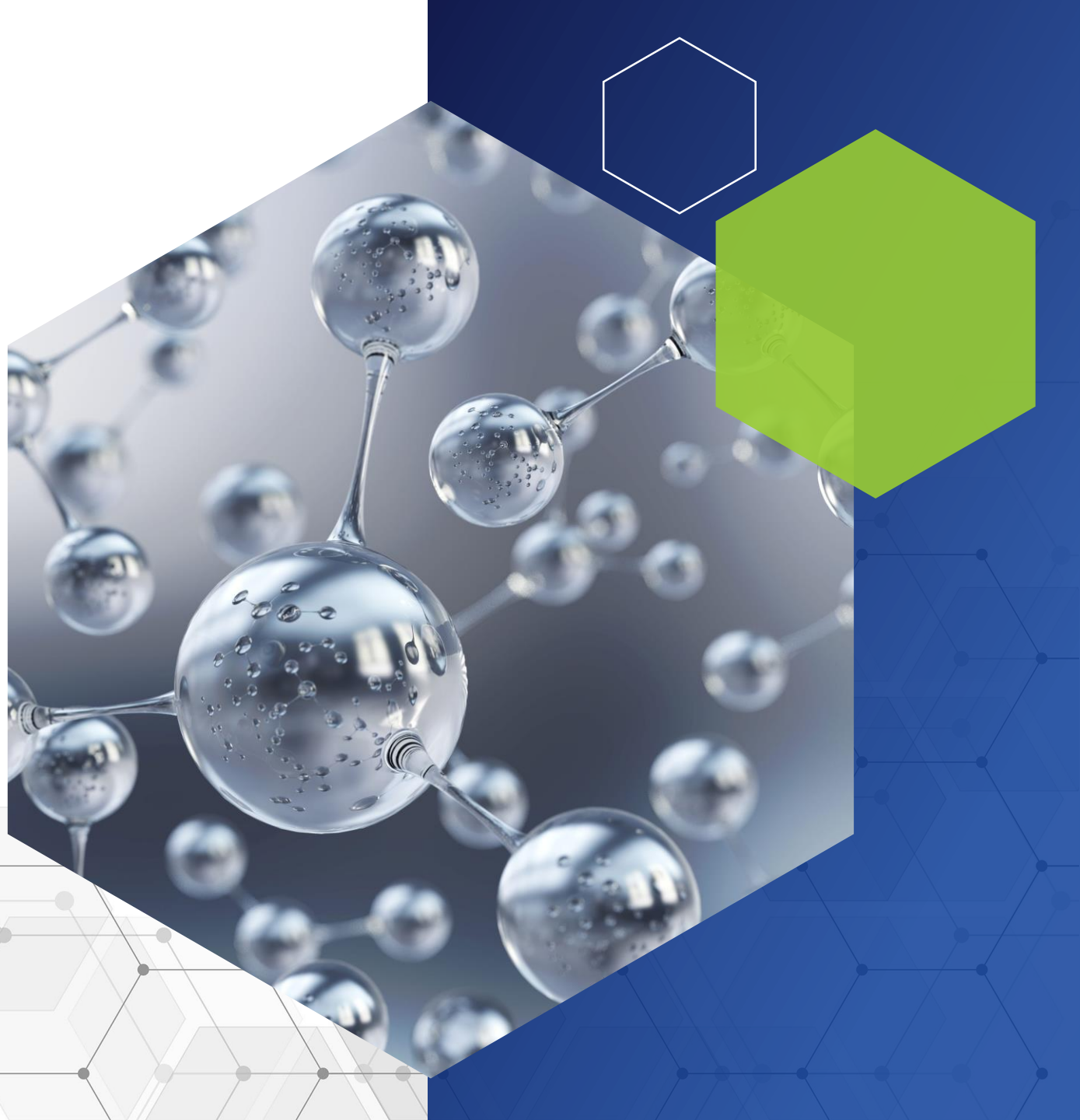




# J.P. Morgan Healthcare Conference

January 2020



# Forward Looking Safe Harbor Statement

*This presentation contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. These statements are often, but not always, made through the use of words or phrases such as "anticipates", "expects", "plans", "believes", "intends", and similar words or phrases. Such statements involve risks and uncertainties that could cause TG Therapeutics' actual results to differ materially from the anticipated results and expectations expressed in these forward-looking statements. These statements are only predictions based on current information and expectations and involve a number of risks and uncertainties. Actual events or results may differ materially from those projected in any of such statements due to various factors, including the risks and uncertainties inherent in clinical trials, drug development, and commercialization. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement and TG Therapeutics undertakes no obligation to update these statements, except as required by law.*

# Fearless Pursuit

- Founded 8 years ago with one goal in mind... to create the best possible treatment options for patients with B-Cell diseases
- Focused on Cancer & Autoimmune disease with registration programs ongoing in:
  - Chronic Lymphocytic Leukemia
  - Marginal Zone Lymphoma
  - Follicular Lymphoma
  - Multiple Sclerosis

