



Annual Shareholders Meeting

NewLink Genetics Corporation

NASDAQ: NLNK  
May 23, 2018

# Cautionary Note Regarding Forward-Looking Statements

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# NewLink Genetics

A Clinical Development Stage Immuno-Oncology Company

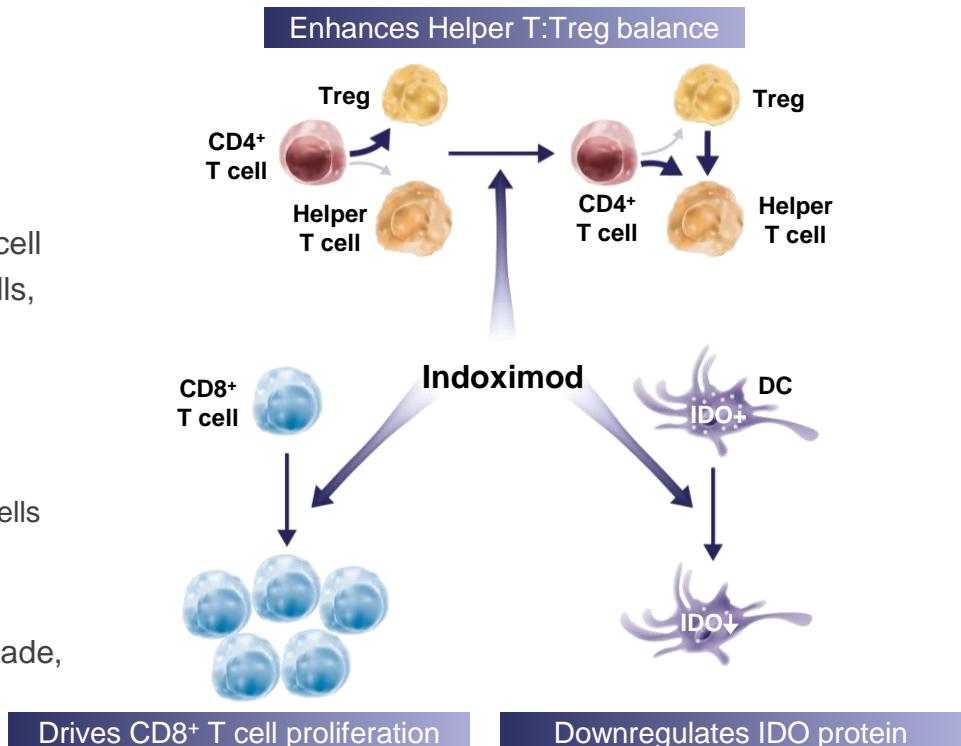
- **Strategic focus on indoximod, an I-O product candidate with a differentiated MOA**
  - Clinical data suggest indoximod combinations may enhance multiple therapeutic modalities
  - Potential to improve patient outcomes across both hematologic cancer and solid tumor indications
- **New data on MOA and emerging clinical data recently presented at AACR**
  - Abstract 3753 – *Indoximod modulates AhR-driven transcription of genes that control immune function*
  - Abstract 10973 – *Front-line therapy of DIPG using the IDO pathway inhibitor indoximod in combination with radiation and chemotherapy*
- **Clinical and operational developments**
  - Finalized novel formulation of indoximod
  - Appointed two new members to the Board of Directors
  - Undertaking clinical program review

