the 4 Ss

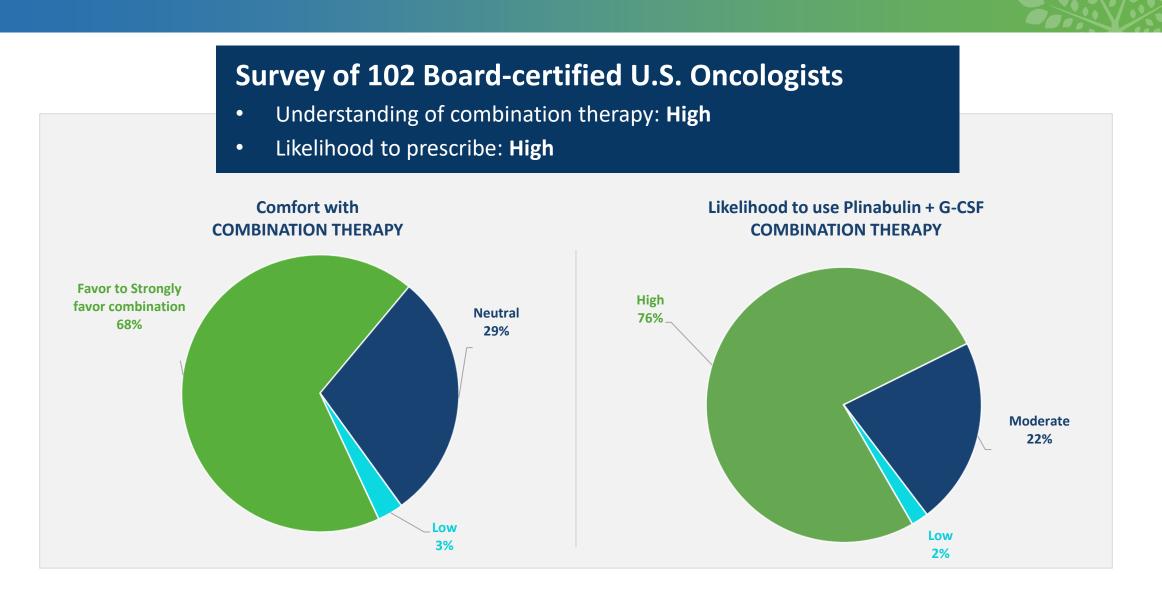


Plinabulin + G-CSF

- Differentiated clinical profile, potential to improve SOC
- Greater clinical control
- Improved outcomes

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Oncologists understand Plinabulin's potential to raise the SoC in CIN





Market research conducted Sep '20; n = 102 board certified U.S. oncologists; data on file BeyondSpring C01 Based on the product information you have just seen, how likely are you to prescribe the Product N + G-CSF combo for CIN in non-myeloid malignancies? C02 How would you rate your comfort with using a combination treatment (e.g. Product N + G-CSF combo) for CIN in non-myeloid malignancies

Plinabulin + G-CSF: "Breakthrough Therapy" with potential to set a new SOC for CIN

Opportunity

- Market size
- ✓ Market growth
- ✓ NCCN guideline change
- Managed care coverage

Unmet need

- Grade 4 neutropenia complications
- CIN: #1 reason for therapy change (4Ds)
- Monotherapy G-CSF not effective
- ✓ 4Ds result in reduced OS

Product differentiation

Plinablulin + G-CSF addresses 3 oncologist needs:

- ✓ Maintains chemo regimen
- Keeps ANC out of the danger zone and thus less FN and less hospitalization
- ✓ Improved A/E profile

Plinabulin+ G-CSF has the potential to:

- Address the oncologist's desire for increased control
- Reduce patient anxiety and fears associated with interrupted therapy and adverse events
- Deliver improved chemotherapy care