125

EMPLOYEES

4

OFFICES

5

MEDICINES UNDER  
DEVELOPMENT

35

CLINICAL TRIALS

5

REGISTRATION-DIRECTED  
AND PHASE 3 CLINICAL  
TRIALS


1

GOAL: TO CURE B-CELL  
DISEASES



# B-Cell Focused Platform

## Clinical Stage Portfolio Overview

Drug Candidate	Mechanism of Action	Stage of Development
<b>Umbralisib</b> (TGR-1202)	 PI3K $\delta$ /CK1 $\epsilon$	Phase 3
<b>Ublituximab</b> (TG-1101)	Anti-CD20	Phase 3
<b>Cosibelimab</b> (TG-1501)	Anti-PD-L1	Phase 1b
<b>TG-1701</b>	BTKi	Phase 1
<b>TG-1801</b>	Anti-CD47/CD19	Phase 1

# UNITY-NHL MZL & FL Rolling Submission Initiated!

## TG THERAPEUTICS ANNOUNCES POSITIVE OUTCOME FROM UNITY-NHL PHASE 2B PIVOTAL TRIAL EVALUATING UMBRALISIB IN PATIENTS WITH RELAPSED/REFRACTORY MARGINAL ZONE LYMPHOMA



February 28, 2019 07:00 ET | Source: TG Therapeutics, Inc.

*Study met the primary endpoint*

*Interim data to be presented*

*Umbralisib was previously granted*

*Conference call to be held to*

NEW YORK, Feb. 28, 2019 (GlobeNewswire) — TG Therapeutics, Inc. (NASDAQ: TGTX) announced today that its Phase 2b pivotal trial evaluating the delta inhibitor, met the primary endpoint. The results met the Company's target guidance of overall response rate (ORR) of at least 40%.

*Interim data to be presented*

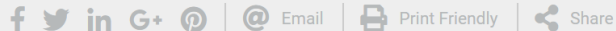
*Admission to the conference*

*by the U.S. Food and Drug Administration*

*Michael J. Gorman, MD, Chief Medical Officer*

*lymphoma*

## TG THERAPEUTICS ANNOUNCES POSITIVE RESULTS FROM THE UNITY-NHL PHASE 2B PIVOTAL TRIAL EVALUATING UMBRALISIB MONOTHERAPY IN PATIENTS WITH RELAPSED/REFRACTORY FOLLICULAR LYMPHOMA



October 28, 2019 07:00 ET | Source: TG Therapeutics, Inc.

*Follicular lymphoma cohort met the primary endpoint of overall response rate (ORR)*

*Umbralisib monotherapy appeared to be well tolerated with a safety profile consistent with previous reports*

### TODAY!

**TG Therapeutics Initiates Rolling Submission of New Drug Application (NDA) to U.S. Food and Drug Administration for Umbralisib as a Treatment for Patients with Previously Treated Marginal Zone Lymphoma and Follicular Lymphoma**

# 2020: Transformational Year w/ Multiple Pivotal Data Sets

## UNITY-NHL Rel/Ref MZL and FL

– *Umbralisib Monotherapy* –

Positive Topline ORR Data

BTD & ODD Granted for MZL

Rolling Submission Initiated,  
Target 1H 2020 Completion

## UNITY-CLL Front Line & Rel/Ref

– *Umbralisib + Ublituximab (U2)* –

Trial Conducted Under SPA

Enrolled ~420 patients

Awaiting PFS Data,  
Target Q1 2020

## ULTIMATE I & II Relapsing MS

– *Ublituximab Monotherapy* –

Trials Conducted under SPA

Enrolled Over 1100 Patients

Awaiting ARR Data,  
Target 2<sup>nd</sup> Half 2020

- *Potential for Approvals in MZL, FL, CLL, SLL and MS from 2020 to 2022*
- *Commercial Preparations Underway*

# Umbralisib:

*Investigational Targeted Therapy with "Best-in-Class" Potential*

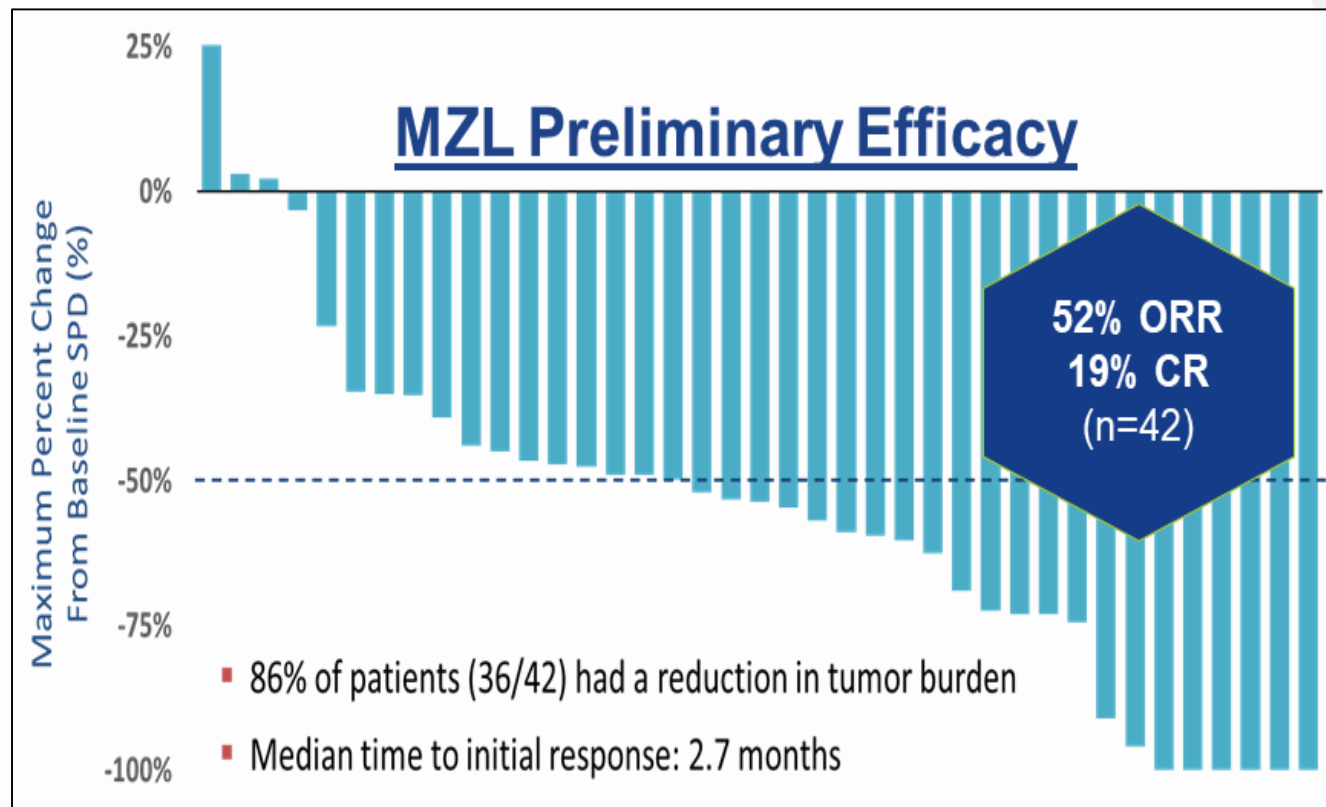
**Next Generation Dual PI3K-  
delta & CK1-epsilon Inhibitor**