Multifaceted approach to cancer driven immunosuppression

# Indoximod Mechanism of Action

## A Unique Approach to Reversing Immunosuppression

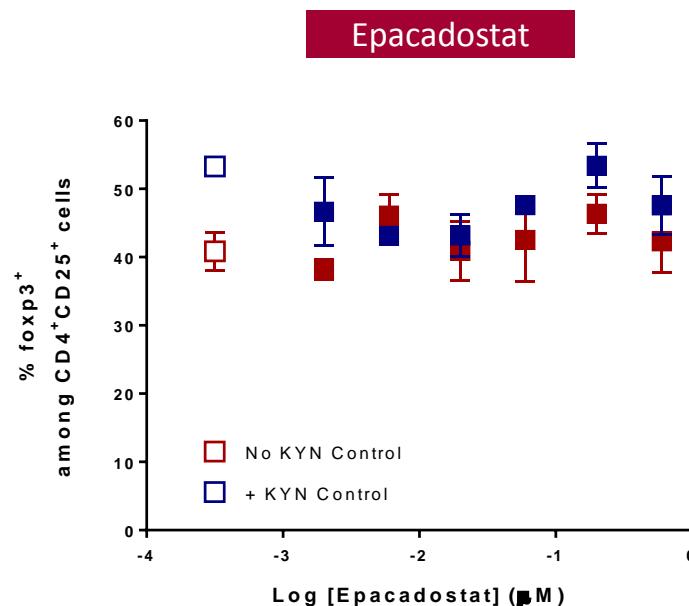
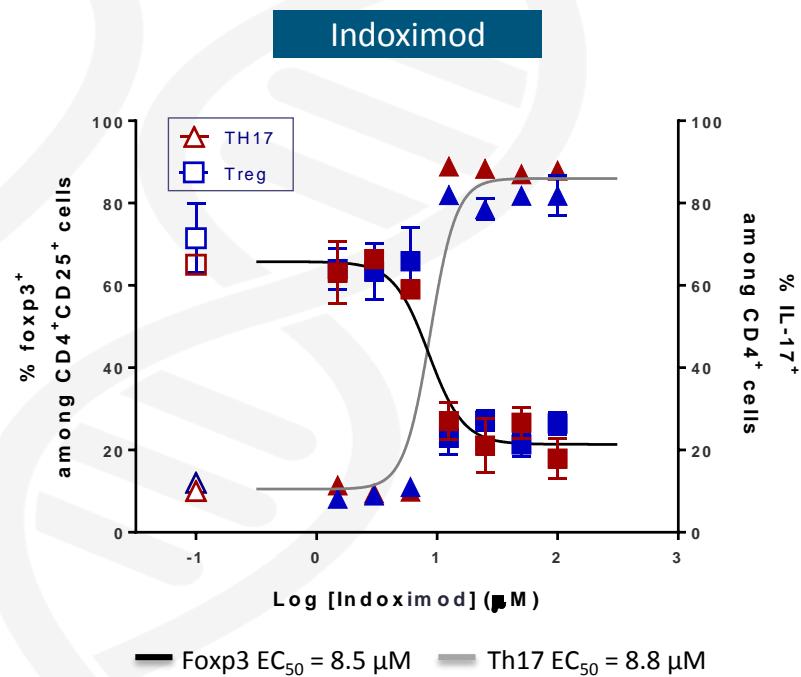
- Orally administered, small-molecule that reverses the immunosuppressive effects of low tryptophan and high kynureanine that result from IDO activity
- Immuno-stimulatory effects of indoximod impact 4 main cell types: CD8<sup>+</sup> T cells, CD4<sup>+</sup> T helper cells, T regulatory cells, and dendritic cells
  - Reverses the effects of low tryptophan by increasing proliferation of effector T cells
  - Drives differentiation into T helper cells vs regulatory T cells
  - Downregulates IDO expression in dendritic cells
- Potential synergy has been shown with checkpoint blockade, chemotherapy, radiation and vaccines



IDO, indoleamine 2,3-dioxygenase; Treg, T regulatory cell; DC, dendritic cell.  
Brincks EL, et al. AACR 2018. Abstract 3753.

