



TREATING SLOW-GROWING NON-HODGKIN LYMPHOMAS

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INTRODUCTION
 Treating Slow-Growing Non-Hodgkin Lymphomas (NHL)



Lizette Figueroa-Rivera
 Sr. Director, Education & Support
 The Leukemia & Lymphoma Society

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DISCLOSURES

Treating Slow-Growing Non-Hodgkin Lymphomas (NHL)

Matthew S. McKinney, MD, has affiliations with Celgene, Epizyme, Kite/Gilead Sciences, Pharamacyclics, and Roche/Genentech (*Consultant*); Beigene, Celgene, Pharamacyclics, Novartis, and Roche/Genentech (*Grant Support*); Kite/Gilead Sciences (*Speakers Bureau*).

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OBJECTIVES

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- Slow-growing non-Hodgkin lymphomas (NHL)
- Treatment advances for slow-growing lymphomas
- Side- effects management
- Ways to effectively communicate with your healthcare team about quality-of-life issues

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WHAT ARE SLOW GROWING (INDOLENT) LYMPHOMAS?

- Lymphomas are cancers that form from part of the blood/lymph system
- Now there are greater than 50 recognized lymphoma diagnoses as recognized by World Health Organization
- Indolent or slow growing or low-grade lymphomas are entities that generally are incurable but do not grow rapidly in the body

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CURRENT LYMPHOMA CLASSIFICATION (WHO 2016)

Mature B-cell neoplasms

Chronic lymphocytic leukemia/small lymphocytic lymphoma
 Monoclonal B-cell lymphocytosis*
 B-cell prolymphocytic leukemia
 Splenic marginal zone lymphoma
 Hairy cell leukemia
 Splenic B-cell lymphoma/leukemia, unclassifiable
 Splenic diffuse red pulp small B-cell lymphoma
 Hairy cell leukemia-variant
 Lymphoplasmacytic lymphoma
 Waldenström' macroglobulinemia
 Monoclonal gammopathy of undetermined significance (MGUS), IgM*
 m heavy-chain disease
 s heavy-chain disease
 a heavy-chain disease
 Monoclonal gammopathy of undetermined significance (MGUS), IgG/M*
 Plasma cell myeloma
 Solitary plasmacytoma of bone
 Extraneoplastic plasmacytoma
 Monoclonal immunoglobulin deposition diseases*
 Extranodal marginal zone lymphoma of mucosa-associated lymphoid tissue (MALT lymphoma)
 Nodal marginal zone lymphoma
 Pediatric nodal marginal zone lymphoma
 Follicular lymphoma
 In situ, follicular neoplasia*
 Duodenal-type follicular lymphoma*
 Pediatric-type follicular lymphoma*
 Large B-cell lymphoma -with IRF4 rearrangement*
 Primary cutaneous follicle center lymphoma
 Mantle cell lymphoma
 In situ mantle cell neoplasia*
 Diffuse large B-cell lymphoma (DLBCL), NOS
 Germinal center B-cell type*
 Activated B-cell type*
 T-cell/histiocyte-rich large B-cell lymphoma
 Primary DLBCL of the central nervous system (CNS)
 Primary cutaneous DLBCL, leg type
 EBV* DLBCL, NOS*

EBV* mucocutaneous ulcer*
 DLBCL associated with chronic inflammation
 Lymphomatoid granulomatosis
 Primary mediastinal (thymic) large B-cell lymphoma
 Intravascular large B-cell lymphoma
 ALK* large B-cell lymphoma
 Plasmablastic lymphoma
 Primary effusion lymphoma
 HHV8* DLBCL, NOS*
 Burkitt lymphoma
 Burkitt-like lymphoma with 11q aberration*
 High-grade B-cell lymphoma, with MYC and BCL2 and/or BCL2 rearrangement* High-grade B-cell lymphoma, NOS*
 B-cell lymphoma, unclassifiable, with features intermediate between DLBCL and classical Hodgkin lymphoma

Hodgkin lymphoma
 Nodular lymphocyte predominant Hodgkin lymphoma
 Classical Hodgkin lymphoma
 Nodular sclerosis classical Hodgkin lymphoma
 Lymphocyte-rich classical Hodgkin lymphoma
 Mixed cellularity classical Hodgkin lymphoma
 Lymphocyte-depleted classical Hodgkin lymphoma
 Posttransplant lymphoproliferative disorders (PTLD)
 Plasmacytic hyperplasia PTLD
 Infectious mononucleosis PTLD
 Florid follicular hyperplasia PTLD*
 Polymorphic PTLD
 Monomorphic PTLD (B- and T-NK-cell types)
 Classical Hodgkin lymphoma PTLD