Umbralisib



- *Demonstrated activity across NHL and CLL*
- *Potentially improved tolerability profile over 1<sup>st</sup> generation compounds*
- *Once daily v. twice daily for 1<sup>st</sup> generation compounds*

# Data Supporting Umbralisib NDA In Previously Treated MZL-*Breakthrough Therapy Designation Granted*



## MZL Preliminary Safety & Tolerability

- Umbralisib was deemed to be well tolerated
- No events of colitis reported
- AE's leading to dose reduction occurred in 6 subjects (9%)
- 10 subjects (14%) discontinued umbralisib due to an AE considered at least possibly related to treatment
- No deaths occurred on study
- Grade 3 infections were limited, occurring in 3 patients (bronchitis, pneumonia, and influenza)

- MZL trial met primary endpoint exceeding target 40% ORR amongst all patients (n=69)
- **INITIATED ROLLING SUBMISSION!**



# UNITY-NHL Umbralisib Monotherapy Follicular Lymphoma (FL)

- TG received guidance from the FDA allowing submission of a single NDA for Follicular Lymphoma (FL) and Marginal Zone Lymphoma (MZL) indications
- Follicular Cohort Met Primary End Point
  - Exceeding 40% ORR hurdle
- Umbralisib monotherapy appeared to be well tolerated with a safety profile consistent with previous reports
- Other PI3K delta's obtained accelerated approved with similar efficacy (range: 42% - 59% ORR)

## UNITY-NHL Trial FL Cohort

*Fully Enrolled*

**Umbralisib  
(TGR-1202)  
Monotherapy**

### Full Enrollment Complete

Target ORR	40-50%
Enrollment Complete	3Q-18
Target Full Data Presentation	2020

# MZL/FL: Incurable Disease with Limited Treatment Options

- Two largest forms of indolent NHL
- ~22,500 new cases per year<sup>1</sup> in MZL and FL
- ~6,000-10,000<sup>2</sup> relapsed indolent patients needing treatment each year



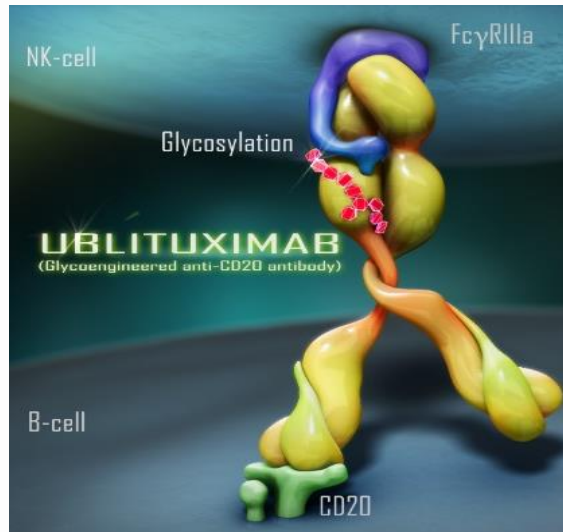
- **R<sup>2</sup>**: recently approved Revlimid plus Rituxan– can cause severe neutropenia and rash; secondary malignancy risks
- **Ibrutinib**: accelerated approval – less than half of patients respond; tolerability issues
- **PI<sub>3</sub>K Delta inhibitors**: none approved for MZL; 3 approved for FL tolerability issues have limited use

- Need for highly active, well-tolerated treatment option for MZL and FL
- **Umbralisib**: Convenient oral daily dosing appears well tolerated with ~40-50% ORR in iNHL

# Ublituximab:

## Investigational Next Generation Anti-CD20 Monoclonal Antibody

US REGULATORY  
SUBMISSION PLAN



CLL (*in combo with umbralisib*)