# Indoximod vs Epacadostat: A Different Mechanism of Action

## *Indoximod Drives Differentiation of Helper vs Regulatory T Cells*



Indoximod reduces Tregs and increases effector T cells

# Indoximod plus Chemotherapy in Acute Myeloid Leukemia (AML)

## *Phase 1/2 Exploring Minimal Residual Disease as a Surrogate Endpoint*

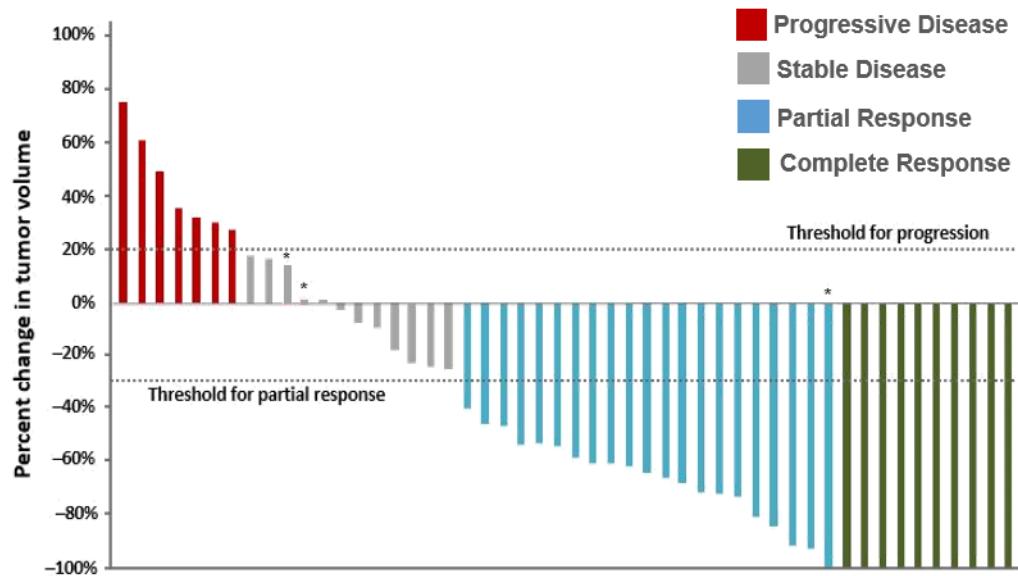
- Phase 1/2 trial for patients with newly diagnosed AML
  - Combination with current standard of care (7+3 chemotherapy)
  - Currently enrolling Phase 1b expansion cohort
  - Minimal residual disease evaluated by sensitive flow cytometry assay
- Data presented by Emadi, et al at EHA, June 2017 (abstract E-012)
  - Indoximod does not appear to add significant toxicity
  - 7/9 patients who completed treatment per protocol achieved morphologic complete response (CR)
  - 7/7 patients who achieved a CR had no evidence of minimal residual disease

Updated AML data anticipated in 2H 2018

# Indoximod (IDO) plus Pembrolizumab (PD-1) in Advanced Melanoma

## Phase 2: Impressive Response Rate and Progression Free Survival (N=51)

Response	N (%)
ORR	<b>31 (61)</b>
CR	10 (20)
PR	21 (41)
SD	10 (20)
DCR	<b>41 (80)</b>
PD	10 (20)
Survival	
Median PFS	<b>12.9 months</b>
PFS at 12 months	<b>56%</b>



Updated melanoma data including subset analysis will be presented at ASCO

\*Patients that progressed due to new non-target lesions.

Note: 1 patient was unevaluable for response due to pleural effusion/collapsed left lung; the patient progressed based on several new non-target lesions.

Zakharia Y, et al. Oral presentation at: Third International Cancer Immunology Conference; September 6-9, 2017; Frankfurt, Germany.

# Early Results for Children with Diffuse Intrinsic Pontine Glioma (DIPG)

## *Encouraging Early Data from Pilot Cohort of 6 Patients*

- DIPG is rapidly fatal with significant unmet need
  - Standard of care treatment is palliative radiation (usually 54 Gy)
  - Median time to progression after radiation is ~6 months<sup>1</sup>
  - No surgical options for patients with DIPG
- Encouraging early results from pilot cohort
  - Phase 1 data suggest indoximod-based immunotherapy enhances conventional therapy (radiation, chemotherapy)<sup>2</sup>
  - All 6 DIPG patients have finished upfront radiation combined with indoximod
  - All 6 patients showed initial improvement in symptoms<sup>3</sup>