Plinabulin in NSCLCFinal phase 3 topline anti-cancer datain 1H 2021

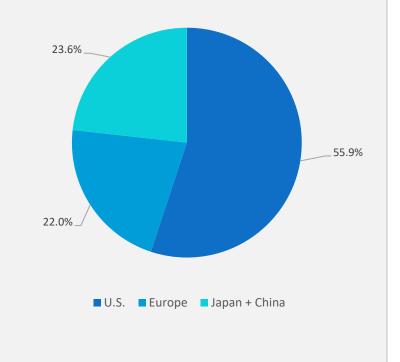


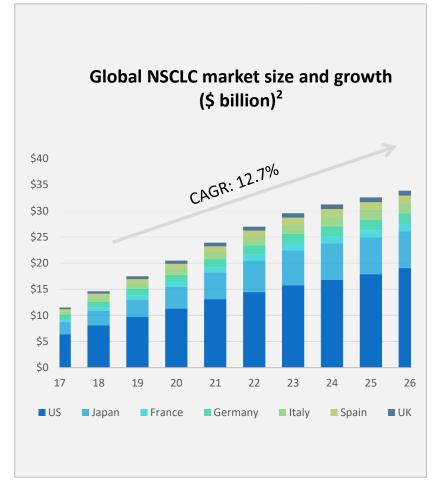
Increasing incidence of non-small cell lung cancer (NSCLC)

Key Investment Highlights:

- ~1.5M NSCLC diagnoses globally
- Key NSCLC drug sales (U.S., Japan, and major EU markets):
 - \$11.5B in 2017
 - Increasing to \$33.9B by 2026
- Primary drivers of growth:
 - Increasing incidence of NSCLC
 - Premium-priced checkpoint inhibitor usage, particularly in the 1st line setting









2nd/3rd line NSCLC patients

	EGFR mutant	EGFR wild type	In EGFR wild type patients		
% of 2L/3L NSCLC patients (western)	15%	85%	Much larger population		
mOS SoC	18.3 months (TKI)	6-8 months (docetaxel)	Much shorter OS		
mOS TKI vs docetaxel ¹		5.4 vs 8.2 months	TKI worse than docetaxel		
Currently available therapies		PD-1 Pemetrexed Ramucirumab + docetaxel Docetaxel	All with significant limitations		
Severe unmet clinical need					

For lung cancer patients infected by COVID-19, death rate is 55%²

Approved therapies fail to address EGFR wild type NSCLC (85% of Western patients) in 2nd and 3rd line treatment

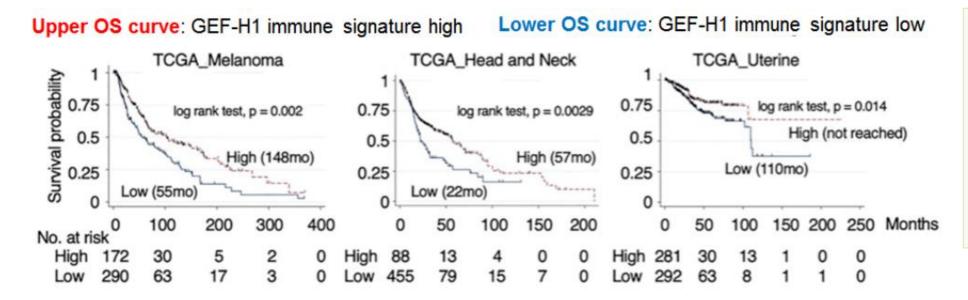
Only four therapies currently approved: Severe Unmet Medical Needs