1H 2020

Multiple Sclerosis

YE 2020

### Ublituximab

Glycoengineered for enhanced potency over 1<sup>st</sup> generation

Activity in rituximab refractory patients

Shorter infusions than all other anti-CD20s (1-1.5 v 3-4 hours)

# Early Clinical Data for Umbralisib and U2 in CLL Support Phase 3



**87% ORR**

*Rel/Ref CLL  
U2 after three months  
In U2+Ven Phase 1 (n=23)*

## PFS: Rel/Ref CLL

Drug/Trial	PFS (mo.)
Umbralisib Phase 1	24
Umbralisib in BTK Intolerant	23.5
U2 Phase 1	28

## UNITY-CLL

*Enrollment Complete*

Randomize

U2

Gazyva +  
CHL

Study Enrollment ~420

Primary Endpoint PFS

Target PFS Readout 1Q20

Patient Population 1L/2L+

***Conducted under Special Protocol  
Assessment***

# CLL Market Opportunity

## *One of the Fastest Growing Global Hematology Markets*

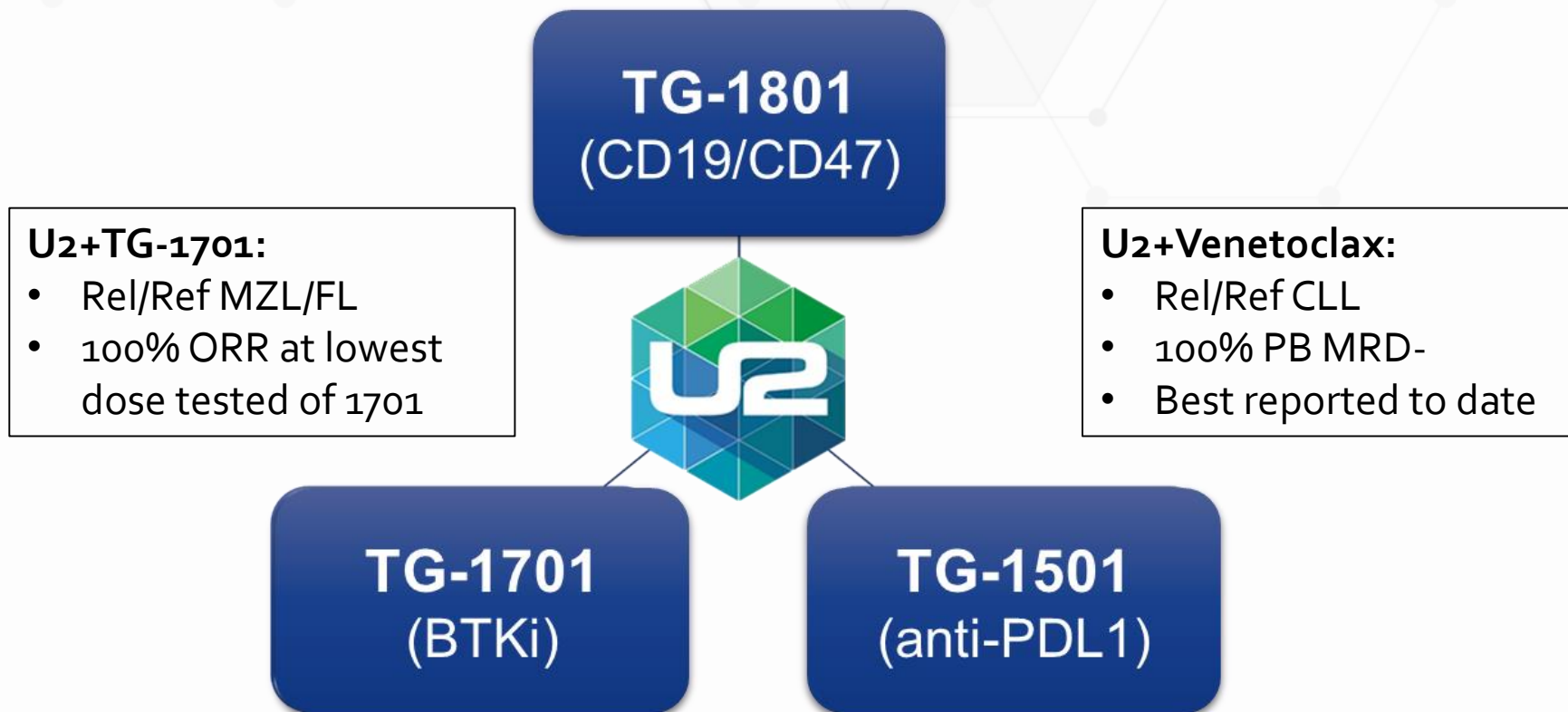
- ~115,000 Americans living with CLL
- ~20,000 newly-diagnosed patients each year
- ~20,000 previously treated patients seeking treatment each year



# U2 Has the Potential to Address Unmet Needs in CLL

- **Over 60,000 patients** in the US have been treated with a BTK-inhibitor creating a large population in the post-BTK setting in need of treatment options
  - **~20% of treatment naïve patients will be deemed poor candidates** for BTK inhibitors due to potential safety/tolerability concerns
  - **Up to 40% of patients will discontinue** BTK inhibitors for efficacy/tolerability issues at median of 17 months from starting therapy in recent real-world analysis
  - **Venetoclax represents a challenge for community practices** due to enhanced monitoring and hospitalization to avoid TLS
- ***Substantial need for additional highly active, well-tolerated treatments for CLL***

