

Practical Management Pearls for Immunotherapy for the Treatment of Acute Leukemia

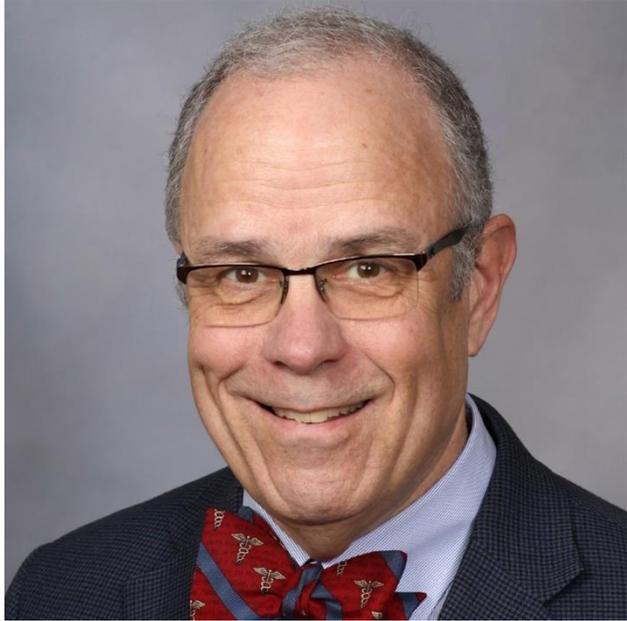
October 14, 2021

11:30 a.m. – 12:30 p.m. ET



The Practical Management Pearls and Case Studies Webinars are part of the Cancer Immunotherapy Clinical Practice Guidelines Advanced Webinar Series supported, in part, by grants from Amgen and Merck & Co., Inc. (as of 9/15/2021)

Webinar faculty



Mark R. Litzow, MD – *Mayo
Clinic Cancer Center*



Daniel A. Arber, MD – *University
of Chicago*

Learning objectives

- Outline practical considerations for diagnostic testing and classification in acute leukemia and the implications for immunotherapy treatment planning
- Appropriately manage challenging and/or uncommon toxicities/irAEs associated with immunotherapy in acute leukemia
- Determine optimal sequencing of immunotherapies in all stages of acute leukemia treatment, including treatment for persistent or relapsed/refractory disease after initial therapy

Outline

- Guideline development
- Diagnostic testing in AL
 - New data in acute leukemia (new classifications)
- Toxicities associated with immunotherapy for acute leukemia
- Sequencing therapies in acute leukemia

Guideline development

Open access **Position article and guidelines**

 Journal for
ImmunoTherapy of Cancer

The Society for Immunotherapy of Cancer (SITC) clinical practice guideline on immunotherapy for the treatment of acute leukemia

Michael M Boyiadzis,¹ Ivan Aksentijevich,² Daniel A Arber,³ John Barrett,⁴
Renier J Brentjens,⁵ Jill Brufsky,¹ Jorge Cortes,⁶ Marcos De Lima,⁷
Stephen J Forman,⁸ Ephraim J Fuchs,⁹ Linda J Fukas,¹⁰ Steven D Gore,¹¹
Mark R Litzow,¹² Jeffrey S Miller,¹³ John M Pagel,¹⁴ Edmund K Waller,¹⁵
Martin S Tallman⁵

Guideline development

- *The Institute of Medicine's Standards for Developing Trustworthy Practice Guidelines* used to develop these recommendations
- Panel consisted of 17 participants, including medical oncologists, hematologists, a hematopathologist, a leukemia research nurse, and a patient advocate
- Recommendations come from literature evidence, supplemented with clinical experience of the panel members where necessary
- Consensus defined as $\geq 75\%$ agreement

Outline

- Guideline development
- **Diagnostic testing in AL**
 - New data in acute leukemia (new classifications)
- Toxicities associated with immunotherapy for acute leukemia
- Sequencing therapies in acute leukemia

Initial Diagnostic Workup of Acute Leukemia

Guideline From the College of American Pathologists and the American Society of Hematology

Daniel A. Arber, MD; Michael J. Borowitz, MD, PhD; Melissa Cessna, MD; Joan Etzell, MD; Kathryn Foucar, MD; Robert P. Hasserjian, MD; J. Douglas Rizzo, MD; Karl Theil, MD; Sa A. Wang, MD; Anthony T. Smith, MLS; R. Bryan Rumble, MSc; Nicole E. Thomas, MPH, CT(ASCP)^{cm}; James W. Vardiman, MD

• **Context.**—A complete diagnosis of acute leukemia requires knowledge of clinical information combined with morphologic evaluation, immunophenotyping and karyotype analysis, and often, molecular genetic testing. Although many aspects of the workup for acute leukemia are well accepted, few guidelines have addressed the different aspects of the diagnostic evaluation of samples from patients suspected to have acute leukemia.

Objective.—To develop a guideline for treating physicians and pathologists involved in the diagnostic and prognostic evaluation of new acute leukemia samples, including acute lymphoblastic leukemia, acute myeloid leukemia, and acute leukemias of ambiguous lineage.

Design.—The College of American Pathologists and the American Society of Hematology convened a panel of

experts in hematology and hematopathology to develop recommendations. A systematic evidence review was conducted to address 6 key questions. Recommendations were derived from strength of evidence, feedback received during the public comment period, and expert panel consensus.

Results.—Twenty-seven guideline statements were established, which ranged from recommendations on what clinical and laboratory information should be available as part of the diagnostic and prognostic evaluation of acute leukemia samples to what types of testing should be performed routinely, with recommendations on where such testing should be performed and how the results should be reported.