	Moving i	nto 1st line	2nd and 3rd lines		
	Nivolumab (PD-1) vs. docetaxel ¹	Pemetrexed vs. docetaxel ²	Ramucirumab + docetaxel vs. docetaxel ³	Plinabulin + docetaxel vs. Docetaxel ⁴	
mOS	+2.8 months (12.2 vs. 9.4) HR = 0.73	+0.4 months (8.3 vs. 7.9) HR = 0.99	+1.4 months (10.5 vs. 9.1) HR = 0.86	+4.6 months (11.3 vs. 6.7) HR < 0.75 ⁵	
ORR	19% vs.12%	9.1% vs. 8.8%	23% vs. 14%	18.4% vs.10.5%	
Grade 3/4 neutropenia	0% vs. 27%	5% vs. 40%	49% vs. 39%	7% vs. 26%	
DOR	17 vs. 6 months	4.6 vs. 5.3 months		12.7 vs. 1 months	
Conclusion	 Introduces potential cytokine storm leading to inflammation Moved into 1st line 	 No efficacy improvement Approved based on low neutropenia rate 	 Modest efficacy benefit Higher severe neutropenia rate than docetaxel 	 Superior efficacy Superior CIN benefit Durable anti-cancer benefit Safety benefit 	



DUBLIN-3 (Study 103): Phase 3 in NSCLC hypothesis based on Plinabulin IO mechanism on activating GEF-H1, a proven target

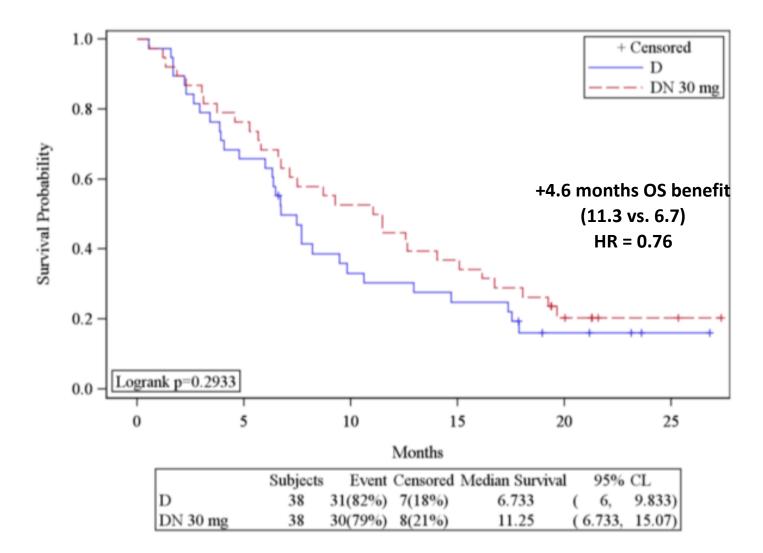
GEF-H1 - higher GEF-H1 immune signatures associated with longer OS in cancer patient ¹



- Anti-cancer activity
- Augmentation of IO effect in combo with chemo or radiation, as antigen generator
- GEF-H1 mechanism driven OS benefit



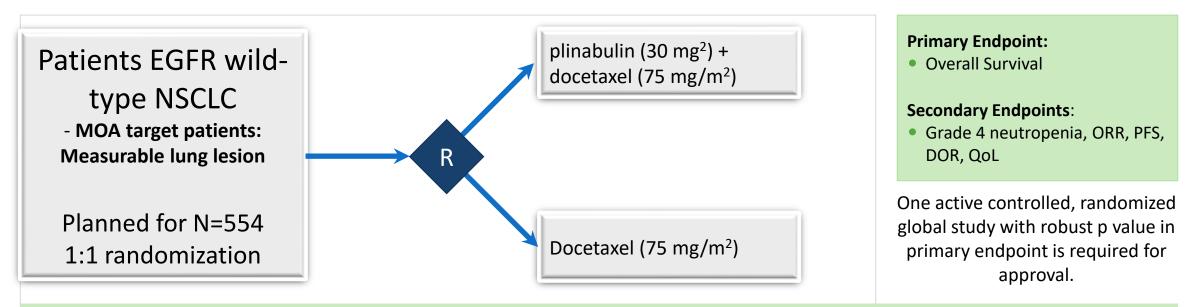
Post-hoc Phase 2 data from Plinabulin in NSCLC in mechanism targeted patients shows overall survival benefit



- Plinabulin MoA- targeted patients: Measurable lung lesion with RECIST 1.1 (CT scan > 1 cm in lung); 70% of NSCLC
- Improved QoL and well tolerated safety profile