Mature T and NK neoplasms

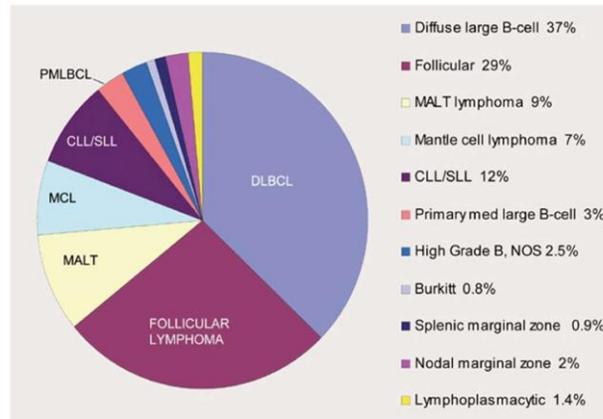
T-cell prolymphocytic leukemia
 T-cell large granular lymphocytic leukemia
 Chronic lymphoproliferative disorder of NK cells
 Aggressive NK-cell leukemia
 Systemic EBV⁺ T-cell lymphoma of childhood*
 Hydroa vacciniforme-like lymphoproliferative disorder*
 Adult T-cell leukemia/lymphoma
 Extranodal NK-/T-cell lymphoma, nasal type
 Enteropathy-associated T-cell lymphoma
 Monomorphic epitheliotropic intestinal T-cell lymphoma*
 Indolent T-cell lymphoproliferative disorder of the GI tract*
 Hepatosplenic T-cell lymphoma
 Subcutaneous panniculitis-like T-cell lymphoma
 Mycosis fungoides
 Sezary' syndrome
 Primary cutaneous CD30⁺ T-cell lymphoproliferative disorders
 Lymphomatoid papulosis
 Primary cutaneous anaplastic large cell lymphoma
 Primary cutaneous gd T-cell lymphoma
 Primary cutaneous CD8⁺ aggressive epidermotropic cytotoxic T-cell lymphoma
 Primary cutaneous acral CD8⁺ T-cell lymphoma*
 Primary cutaneous CD4⁺ small/medium T-cell lymphoproliferative disorder
 Peripheral T-cell lymphoma, NOS
 Angioimmunoblastic T-cell lymphoma
 Follicular T-cell lymphoma*
 Nodal peripheral T-cell lymphoma with TFH phenotype*
 Anaplastic large-cell lymphoma, ALK⁺
 Anaplastic large-cell lymphoma, ALK⁻*
 Breast implant-associated anaplastic large-cell lymphoma*

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DISTRIBUTION OF LYMPHOMA SUBTYPES



Jaffe, WHO 2008

Indolent lymphomas:

Follicular lymphoma
Marginal zone or MALT lymphoma
Lymphoplasmacytic lymphoma/
Waldenström
macroglobulinemia
CLL/SLL

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QUESTIONS TO ASK AT DIAGNOSIS?

- Is the biopsy sample adequate to make the diagnosis?
- What is stage?
 - Mostly important for limiting treatment, less for prognosis
- What markers indicate the patient's prognosis?
 - Different than same question having to do with staging
- What is the best observation or treatment plan?

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SPECIAL CHALLENGES OF LIVING WITH SLOW GROWING LYMPHOMAS

- Most indolent lymphomas are incurable, and patients deal with chronically
- Indolent lymphomas can cause serious health problems
- Therapies for lymphoma can have significant side effects
- It is important to address social, family, mental and financial stressors brought on by the challenges of dealing with a slow growing lymphoma
- Patients deserve a personalized “30 year plan”

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LYMPHOMA TREATMENT OPTIONS/MODALITIES

- Chemotherapy
- Radiation
- Immunotherapies
(antibodies, radioimmunotherapy, checkpoint inhibitors, bispecific antibodies)
- Small molecule inhibitors
- Stem cell transplant (autologous = self, allogeneic = donor infusion)
- Cell therapy (chimeric antigen receptor modified T cells = CAR T cells)

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LOW GRADE/INDOLENT LYMPHOMA PRINCIPLES OF TREATMENT

- Early stage (usually stage I) lymphomas may be amenable to curative radiation treatment
- Otherwise treatment should only be administered for symptoms and using GELF or similar criteria (iwCLL18, IWWM etc.)

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STAGE I-II DISEASE

- 100 pts w/stage I/II FL Radiation +/- chemotherapy
- Freedom from Tx Failure (FFTF)
 - 46% 10 years
 - 39% 15 years
- Overall survival:
 - 10 year 75%
 - 15 year 62%
 - 57% deaths from lymphoma
- No difference in outcomes +/- chemotherapy

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Guadagnolo et al. Int J Rad Onc Biol Phys 2006;64, 928-934.



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WATCH AND WAIT STRATEGIES FOR LOW GRADE (INDOLENT) LYMPHOMAS

	Watch and Wait	ProMACE-MOPP + XRT
Patients	41	43
Alive off therapy	5/16 (31%)	25/43 (58%)
Alive without disease	5/41 (12%)	22/43 (51%)
Alive, continuously free of disease	0/41 (0%)	22/43 (51%)
Alive	34/41 (83%)	36/43 (84%)

TREATMENT PROGRAMS FOR INDOLENT LYMPHOMAS (ADVANCED DISEASE)

- Goal of treatment is to decrease symptoms and improve patient survival; patients doing well do not need treatment
- Several regimens exist for follicular lymphoma others
- Bendamustine based regimens give longest response in most patients
- We may be moving toward chemotherapy free approaches
- Relapsed disease may also be treated with only novel agents

GELF CRITERIA

- Single node > 7 cm
- More than nodal sites > 3 cm
- Systemic symptom(s)
- Compression syndrome or serous effusion
- Cytopenia
- Lymphocyte count > 50,000/ μ L

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Journal of Clinical Oncology 1997; 15: 1110-7.