# B-Cell Platform Provides Next Gen Combo's



## Ublituximab + Umbralisib (U2) + Ibrutinib

Response Rate Observed with Triple Therapy

Type	Pts (n)	CR <sup>+</sup> (n)	PR (n)	ORR 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%)
DLBCL	6	-	1	1 (17%)
<b>Total</b>	<b>36</b>	<b>10</b>	<b>20</b>	<b>30 (83%)</b>

Nastoupil et al, Lugano 2017

## Ublituximab + Umbralisib + Pembro

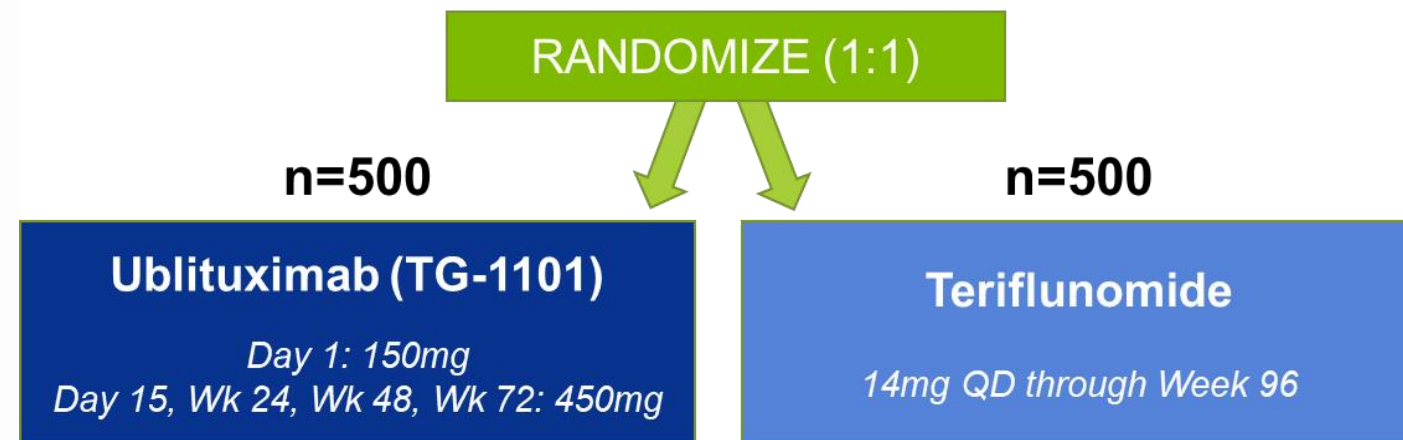
Response Rate Observed with Triple Therapy

Group	N	CR N (%)	PR N (%)	ORR N (%)
CLL	10	1 (10%)	8 (80%)	<b>9 (90%)</b>
RT	4	2 (50%)	0	<b>2 (50%)</b>

Mato, et al. ASH 2018

# Ublituximab in Relapsing Forms of Multiple Sclerosis

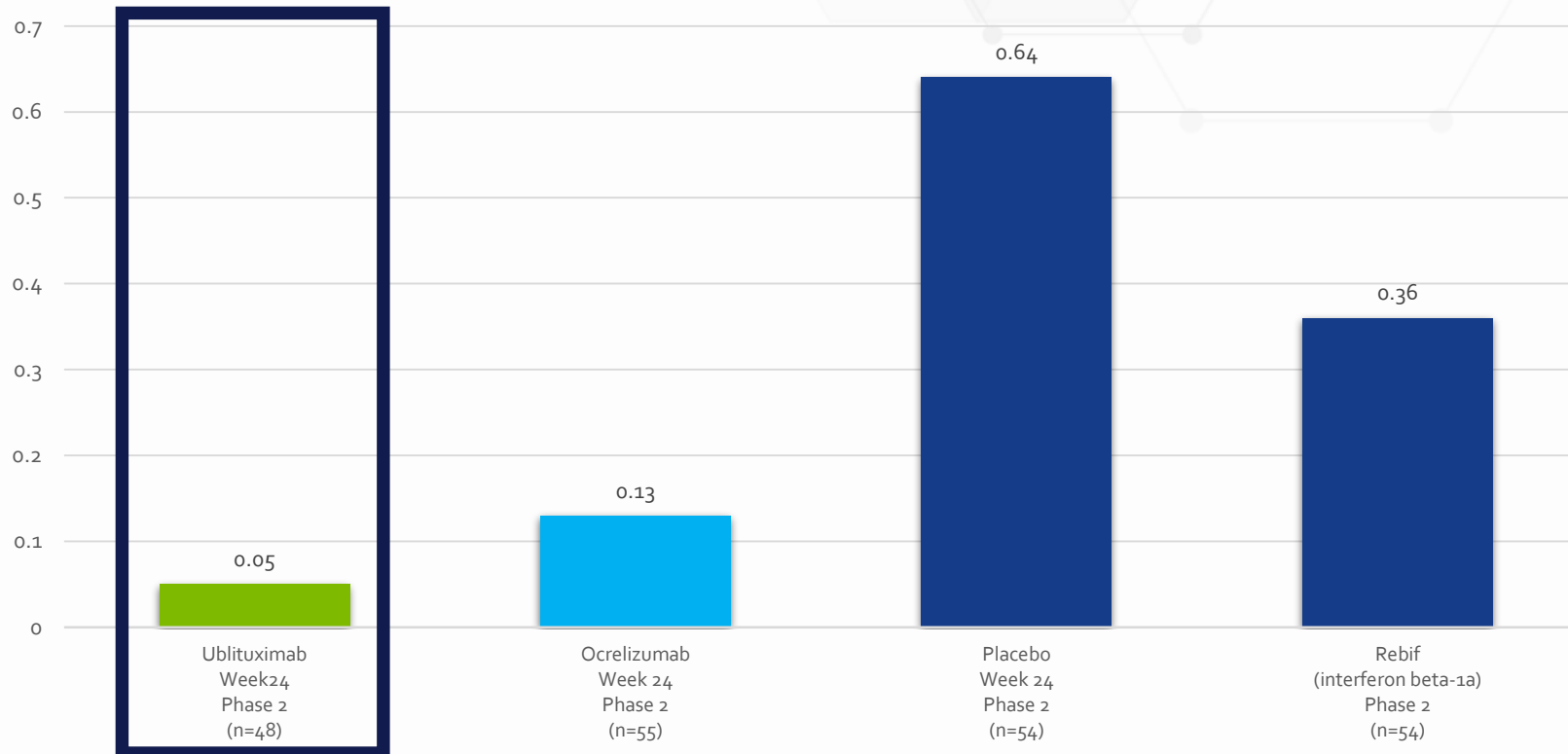
- ~1M Americans living with MS
- Completed Phase 2, data presented at multiple conferences
- Fully Enrolled Phase 3 ULTIMATE Trials
  - Special Protocol Assessment (SPA)
  - **Topline Data Targeted for 2H 2020**