DUBLIN-3 (Study 103): Phase 3 in NSCLC – second interim analysis completed; DSMB recommended trial to continue

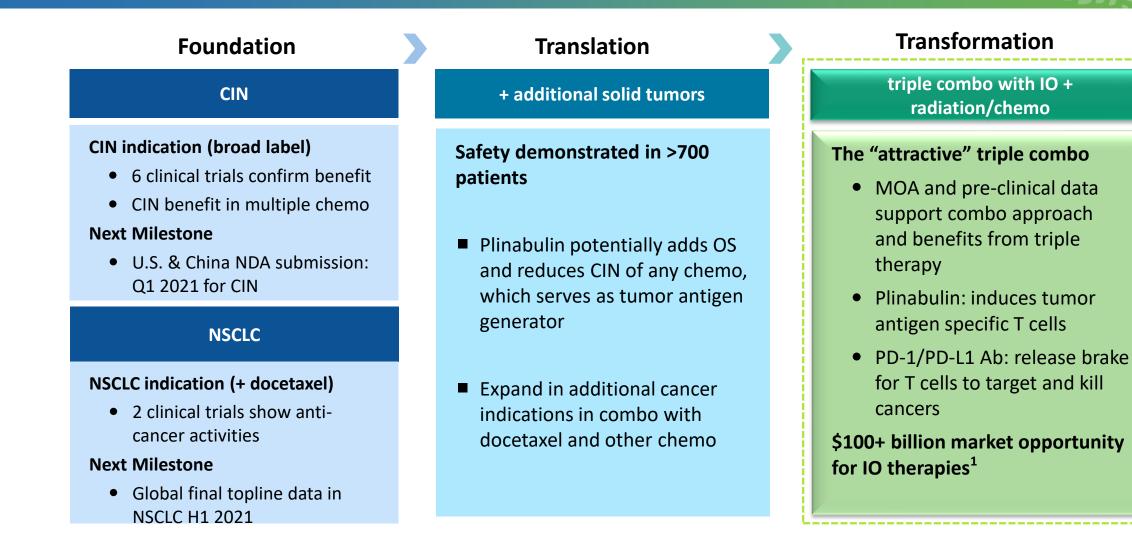
DUBLIN-3 Phase 3 in NSCLC Trial Design



- First interim analysis completed in Q1 2019 at 1/3 patient mortality. DSMB recommended trial to continue without modification (HR < 0.75 based on mOS)
- Second interim analysis completed in Q2 2020 at 2/3 patient mortality. DSMB recommended trial to continue without modification
- Final analysis: 439 patient mortality; study succeeds if p < 0.046 for mOS 1H 2021</p>

BeyondSpring

Plinabulin: "pipeline in a drug" for multiple cancer indications





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Near-term milestones will create significant value for Plinabulin in the next 12 to 36 months

- CIN in global markets -> compelling value creation potential from improving SOC, and large opportunity for life cycle management
- NSCLC in global markets -> large and growing patient population with large unmet medical need
- IO Combos -> multiple cancers and improving current therapies

Plinabulin life-cycle management related to CIN **IO Combos** Various cancers Clin.Data: Q4 2021 NSCLC Final topline global Ph 3 data: H1 2021 CIN U.S. + China NDA submission Q1 2021







SEED is a majority-owned subsidiary pursuing "Molecular Glue" targeted protein degradation to degrade diseasecausing proteins previously believed to be undruggable

- \$800M collaboration with Lilly on three targets
- Own targets (e.g., KRAS)
- Structure conducive to having additional collaborations

BeyondSpring: Key Highlights



Mission

Committed to raising the standard of care for cancer patients in the largest global markets with first-in-class treatments that improve lives and clinical outcomes for millions of patients in need

2 near-NDA Assets Global Market Opportunities

PLINABULIN: Raising SOC in CIN & NSCLC

- First-in-Class immune agent
- New Chemical Entity
- ✓ IP through 2036 in 36 jurisdictions