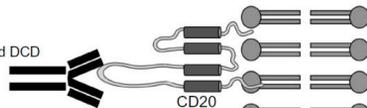


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CD20—MONOCLONAL ANTIBODY IMMUNOTHERAPY

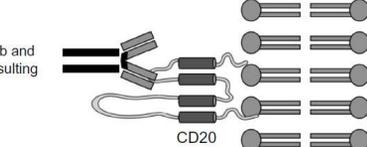
Rituximab:

Binds to large loop of CD20 resulting in CDC, ADCC, and DCD



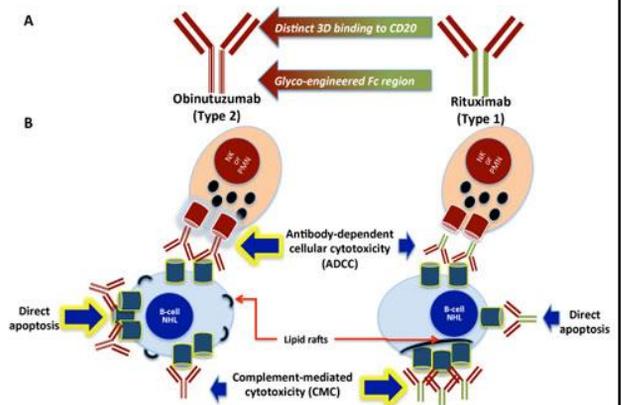
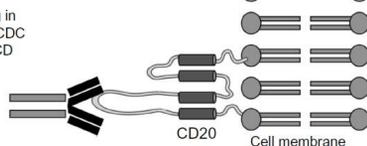
Ofatumumab:

Binds to small loop of CD20 slower off time than rituximab and improved binding of C1q, resulting in improved CDC



Obinutuzumab:

Fc₁ region modified resulting in improved ADCC but worse CDC type 2 Ab, with improved DCD



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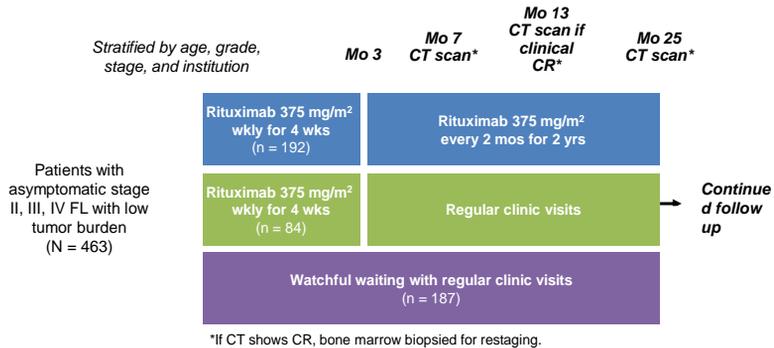
Blood and Lymphatic Cancer: Targets and Therapy 2015 5: 43—53



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RITUXIMAB VS WATCHFUL WAITING IN ASYMPTOMATIC FL: IS TREATMENT NEEDED?

Randomized phase III study



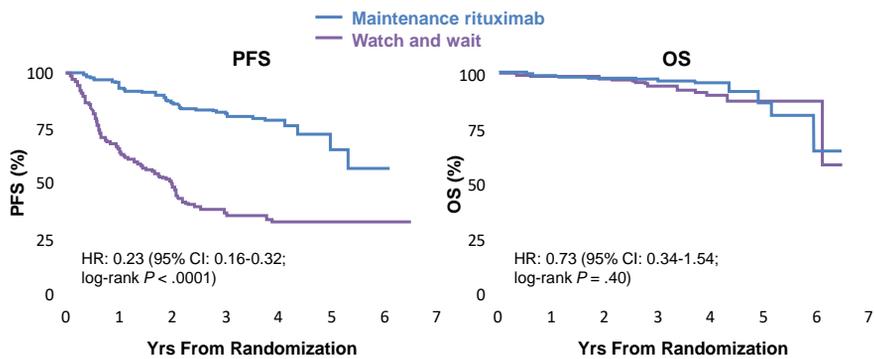
Slide credit: clinicaloptions.com

BEATING CANCER IS IN OUR BLOOD.

Ardeshna KM, et al. Lancet Oncol. 2014;15:424-435.



RITUXIMAB VS WATCHFUL WAITING IN ASYMPTOMATIC FL: SURVIVAL



Slide credit: clinicaloptions.com

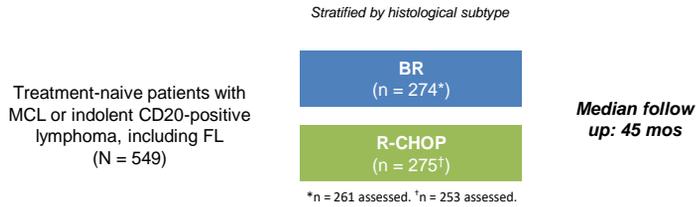
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Ardeshna KM, et al. Lancet Oncol. 2014;15:424-435.



STIL NHL 1-2003: BR VS R-CHOP IN NEWLY DIAGNOSED FL

Randomized, open-label phase III noninferiority trial



BR: bendamustine 90 mg/m² on Days 1-2; rituximab 375 mg/m² on Day 1; 4-wk cycles for 6 cycles max.
R-CHOP: cyclophosphamide 750 mg/m² on Day 1; doxorubicin 50 mg/m² on Day 1; vincristine 1.4 mg/m² on Day 1; prednisone 100 mg on Days 1-5; rituximab 375 mg/m² on Day 1; 3-wk cycles for 6 cycles max. No maintenance or consolidation treatment given.

- Primary endpoint: noninferiority of BR vs R-CHOP for PFS (decrease < 10% at 3 yrs)
- Secondary endpoints: response rate, time to next treatment, EFS, OS, safety

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Slide credit: clinicaloptions.com

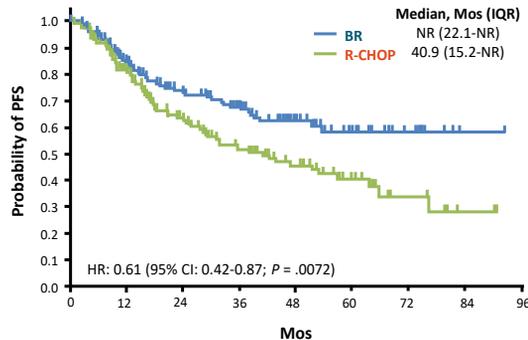
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Rummel MJ, et al. Lancet. 2013;381:1203-1210.



