

TarGeting B-Cell Diseases

## Forward Looking Safe Harbor Statement

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## **Our Goal**

# To develop the best possible treatment for B-cell diseases

Ideally, to cure these diseases



## What are B-cell Diseases?

B-cell diseases refer to conditions that are associated with aberrant B-cells or B-cell functions, including:

## Chronic Lymphocytic Leukemia (CLL)

- Non-Hodgkin's Lymphoma
  - Follicular Lymphoma (FL)
  - Marginal Zone Lymphoma (MZL)
  - Diffuse Large B-cell Lymphoma (DLBCL)

#### Autoimmune Diseases

 Multiple Sclerosis (MS), Rheumatoid Arthritis (RA) and Lupus (SLE)



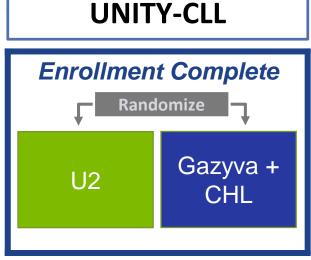
Umbralisib (TGR-1202)	Ublituximab (TG-1101)
Next Generation PI3K delta inhibitor	Next Generation anti-CD20 monoclonal antibody
Overcomes 1 <sup>st</sup> generation Toxicity	Glycoengineered for enhanced potency over 1st generation
Activity across NHL and CLL	Activity in Rituxan refractory patients
Once daily dosing vs. BID	Shorter infusions than all other anti-CD20s (1.5 v 3-4 hours)





## For the Treatment of CLL

- There are ~115,000 Americans living with CLL and ~20,000 newly diagnosed each year
- expected to be the only novel doublet approved for BOTH newly-diagnosed and relapsed patients
- Possible accelerated approval based on ORR; Full approval based on PFS



Total Enrollment	420
Target ORR Improvement	15%
Complete Enrollment	4Q17
Top-Line ORR	2Q18
Target NDA/BLA Filing	4Q18

# Company's Assumptions UNITY-CLL: ORR Endpoint

Targeting ~15% improvement in ORR (with Minimum detectable difference of ~13%)

	Gazyva+CHL	U2
Treatment Naïve	75-78%	88-92%
Relapsed/ Refractory	55-60%	78-82%
Blended ORR	67-71%	84-88%

# **UNITY-CLL PFS Endpoint**

Integrated Analysis: Umbralisib & U2 (n=27)	Helios: Benda + Rituxan (n=183)	Umbralisib + Gazyva + CHL (n=15)	CLL-11: Gazvya + CHL (n=289)	
Rel/Ref		Front Line		
24+ months*	13.3 months	~36+months**	26.7months	

<sup>\*</sup> Median PFS for Umbralisib Monotherapy: 24 Months; Median PFS and DOR not reached for Umbralisib + Ublituximab ('U2')



<sup>\*\*</sup> Median PFS not reached with longest patient on 43+ months

# **GENUINE Update**

MURANO Ph. 3 GENUINE Ph. 3\*

	Venetoclax + Rituxan	Ublituximab + Ibrutinib
Median Number of Prior Lines	1	3
ORR (IRC)	92.3%	81%
CR (IRC)	8.3%	10%

\*GENUINE Data Update as of August 2017

#### **Expedited Programs for Serious Conditions – Drugs and Biologics**

Demonstrating meaningful benefit over available therapy

Provides efficacy comparable to those of available therapy, while (1) avoiding serious toxicity that occurs with available therapy...



# **Umbralisib and U2 in NHL**



**GLYCOENGINEERED UBLITUXIMAB + PI3K DELTA UMBRALISIB (TGR-1202)** 

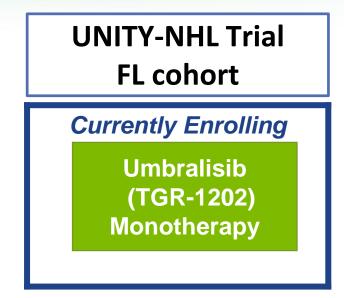
## PREVIOUSLY TREATED NHL PATIENTS

Includes three cohorts: FL, MZL and DLBCL



# Umbralisib and U2 in Follicular Lymphoma (FL)

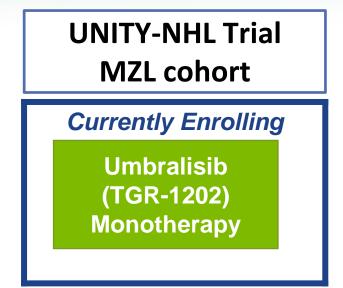
- Approximately 15,000 new cases per year with ~7,500 relapsed patients needing treatment per year
- 53% ORR for umbralisib single agent at higher doses in r/r FL in Phase 1 at higher doses
- Defined path for accelerated approval based Copanlisib approval



Target Enrollment	~100
Target ORR	45-55%
Complete Enrollment	Mid-18
Top-Line Data	1H19

# Umbralisib and U2 in Marginal Zone Lymphoma (MZL)

- Approximately 7,500 new cases per year, with ~3,000 relapsed patients needing treatment each year
- Ibrutinib recently approved with 46% ORR
- Defined path for accelerated approval based on recent ibrutinib approval



Target Enrollment	~60
Target ORR	40-50%
Complete Enrollment	4Q18
Top-Line Data	1H19

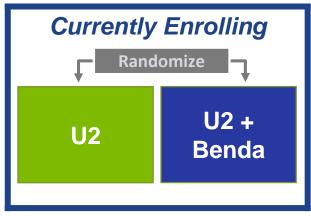


# Umbralisib and U2 in Diffuse Large B-cell Lymphoma (DLBCL)

 US annual incidence of ~20,000 new cases per year of which ~50-60% cured with front-line treatment