# MS – Phase 2 ARR Comparison

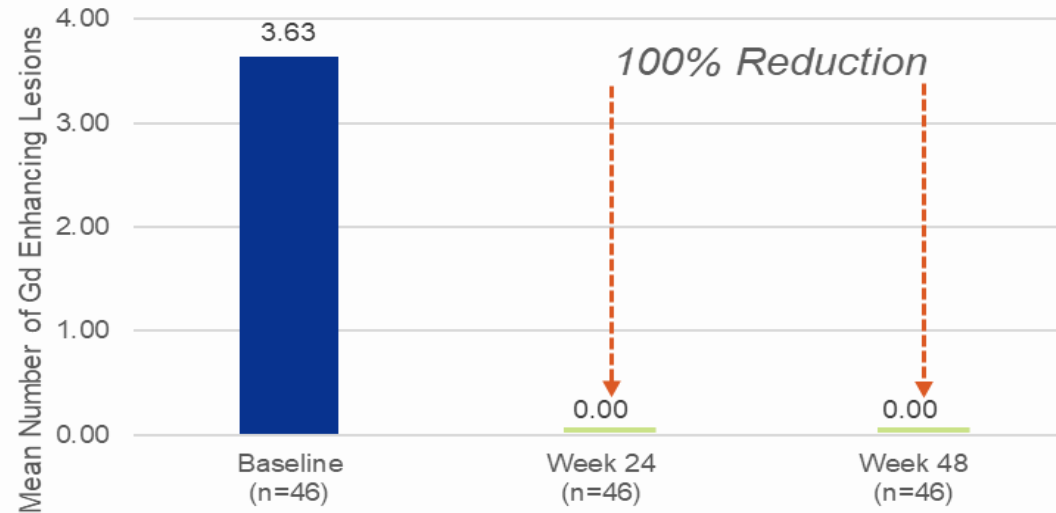
Annualized Relapse Rate (ARR) Comparator



- **Final Ublituximab Phase 2 Data at Week 48**
  - 48 patients through 48 weeks of treatment
  - Annualized Relapse Rate of .07

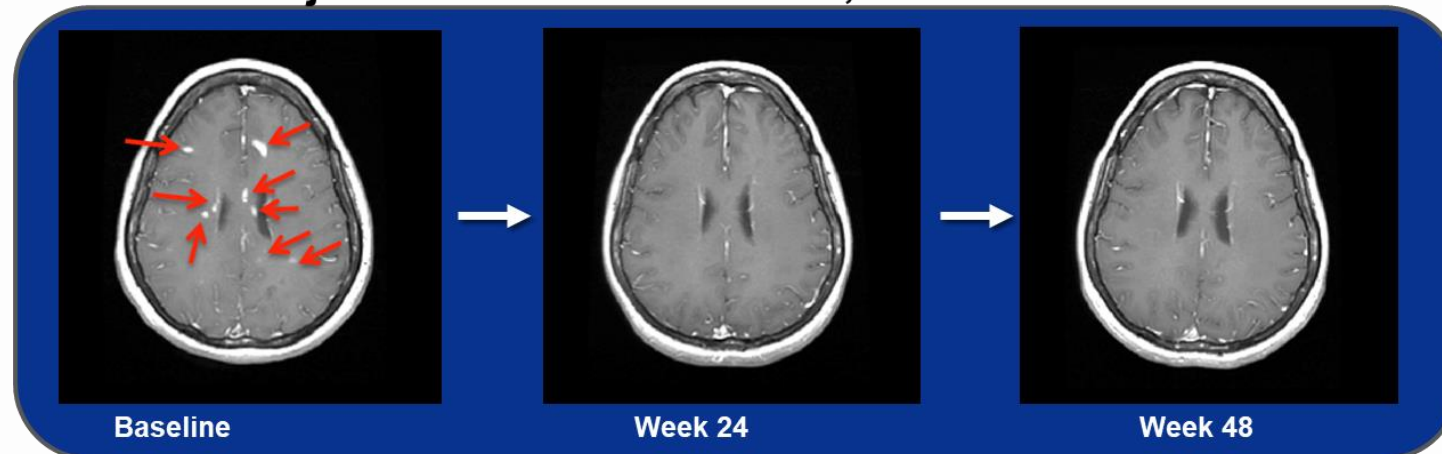
# Ublituximab Phase 2: MRI-Gd Enhancing Lesions