STIL NHL 1-2003: PFS IN FL

No OS difference between treatment arms (for both, median OS: NR)



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Slide credit: clinicaloptions.com

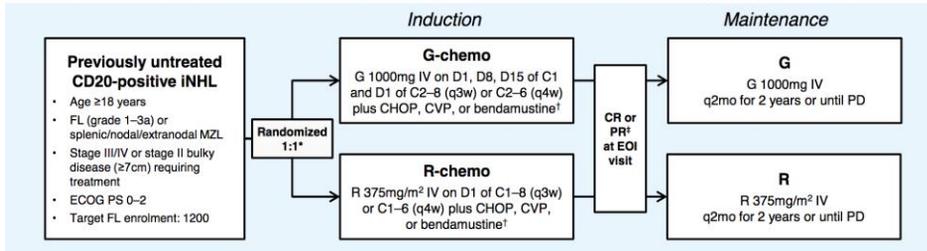
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Rummel MJ, et al. Lancet. 2013;381:1203-1210.



OBINUTUZUMAB BASED CHEMOIMMUNOTHERAPY FOR FL: PHASE III GALLIUM STUDY

International, open-label, randomized Phase III study



Primary endpoint

- PFS (INV-assessed in FL)

Secondary and other endpoints

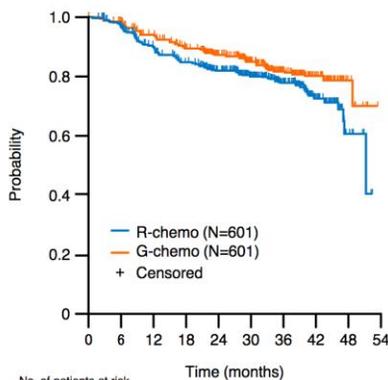
- PFS (IRC-assessed)[§]
- OS, EFS, DFS, DoR, TTNT
- CR/ORR at EOI (+/- FDG-PET)
- Safety

*FL and MZL pts were randomized separately; stratification factors: chemotherapy, FL/PI (FL) or IPI (MZL) risk group, geographic region; [†]CHOP q3w × 6 cycles, CVP q3w × 8 cycles, bendamustine q4w × 6 cycles; choice by site (FL) or by pt (MZL); [‡]Pts with SD at EOI were followed for PD for up to 2 years; [§]Confirmatory endpoint

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PROGRESSION FREE SURVIVAL WITH OBINUTUZUMAB VERSUS RITUXIMAB WITH CHEMOTHERAPY IN FL



	R-chemo, n=601	G-chemo, n=601
Pts with event, n (%)	125 (20.8)	93 (15.5)
3-yr PFS, % (95% CI)	77.9 (73.8, 81.4)	81.9 (77.9, 85.2)
HR (95% CI), p-value*	0.71 (0.54, 0.93), p=0.0138	

Median follow-up: 34.5 months

No. of patients at risk	0	6	12	18	24	30	36	42	48	54
R-chemo (N=601)	601	563	500	460	372	263	160	66	10	0
G-chemo (N=601)	601	569	528	491	385	270	162	73	10	0

*Stratified analysis; stratification factors: chemotherapy regimen, FL/IPI risk group, geographic region

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DIFFERENCES IN OBINUTUZUMAB VERSUS RITUXIMAB-CHEMOTHERAPY TOXICITIES

% (n)	R-chemo (n=597)	G-chemo (n=595)
Any AE	98.3% (587)	99.5% (592)
Grade ≥3 AEs (≥5% in either arm)	67.8% (405)	74.6% (444)
Neutropenia	37.9% (226)	43.9% (261)
Leucopenia	8.4% (50)	8.6% (51)
Febrile neutropenia	4.9% (29)	6.9% (41)
IRRs*	3.7% (22)	6.7% (40)
Thrombocytopenia	2.7% (16)	6.1% (36)
Grade ≥3 AEs of special interest by category (selected)		
Infections†	15.6% (93)	20.0% (119)
IRRs‡	6.7% (40)	12.4% (74)
Second neoplasms §	2.7% (16)	4.7% (28)
SAEs	39.9% (238)	46.1% (274)
AEs causing treatment discontinuation	14.2% (85)	16.3% (97)
Grade 5 (fatal) AEs	3.4% (20)	4.0% (24)**
Median (range) change from baseline in IgG levels at end of induction, g/l¶	-1.46 (-16.4–9.1)††	-1.50 (-22.3–6.5) ††

*As MedDRA preferred term; †All events in MedDRA System Organ Class 'Infections and Infestations'; ‡Any AE occurring during or within 24h of infusion of G or R and considered drug-related; §Standardized MedDRA query for malignant or unspecified tumors starting 6 mo after treatment start; ¶Ig levels were measured during screening, at EOI and end of maintenance and during follow-up; **Includes patient who died after clinical cut-off date from AE starting before cut-off date; ††n=472; ††n=462

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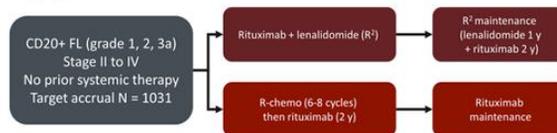


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CHEMOTHERAPY FREE APPROACH IN FOLLICULAR LYMPHOMA (FL)

RELEVANCE Trial

Ongoing Phase 3 Trial—Lenalidomide + Rituximab^[a]



- R-chemo: investigator's choice of R-CHOP, R-CVP, or BR
- Primary endpoint: CR/Cru rate at 120 wk, PFS
- Secondary endpoint: EFS, TTNT, OS, MRD using PCR, and HRQoL
- In a single-center trial, patients with untreated FL who received the combination of rituximab + lenalidomide had an ORR of 98% and a CR rate of 87%^[b]

a. ClinicalTrials.gov. NCT01650701.