CIN: Combo with G-CSF

- ✓ Final Ph 3 topline data Nov 2020
- ✓ NDA submission early 1Q 2021
- Market: \$4.5B (US)
- Breakthrough Designation (US, China)

NSCLC: Combo with docetaxel

- Final Ph 3 data 1H2021
- Early 2022 NDA submission
- ✓ \$30B+ global market

Broad Pipeline

PLINABULIN: A pipeline in a drug

- Triple combo w/IO agents and radiation/chemo
- Expansion to additional solid tumors

Targeted Protein Degradation Platform

- Seed Therapeutics (Subsidiary)
- Collaboration with Eli Lilly

Three Pre-Clinical IO Agents

Global Capabilities Continuous Innovation

Strong clinical development

- Enrolled 1,000+ patients to final filing stage for CIN and NSCLC
- Dual U.S. and China development strategy
- ✓ Strong clinical investigator network

Deep Regulatory Expertise

Commercialization Planning Underway



Appendix



Plinabulin – first-in-class agent (a 20-year journey)

2000-2005: NCE discovered 20 years ago from sea microbes, then optimized

- In 2000, new chemical entity (NCE) Halamide class compound was discovered from sea microbes by Nereus Pharmaceuticals based in San Diego, CA
- 300+ derivatives were made and screened through colon cancer cell lines, and Plinabulin was found to be most efficacious and safe compound to enter clinics

2005-2010: Early clinical studies showed unique clinical profile for Plinabulin, a tubulin binder; but did not know why

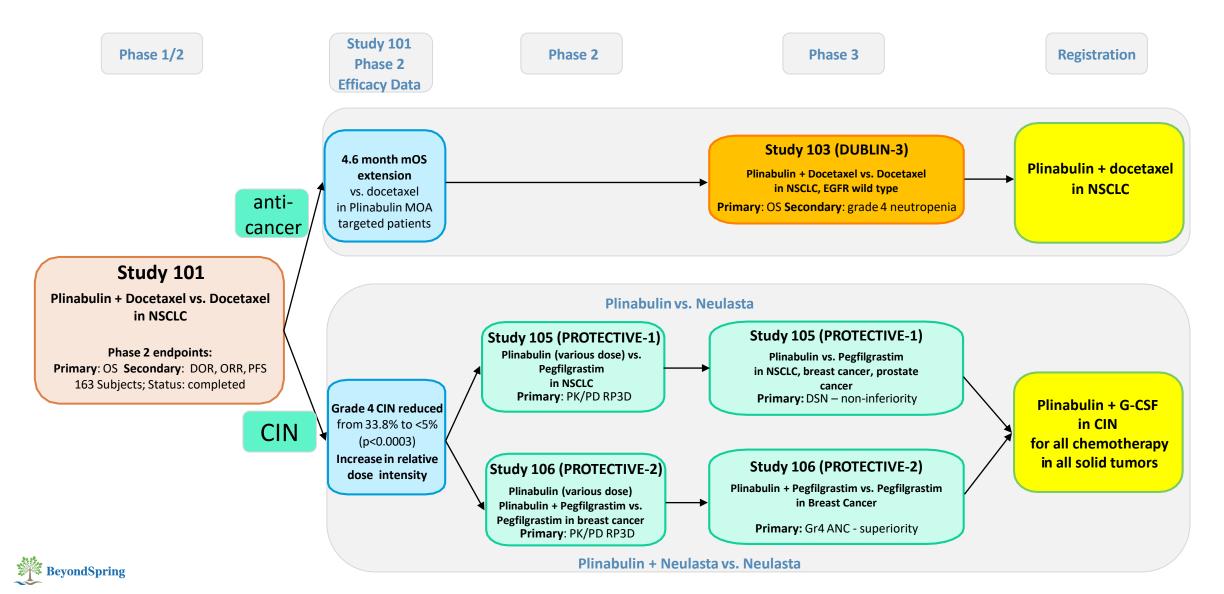
- Early clinical studies in NSCLC in Plinabulin + docetaxel showed durable anti-cancer benefit, and serendipitous finding of CIN benefit
- Mechanism as a tubulin binder cannot explain clinical profile