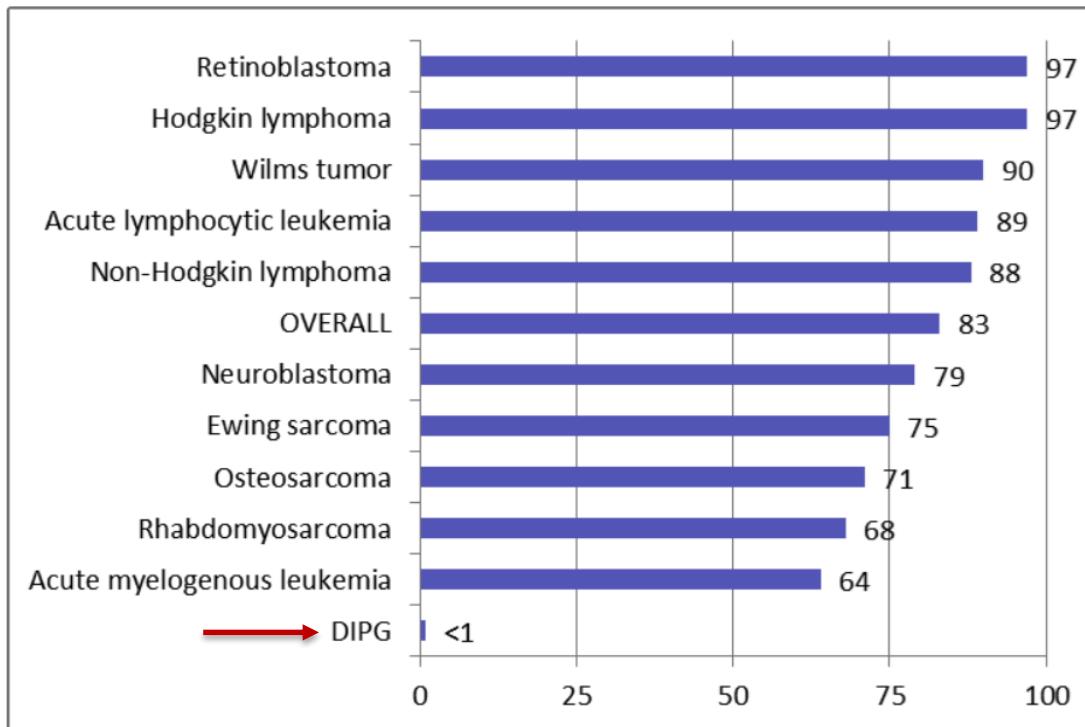
1. Wolff JE, et al. *J Neurooncol.* 2012;106(2):391-397. 2. Cohen KJ, et al. *Neuro Oncol.* 2011;13(4):410-416.