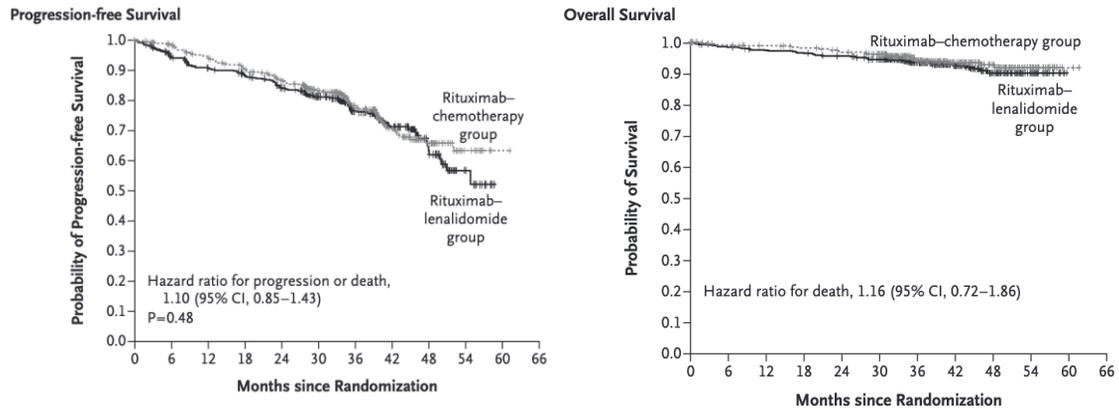
b. Fowler NH, et al. *Lancet Oncol.* 2014;15:1311-1318.

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SIMILAR OUTCOMES WITH CHEMO-FREE APPROACH IN FL



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New England Journal of Medicine 2018



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ADVANCED FOLLICULAR LYMPHOMA APPROACH

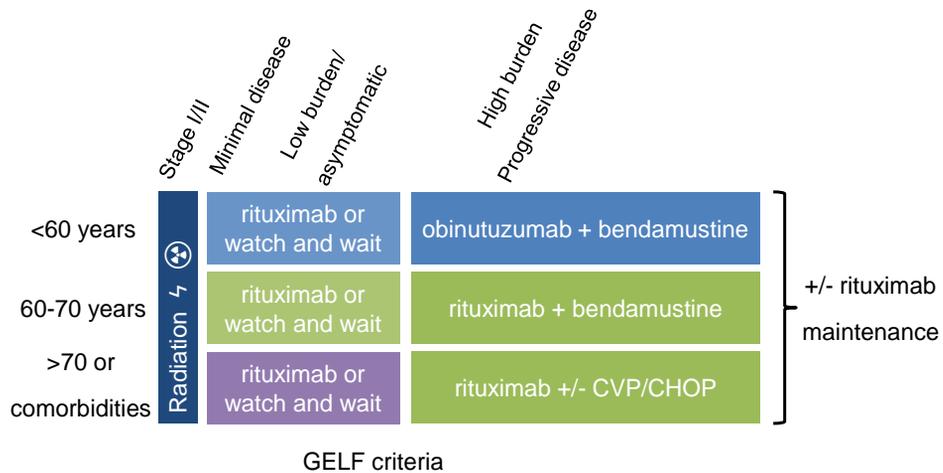
- I recommend observation for patients not symptomatic from their lymphoma
- If treatment is needed options range from chemo-free approach to aggressive regimens such as obinutuzumab-bendamustine
- Each patient's treatment must be individualized based on preferences and underlying health
- Most patients need multiple specific treatment regimens over many years
- CD20 antibody maintenance can be offered but is an individualized decision—it improves progression free time but not overall survival

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SIMILAR OUTCOMES WITH CHEMO-FREE APPROACH IN FL



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APPROACH TO RELAPSED FOLLICULAR LYMPHOMA

- Most patients will undergo multiple therapies for follicular lymphoma
- Treatment approach should be individualized but we generally look toward novel agents (e.g. lenalidomide/rituximab)
- Multiple new therapies recently approved so we are moving away from chemotherapy and stem cell transplant.

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IMPORTANT RECENT FDA APPROVALS FOR NEW LYMPHOMA DRUGS

- Follicular lymphoma
 - Obinutuzumab frontline treatment
 - Lenalidomide with rituximab
 - Duvelisib
 - Tazemetostat
 - Umbralisib
 - Axicabtagene ciloleucel
- Marginal zone lymphoma
 - Ibrutinib
 - Lenalidomide with rituximab
 - Umbralisib
- Waldenstrom macroglobulinemia
 - Ibrutinib with rituximab (the only FDA approved therapy in Waldenstrom's)
 - Zanubrutinib indication filed with FDA

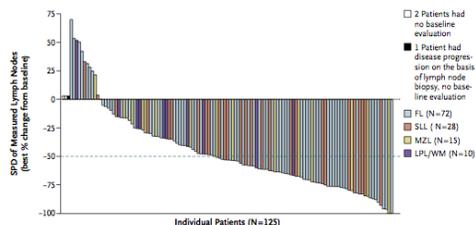
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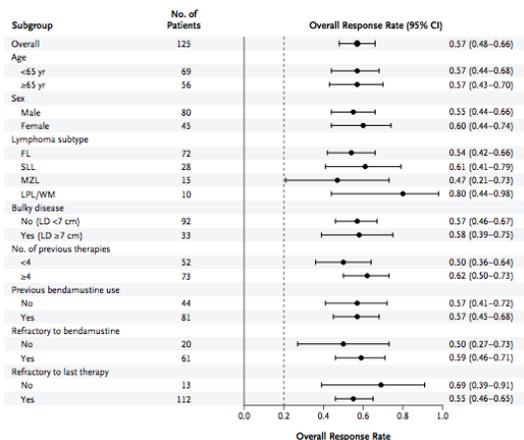
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PI-3 KINASE INHIBITORS FOR FL