T1 Gd Enhancing Lesions Baseline vs. Week 24 & Week 48

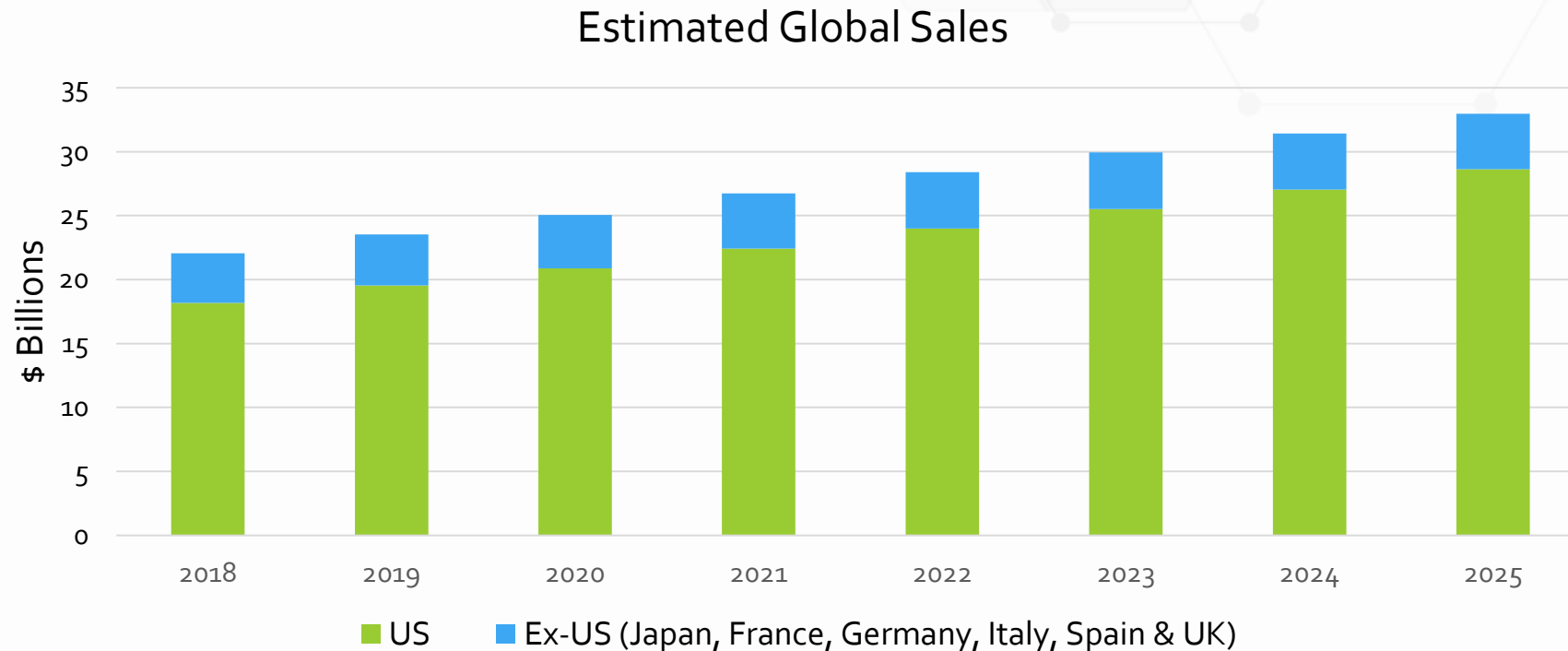


- No T1 Gd-enhancing lesions were detected in any subjects at Week 24 or Week 48 (100% reduction;  $p=0.003$ )

Subject T1 Gd MRI at Baseline, Week 24 & Week 48



# Significant Opportunity for Ublituximab in MS














Global Prevalence = ~2.3 Million

Global Market Size >\$30 Billion by 2025

- **Ocrelizumab >\$2 Billion in 2018 annual sales; on track to achieve ~\$4B in 2019**