2 Johnson T, et al. SNO 2017.

3 Johnson T, et al. AACR 2018. Plenary #10973.

# DIPG Prognosis Worst of All Pediatric Cancers

## Pediatric Cancer 5-Year Survival Rates



# Encouraging Early Data in DIPG

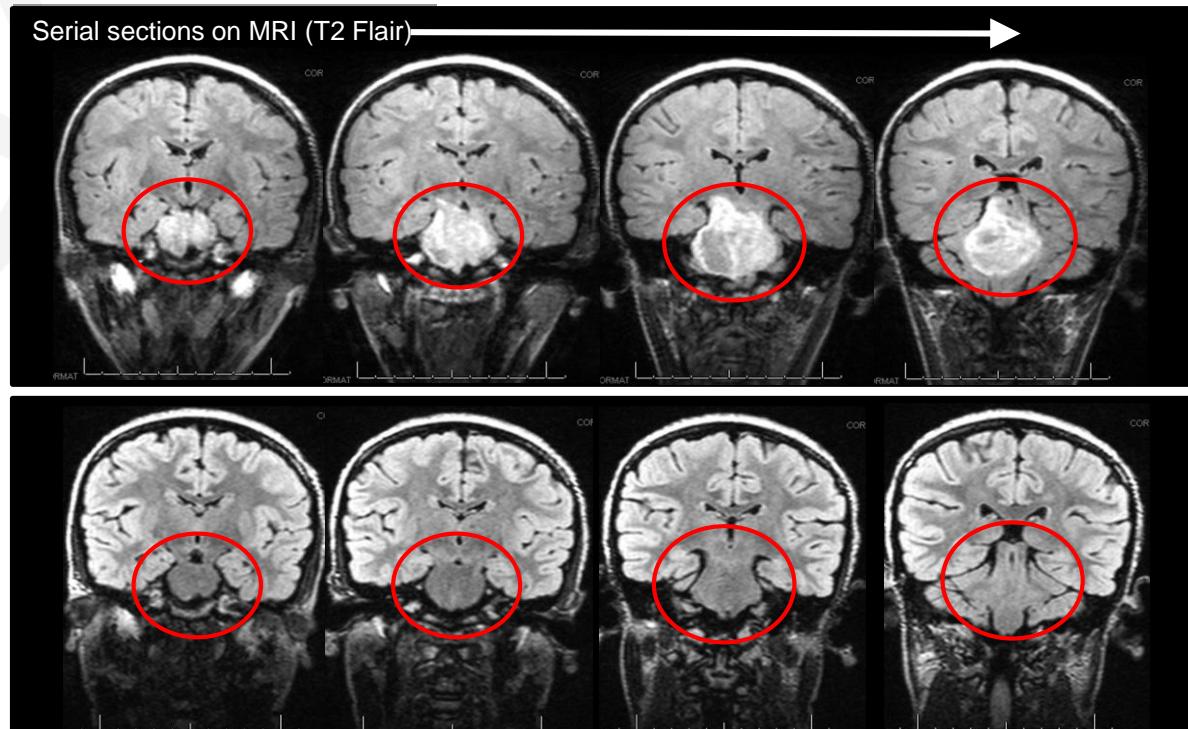
*Response for 9.4-Year-Old Male with Newly Diagnosed DIPG*

Baseline  
(pretreatment)

DIPG scans reviewed by  
Tina Young-Poussaint, M.D.,  
Boston Children's Hospital

Patient 037 classified as:  
“Significant response”

After 6 weeks of  
indoximod +  
radiation (54 Gy)



Patient remains neurologically normal at 6 months from initial treatment

## Upcoming Milestones 2018

- Abstracts accepted for presentation at ASCO Annual Meeting, June 2018
  - Abstract 9512 – *Phase 2 trial of the IDO pathway inhibitor indoximod plus checkpoint inhibition for the treatment of patients with advanced melanoma*
  - Abstract 4015 – *Phase 2 trial of the IDO pathway inhibitor indoximod plus gemcitabine / nab-paclitaxel for the treatment of patients with metastatic pancreas cancer*
- Abstract accepted for presentation at the 18<sup>th</sup> International Symposium on Pediatric Neuro-Oncology (ISPNO), July 2018
  - *Radio-immunotherapy using the IDO pathway inhibitor indoximod for children with newly-diagnosed DIPG*
- Data from Phase 1b trial of indoximod plus standard-of-care chemotherapy for patients with newly diagnosed acute myeloid leukemia (AML) intended to be presented 2H 2018

Data for indoximod in different therapeutic combinations across multiple cancer indications

## Financial Position

Q1 2018 End Cash and Equivalents	\$143.9 million
YE 2018 Cash Projected	To be updated on Q2 call
Shares Outstanding as of March 31, 2018	37.2 Million

Resources sufficient to support focused clinical development of indoximod

# NewLink Genetics – Key Takeaways

## *Indoximod, an Immuno-Oncology Candidate with Differentiated MOA*

- Indoximod has a differentiated mechanism of action (MOA)
  - Reverses the effects of low tryptophan by increasing proliferation of effector T cells
  - Drives differentiation into T helper cells vs regulatory T cells
  - Downregulates IDO expression in dendritic cells
- Promising clinical activity of indoximod in combination with
  - Chemotherapy in AML
  - Checkpoint blockade in melanoma
  - Radiation and chemotherapy in DIPG
- Additional indoximod data to be presented at upcoming medical conferences
  - Melanoma & Pancreatic Cancer: Final Phase 2 results at ASCO, June 2018
  - DIPG: Updated Phase 1b data at ISPNO, July 2018
  - AML: Updated Phase 1b data intended 2H 2018

Differentiated MOA demonstrating clinical activity for multiple combinations and indications