- Phosphatidylinositol-3-kinase delta (PI3K δ) inhibitor
- 57% response rate, 1.9 months to response, durability 12.5 months
- Phase II protocol
 - 125 patients
 - Refractory/relapsed within 6 months of rituximab/alkylator



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Gopal, et. al. NEJM (2014)



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CONSIDERATIONS FOR PI3K INHIBITOR SELECTION

- All 3 FDA-approved PI3K inhibitors have shown similar efficacy in the setting of relapsed/refractory FL
- Different toxicity profiles may factor into choice of PI3K inhibitor, particularly in patients with comorbidities
 - Hepatic toxicity and immune-related colitis are the most clinically concerning with idelalisib and duvelisib, hyperglycemia and hypertension with copanlisib
- Route of administration is another difference among PI3K inhibitors
 - Idelalisib and duvelisib are taken orally, copanlisib is administered by IV
- Choice of therapy should be individualized

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PRECISION MEDICINE HAS ARRIVED FOR FOLLICULAR LYMPHOMA--TAZEMETOSTAT

- 25-30% of follicular lymphomas have mutation in gain of function mutations in *EZH2* (most often codon Y646)
- Tazemetostat is a selective *EZH2* inhibitor that reduces *EZH2* mutant related H3K27me3
- Tazemetostat approved 6/18/2020 for FL after 2 or more lines of therapy in *EZH2*-mutated FL
- Side effect profile favorable relative to lenalidomide, PI-3 kinase inhibitors

	EZH2 mutated lymphoma	EZH2 wild type
Partial response rate %	57	34
Complete response %	12	4

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UPDATE(S) ON UPCOMING NEW THERAPIES

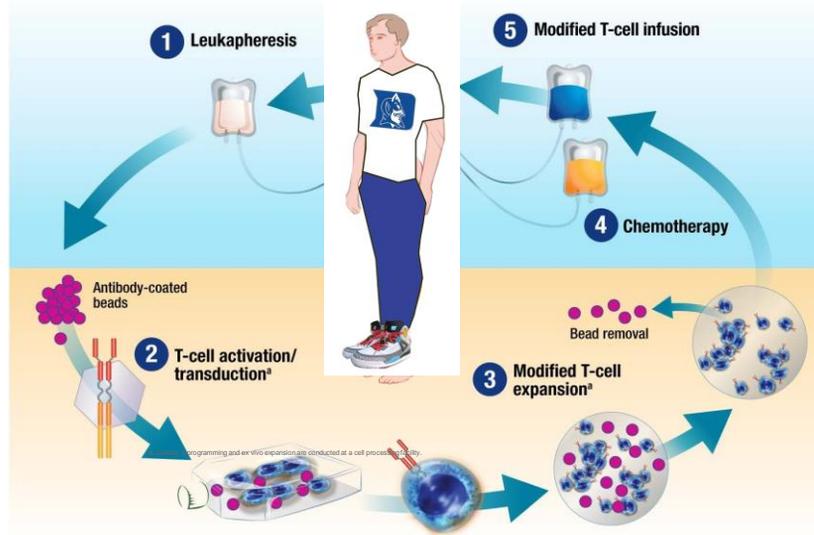
- Novel approaches can be classified into 3 types:
 - New applications of existing therapies (e.g. stem cell transplantation in certain subgroups or new combinations)
 - Molecularly targeted agents
 - Specifically pairing characteristics of patient's tumor to a drug
 - May be guided by new laboratory studies
 - Targeted "Smartbomb" delivery of chemotherapy agents in tumor cells
 - Immunotherapy
 - Immune "checkpoint" blockade
 - Modified activated T cell therapies
 - T cell engaging bi-specific antibodies (BiTEs)

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CTL019 IS DESIGNED TO HUNT AND DESTROY CD19-POSITIVE B-CELL CANCERS IN PATIENTS



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Figure courtesy of Novartis



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CAR T CELL TREATMENT IN LYMPHOMAS

- B cell lymphoma can be treated CAR T cells directed against the CD19 protein (among others)
- Response rates high in studied patients with lymphoma where other therapies have failed
- Therapy is complicated, expensive and requires inpatient hospitalization for side effect monitoring
- Numerous trials are now evaluating CAR T cells for other lymphoma types
 - Recent approvals for mantle cell lymphoma, follicular lymphoma

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ZUMA-5: PHASE II TRIAL OF AXICABTAGENE CILOLEUCEL (AXI-CEL) IN HIGH-RISK R/R INDOLENT NHL

Patients with high risk* indolent FL or MZL after ≥ 2 prior lines of CIT; ECOG PS 0/1; no CNS involvement or transformed disease (planned N = 160; n = 96 for efficacy analysis†)

Leukapheresis

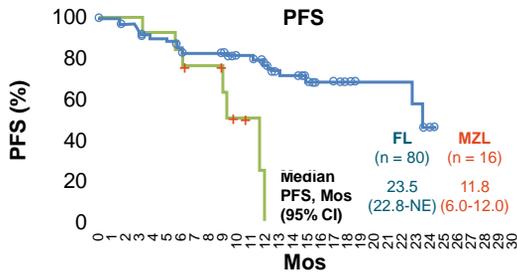
Lymphodepleting Conditioning Regimen

Cyclophosphamide + Fludarabine on Days -5 to -3

Axicabtagene Ciloleucel[®] IV on Day 0

Patients followed up to 15 yrs for safety

*High risk: with POD24, relapse post ASCT, or PD within 6 mos of second-line CIT or beyond.
 †n = 80 with FL and ≥ 9 mos of follow-up; n = 16 with MZL and ≥ 1 mo of follow-up. Axi-cel: CD19-directed CAR T-cell therapy.