2010-2020: Research collaboration with experts in protein structure, immune and CIN field shed light on Plinabulin's unique MOA, leading to targeted clinical development and extended/strong patent protection to 2036

- Four MOA papers published in 2019, collaborating with leading scientists from Univ. of Basel, Fred Hutch, Mass General and MD Anderson
- Data from additional manufacturing and research provide strong basis for extended and strong patent protection to 2036 in 36 jurisdictions
- Design registration studies in CIN and NSCLC based on Plinabulin MOA and chart targeted clinical development programs.



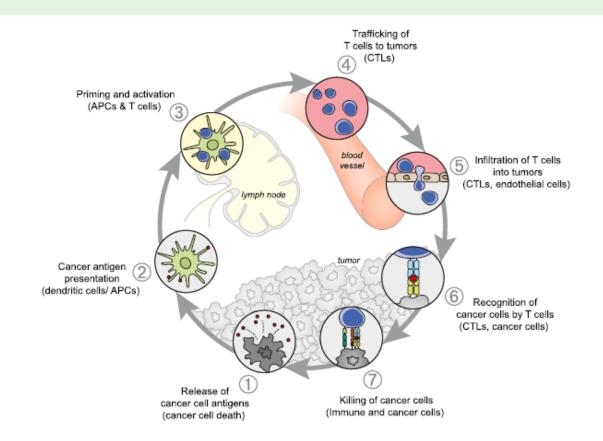
Plinabulin Registration Studies



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Plinabulin induces dendritic cell DC maturation, a key step in initiating anti-cancer immunotherapy

- The generation of cancer immunity is a cyclic process. In principle, the response of T cells to tumor cells should be amplified and expanded
- Initiating anti-cancer immunity includes antigen release, presentation and activation of cancer antigen-specific T cells.
 Dendritic cells are the most important antigen-presenting cells





Global neutropenia market

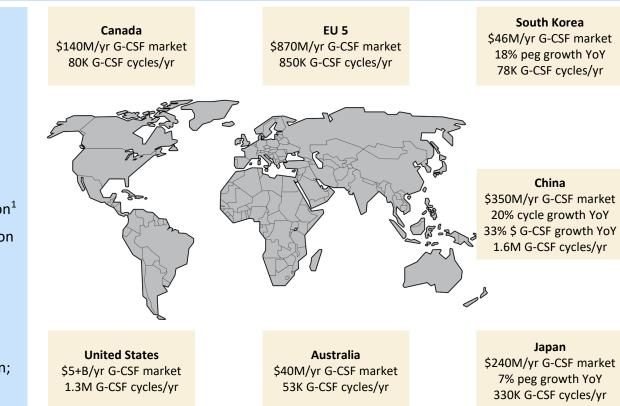
- CIN market = \$7 billion in 2017 (mainly used with high risk chemo)
- China and the U.S. represent 2/3 of all G-CSF global therapy
- Plinabulin positioning: Plinabulin as mono <u>or</u> combo therapy improves on the standard of care in the treatment of Chemotherapy Induced Neutropenia; potential for improved chemotherapy outcome

Addressable market

- Combination with G-CSF to
 - Improved neutropenia
 - Reduced bone pain
 - Improve compliance & persistency with chemo

Global

- G-CSF cycles/year = 4.3 million¹
- G-CSF market value = \$7 billion
- Plinabulin + G-CSF: reduced neutropenia and improved bone pain; potential for improved compliance and persistency with chemo
- Plinabulin: reduced bone pain; improved thrombocytopenia and immune function



Note: ¹ https://onlinelibrary.wiley.com/doi/full/10.3322/caac.21338. G-CSF market growth based on IQVIA data and DDM MD data Q3 '16 to Q2 '18. Standardized G-CSF units







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