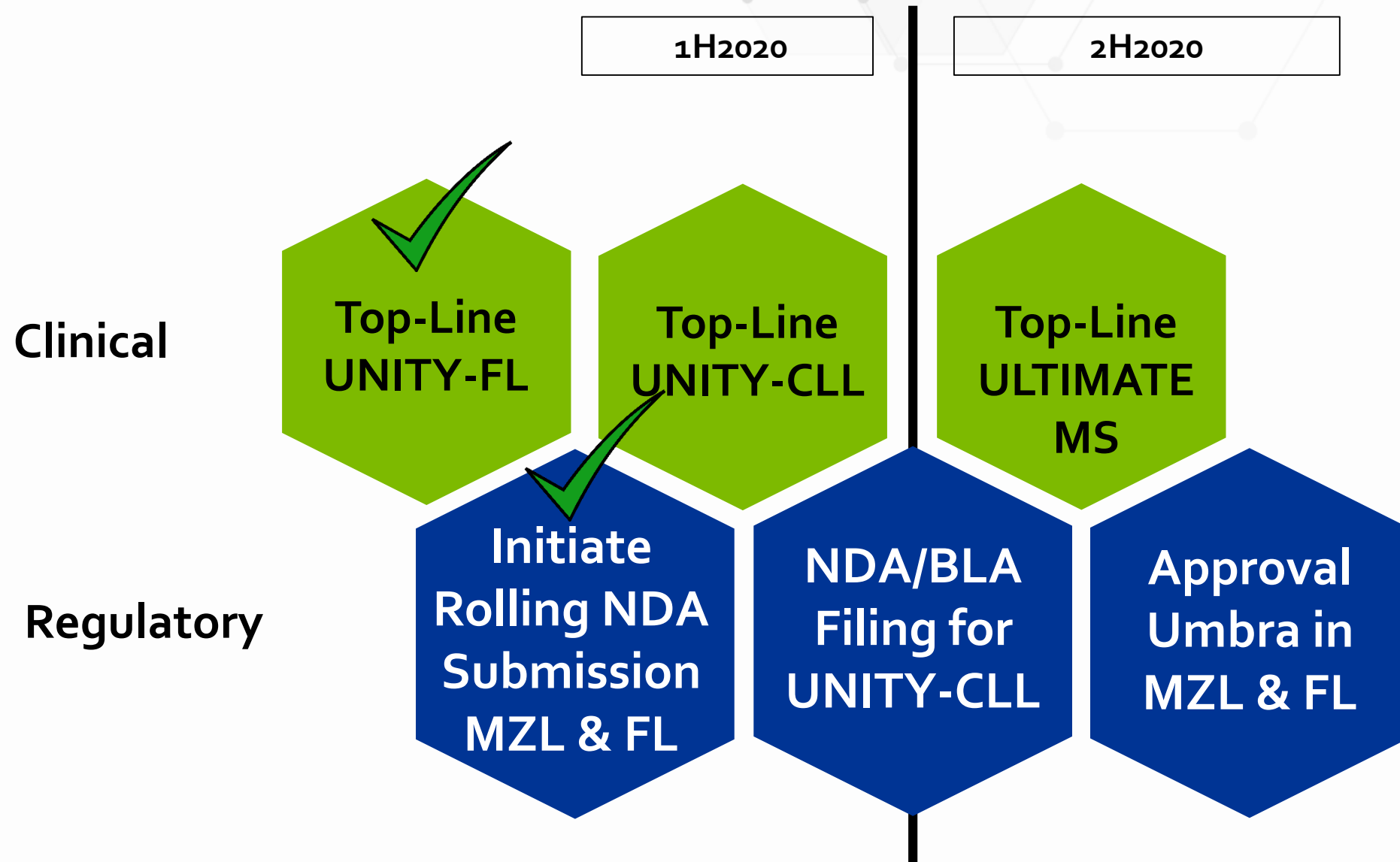
# Multiple Treatment Options Coexist & Account for Meaningful Market Share

Class	Drug	U.S. Approval			Global 2018 Revenues by Drug/Class (\$ in millions)
		Date	Route of Administration	Dosing	
Interferons	 BETASERON <sup>®</sup> <small>(interferon beta-1a)</small>	1993	Subcutaneously	1x / 2 days	\$643
	 AVONEX <sup>®</sup> <small>(interferon beta-1a)</small>	1996	Intramuscularly	1x / week	\$1,915
	 Rebif <sup>™</sup>	2002	Subcutaneously	3x / week	\$1,732
	 EXTAVIA <sup>®</sup>	2009	Subcutaneously	1x / 2 days	\$162
	 plegri <sup>dy</sup> <small>(peginterferon beta-1a)</small>	2014	Subcutaneously	1x / 2 weeks	\$448
IV Potent Immunomodulators	 TYSABRI <sup>®</sup> <small>(natalizumab)</small>	2004	Intravenously	1x / 4 weeks	\$1,864
	 LEMTRADA <sup>®</sup> <small>(alemtuzumab)</small>	2014	Intravenously	3x / year	\$475
T-Cell Targeted Orals	 GILENYA <sup>®</sup> <small>(tingolimod)</small>	2010	Orally	1x / day	\$3,380
	 AUBAGIO <sup>®</sup> <small>(teriflunomide)</small>	2012	Orally	1x / day	\$1,945
	 Tecfidera <sup>®</sup> <small>(dimethyl fumarate)</small>	2013	Orally	2x / day	\$4,274
CD20s	 OCREVUS <sup>®</sup> <small>(ocrelizumab)</small>	2017	Intravenously	2x / year	\$2,406
	Ofatumumab	2020	Subcutaneously	1x / 4 weeks	TBD
	Ublituximab	2021	Intravenously	2x / year	TBD

# Ublituximab Potential Value Proposition in MS

- Equal to better activity with comparable safety
- Convenience of 1 hour infusion every 6 months v. 3-4 hours for Ocrelizumab
- Strategically priced to optimize patient access
- ***Estimate \$1-2B annual market opportunity in the US alone for ublituximab in MS***

# Targeted Key Data & Potential Filings/Approvals



## Key Financial Statistics

**Ticker:** TGTX (NASDAQ)

**Price:** \$14.10 (close on 1/15/2020)

**Shares:** ~112M (fully-diluted, as of 12/31/19)

**Cash:** ~\$140M (as of 12/31/19)



# TG Therapeutics

**NASDAQ: TGTX**