- Manageable toxicity profile with axi-cel; early onset of AEs, generally reversible
 - 1 grade 4, no grade 5 neurologic events; events ongoing in 4 patients at data cutoff
 - 1 grade 5 CRS event; no ongoing CRS at data cutoff

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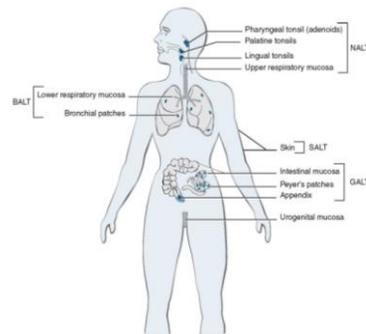
Jacobson. ASCO 2020. Abstr 8008. NCT03105336.



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MARGINAL ZONE LYMPHOMA

- 3 types:
 - Extranodal (MALT) lymphomas
 - Mucosa associated lymphatic tissue
 - Nodal MZL
 - Splenic MZL
- Association with chronic antigenic stimulation by infection or autoantigens in lymph tissues
- 70% are mucosal associated lymphoid tissue (MALT) lymphomas
- Gastric MALT lymphoma in 30% of cases



Ocular adnexal marginal zone lymphoma

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TREATMENT FOR MZL/MALT LYMPHOMAS

- Consideration for cure in early-stage disease (gastric MALT most common scenario)
- Treat infection if present followed by observation (*H. pylori* eradication and upper stomach endoscopy surveillance in gastric MALT)
- Radiation can be considered in early-stage disease if antibiotic treatment not successful
- Rituximab, chemotherapy used for extensive stage symptomatic lymphoma
- New agents approved for MZL include lenalidomide, ibrutinib, umbralisib after failure of chemo/immunotherapy

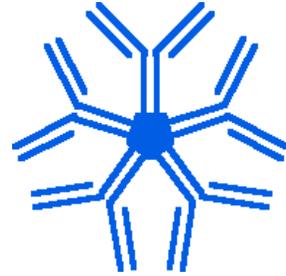
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WALDENSTRÖM MACROGLOBULINEMIA/ LYMPHOPLASMACYTIC LYMPHOMA (WM/LPL)

- Waldenström macroglobulinemia (WM) is an indolent process where an underlying LPL or MZL secretes IgM protein
- IgM can cause blood hyperviscosity and that can cause seizures, bleeding, vision changes
- Most common WM/LPL symptoms are fatigue, anemia, neuropathy
- Treatment aimed at alleviating symptoms of WM/LPL
- Rituximab, chemotherapy, proteasome inhibitors and ibrutinib are most commonly used treatments



IgM pentamer
complex

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NEW AGENTS IN LYMPHOMA AND WHAT TO LOOK FOR NEXT

- Novel cell therapies and new agents are offering new options for patients across diseases
- Treatment of chemotherapy-refractory diffuse large B cell lymphoma example of progress in the field
- Upcoming advances to look for:
 - Better combination treatment for T cell lymphomas
 - CAR T cell approvals outside of DLBCL (?mantle cell or aggressive FL)
 - Chemotherapy free approaches
 - New molecules with activity

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SUMMARY (1)

- There are many complex treatment programs for various lymphomas
- Hopefully we will continue to come up with new treatments and cure more patients

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SIDE EFFECTS OF SLOW GROWING LYMPHOMA TREATMENTS

Medication	administration	Most common side effects
rituximab	IV	infusion reactions, infections
bendamustine	IV	low blood counts, infections, rash, fatigue, nausea
CHOP	IV and oral	alopecia, low blood counts, infections, heart toxicity, neuropathy, nausea
lenalidomide	oral	low blood counts, diarrhea, rash
tazemetostat	oral	nausea, low blood counts
idelalisib	oral	liver toxicity, low blood counts, infection, diarrhea, lung damage
duvelisib	oral	liver toxicity, low blood counts, infection, diarrhea, lung damage
copanlisib	IV	low blood counts, diarrhea, high blood sugar, high blood pressure
axicabtagene ciloleucel	hospitalization	low blood counts, cytokine release syndrome, neurotoxicity
ibrutinib	oral	bleeding, atrial fibrillation, rash, joint and muscle pain
radiation	daily treatments on gantry	skin burn, nausea, fatigue, organ damage, risk for secondary leukemia

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TALKING WITH YOUR DOCTOR ABOUT SIDE EFFECTS OF TREATMENT

- Side effects, route of administration, schedule and cost are important factors to consider in selecting and dealing with lymphoma therapy
- There are a wide range of toxicities/side effects across treatment options
- There are a number of simple solutions to side effects (e.g. steroids for rash, caffeine for acalabrutinib headache, anti-diarrheals, etc.)
- Knowledge is power—know the potential side effects
- Communication is key—make sure you share symptoms and concerns with your physician and how these affect you

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HOW TO HELP FRIENDS/FAMILY DEAL WITH LYMPHOMA

- Caregivers are extremely important for lymphoma patients
- Make sure patient is comfortable with your involvement
- Respect patient's views and wishes
- Be another set of eyes/ears but not their doctor
- Seek out resources as needed (LLS, Lymphoma Research Foundation, NCI PDQ, etc.)