 For those not cured, ~20% will be eligible for, and obtain a cure from, transplant

 Nothing approved for the ~6,000 relapsed or refractory patients who are not eligible for transplant UNITY-NHL Trial DLBCL Cohort



Target Enrollment	~200
Target ORR	40-50%
Complete Enrollment U2 & U2+Benda	1Q & Mid-18
Top-Line Data	1H19

Possible accelerated approval

# **Umbralisib** and U2

U.S. Market Opportunity (Company Estimates)

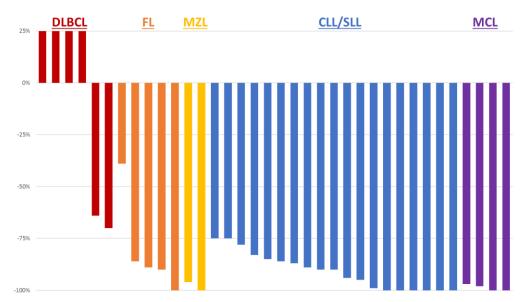
Current Regimen	Disease	Patients Needing Treatment/Year
U2	CLL	~20,000
Umbra	FL	~7,500
Umbra	MZL	~3,000
U2 + Benda	DLBCL	~6,000

CLL & NHL represents a multi-billion dollar opportunity for Umbralisib and U2

# **Newly In-Licensed BTK inhibitor**

#### **Best Percent Change from Baseline in Disease Burden**

Ublituximab + Umbralisib + **Ibrutinib** 



Туре	Pts (n)	CR <sup>†</sup> (n)	<b>PR</b> (n)	ORR n (%)	<b>SD</b> (n)	<b>PD</b> (n)
CLL/SLL	19	6	13	19 (100%)	-	-
MZL	2	1	1	2 (100%)	-	-
MCL	4	2	2	4 (100%)	-	-
FL	5	1	3	4 (80%)	1	-
DLBCL	6	-	1	1 (17%)	-	5
Total	36	10	20	30 (83%)	1	5

# **Ublituximab in Multiple Sclerosis**

- A new study by the Nat'l MS
   Society estimates that ~1,000,000

   Americans are living with MS
- Recently approved anti-CD20 (ocrelizumab) with first year sales approaching \$1B
- Will compete on price and convenience
- Phase 3 Trials under Special Protocol Assessment



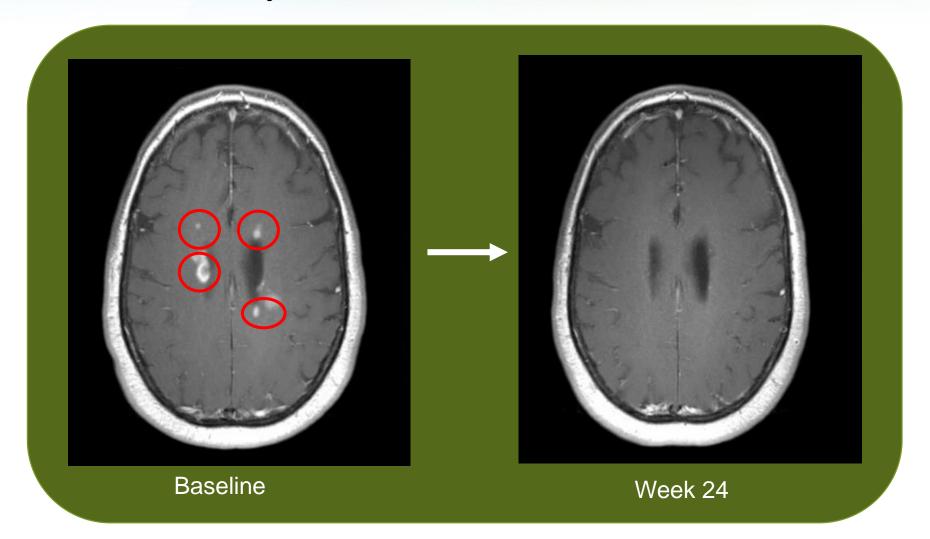
Target Enrollment	~850
Complete Enrollment	1Q 19

# Clinical Endpoints at Week 24

Endpoint	TG-1101 Phase 2 (N=24) (24 Weeks)	Ocrelizumab Phase 2 (N=55) (24 Weeks)	Opera I&II (96 Weeks)
Annualized Relapse Rate	0.09	0.13	0.156
% Relapse Free	95.8%	87%	80%

### T1-Gd MRI at Baseline and Week 24: Study Subject

#### Subject T1-Gd MRI at Baseline and Week 24



# **Corporate & Financial**

#### **Key Financial Statistics**

Ticker: TGTX (NASDAQ)

**Price:** \$9.20 (close on January 10, 2018)

Shares: ~72M (fully-diluted)

Cash: ~\$85M (as of December 31, 2017)

Runway: Into first half 2019

# **Key Goals and Objectives for 2018**

Q1

- Present Updated MS Phase 2 Data
- Complete Enrollment in U2 arm of DLBCL cohort of UNITY-NHL

Q2

Top-Line ORR Results from UNITY-CLL

Q3

- Potential GENUINE BLA filing for Accelerated Approval
- Complete Enrollment in FL cohort of UNITY-NHL
- Complete Enrollment in U2+B arm of DLBCL cohort of UNITY-NHL

Q4

- Complete Enrollment in MZL cohort of UNITY-NHL
- Potential UNITY-CLL BLA/NDA for Accelerated Approval

# TG Therapeutics

**NASDAQ: TGTX**