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SARS-COV-2/COVID-19 IN SLOW GROWING LYMPHOMA PATIENTS—IMPORTANT CONSIDERATIONS

- Slow growing lymphoma patients appear to have worse outcomes with COVID-19 illness
- Systemic therapies likely reduce immunity to clearing infection and responding to vaccination (largely extrapolating data regarding rituximab and influenza vaccination).
- SARS-CoV-2 vaccination is very safe for slow growing lymphoma patients
- SARS-CoV-2 immunity in blood cancer patients is being actively studied (including by the LLS!)

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HOW I ADVISE SLOW GROWING LYMPHOMA PATIENTS ON SARS-COV-2 VACCINATION

- No restriction for treatment naïve patients
- For patients symptomatic from localized/contained disease consider low dose radiation with goal of control for 6-12 months and get vaccinated
- If patient is doing well on maintenance rituximab/obinutuzumab consider holding for 1-3 cycles and administering SARS-CoV-2 vaccine
- Consider holding oral agent for 3-4 months if patient in remission
- Advise against vaccination if recent chemotherapy/immune therapy; wait 4-6 months from last treatment
- Patients should know they may not fully respond to vaccination

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SUMMARY AND WORDS OF ADVICE AND HOPE:

1. There is lots of hope for treatment/"cures" and for new therapies.
2. The devil is in the details; Don't hesitate to seek out help from an expert.
3. We lack ways to prevent/detect lymphomas early (with rare exception) and rare for them to be inherited.
4. We are hopeful that we can cure these diseases in the future.



"Hope and fear cannot occupy the same space.
Invite one to stay"

– Maya Angelou

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Q&A SESSION

Treating Slow-Growing Non-Hodgkin Lymphomas (NHL)

- **Ask a question by phone:**
 - Press star (*) then the number 1 on your keypad.
- **Ask a question by web:**
 - Click "Ask a question"
 - Type your question
 - Click "Submit"

Due to time constraints, we can only take one question per person. Once you've asked your question, the operator will transfer you back into the audience line.

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LLS EDUCATION & SUPPORT RESOURCES

HOW TO CONTACT US:

To contact an **Information Specialist** about disease, treatment and support information, resources and clinical trials:

-  **Call: (800) 955-4572**
Monday to Friday, 9 a.m. to 9 p.m. ET
-  **Chat live online: www.LLS.org/InformationSpecialists**
Monday to Friday, 10 a.m. to 7 p.m. ET
-  **Email: infocenter@LLS.org**
All email messages are answered within one business day.

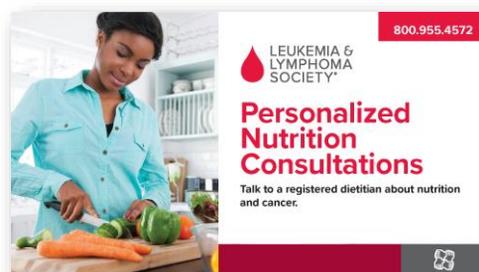
CLINICAL TRIAL SUPPORT CENTER

Work one-on-one with an LLS Clinical Trial Nurse Navigator who will help you find clinical trials and personally assist you throughout the entire clinical-trial process.

www.LLS.org/Navigation



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NUTRITION CONSULTATIONS

Our registered dietitian has expertise in oncology nutrition and provides free one-on-one consultations by phone or email.

www.LLS.org/Consult



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LLS EDUCATION & SUPPORT RESOURCES



Online Chats

Online Chats are free live sessions, **moderated by oncology social workers**. To register for one of the chats below, or for more information, please visit www.LLS.org/Chat.



LLS Online Community

Community of blood cancer patients, survivors and caregivers supporting each other and giving trusted information and resources, please visit www.LLS.org/Community.



Patient Podcast

The Bloodline with LLS is here to remind you that after a diagnosis comes hope. To listen to an episode, please visit www.TheBloodline.org.

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LLS EDUCATION & SUPPORT RESOURCES

877.557.2672

LEUKEMIA & LYMPHOMA SOCIETY™

Help With Finances

The Leukemia & Lymphoma Society (LLS) offers financial assistance* to help individuals with blood cancer:

The **LLS Patient Aid** Program provides financial assistance to blood cancer patients in active treatment. Eligible patients will receive a \$100 stipend. Visit www.LLS.org/PatientAid

The **Urgent Need** Program, established in partnership with Maggie's Lane, helps pediatric and young adult blood cancer patients, or adult blood cancer patients who are enrolled in clinical trials, with acute financial need. The program provides a \$500 grant to assist with non-medical expenses, including utilities, rent, mortgage, food, lodging, dental care, child care, elder care, and other essential needs. Visit www.LLS.org/UrgentNeed

The **Susan Lang Pay-It-Forward Patient Travel Assistance** Program provides blood cancer patients a \$500 grant to assist with transportation and lodging-related expenses. Visit www.LLS.org/Travel

The **Co-Pay Assistance** Program offers financial support toward the cost of insurance co-payments and/or insurance premiums for prescription drugs. Visit www.LLS.org/CoPay

*Funding for LLS Co-pay Assistance Program is made possible through the generous funding from the LLS Financial Assistance program's primary donor, the Susan Lang Foundation, and other donors.

The Leukemia & Lymphoma Society (LLS) offers the following financial assistance programs to help individuals with blood cancer:

www.LLS.org/Finances



To order free materials: www.LLS.org/Booklets

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THANK YOU



We have one goal: A world without blood cancers