Conclusions.—The guideline provides a framework for the multiple steps, including laboratory testing, in the evaluation of acute leukemia samples. Some aspects of the guideline, especially molecular genetic testing in acute leukemia, are rapidly changing with new supportive literature, which will require on-going updates for the guideline to remain relevant.

(*Arch Pathol Lab Med.* 2017;141:1342–1393; doi: 10.5858/arpa.2016-0504-CP)

Accepted for publication December 9, 2016.

Published as an Early Online Release February 22, 2017.

Supplemental digital content is available for this article at www.archivesofpathology.org in the October 2017 table of contents.

From the Department of Pathology, University of Chicago,

ASCO special article

Initial Diagnostic Work-Up of Acute Leukemia: ASCO Clinical Practice Guideline Endorsement of the College of American Pathologists and American Society of Hematology Guideline

Valérie de Haas, PhD¹; Nofisat Ismaila, MD²; Anjali Advani, MD³; Daniel A. Arber, MD⁴; Raetasha S. Dabney, MD⁵; Dipti Patel-Donnelly, MD⁶; Elizabeth Kittlas, LMSW⁷; Rob Pieters, MD, PhD¹; Ching-Hon Pui, MD⁸; Kendra Sweet, MD⁹; and Ling Zhang, MD⁹

abstract

PURPOSE The College of American Pathologists (CAP) and the American Society of Hematology (ASH) developed an evidence-based guideline on the initial diagnostic work-up of acute leukemia (AL). Because of the relevance of this topic to the ASCO membership, ASCO reviewed the guideline and applied a set of procedures and policies for endorsing clinical practice guidelines that have been developed by other professional organizations.

METHODS The CAP-ASH guideline on initial diagnostic work-up of AL was reviewed for developmental rigor by methodologists. Then, an ASCO Endorsement Expert Panel updated the literature search and reviewed the content and recommendations.

RESULTS The ASCO Expert Panel determined that the recommendations from the guideline, published in 2016, are clear, thorough, and based on the most relevant scientific evidence. ASCO fully endorsed the CAP-ASH guideline on initial diagnostic work-up of AL and included some discussion points according to clinical practice and updated literature.

CONCLUSION Twenty-seven guideline statements were reviewed. Some discussion points were included to better assess CNS involvement in leukemia and to provide novel insights into molecular diagnosis and potential markers for risk stratification and target therapy. These discussions are categorized into four sections: (1) initial diagnosis focusing on basic diagnostics and determination of risk parameters, (2) molecular markers and minimal residual disease detection, (3) context of referral to another institution with expertise in the management of AL, and (4) reporting and record keeping for better outlining and follow-up discussion. Additional information is available at: www.asco.org/hematologic-malignancies-guidelines.

CAP/ASH Guideline

Expert Panel	Advisory Panel	CAP/ASH Staff
Daniel Arber, MD, Co-Chair, CAP	Frederick R. Appelbaum, MD	Nicole Thomas, MPH, CT(ASCP), CAP
James Vardiman, MD, Co-Chair, ASH	Clara Bloomfield, MD	Robert Plovnick, MD, ASH
Michael Borowitz, MD, PhD, ASH	William L. Carroll, MD	Robert Kunkle, ASH
Melissa Cessna, MD, CAP	Laura Housley, Patient Advocate	Kendall Alexander, MPH, ASH
Joan Etzell, MD, CAP	Jerry Hussong, MD	R. Bryan Rumble, MSc, methodologist
Kathryn Foucar, MD, ASH	Steven H. Kroft, MD, FASCP	Christina Lacchetti, MHSc, methodologist
Robert Hasserjian, MD, ASH	Michelle Le Beau, PhD	Tony Smith, MLS, CAP, Medical Librarian
J. Douglas Rizzo, MD, ASH	Martin S. Tallman, MD	
Karl Theil, MD, CAP		
Sa Wang, MD, CAP		

CAP/ASH Guideline

- Developed key questions
- Performed baseline survey of pathologists and hematologists (Arch Pathol Lab Med (2017) 141(8):1101-1106)
- Performed meta-analysis
- Public comment period
- Resulted in 27 guideline statements regarding the initial workup of acute leukemia (Arch Pathol Lab Med (2017) 141(10):1342–1393)

CAP/ASH Guideline

- Wide variety of recommendations, including
 - Immunophenotyping
 - Genetic and molecular genetic testing
 - Classification

Immunophenotyping

Table 2 Recommended immunotherapy-centric diagnostic markers for acute leukemia

Disease type	Marker	Agents for consideration
Acute lymphoblastic leukemia	CD19	Blinatumomab
	CD19	Tisagenlecleucel (patients aged ≤ 25 years)
	CD22	Inotuzumab ozogamicin
	CD20	Rituximab
Acute myeloid leukemia	CD33	Gemtuzumab ozogamicin

Immunophenotyping – Baseline Survey

	AML Response Percent	AML Response Count	ALL Response Percent	ALL Response Count
Morphologic assessment	100.0%	234	99.1%	232
Flow cytometric analysis	99.1%	232	98.3%	229
Conventional cytogenetics (karyotype)	96.2%	225	96.6%	225
Molecular testing	78.2%	183	54.9%	128

Genetic and Molecular Genetic Testing (Karyotype; Mutational studies)

	AML Response Percent	AML Response Count	ALL Response Percent	ALL Response Count
Morphologic assessment	100.0%	234	99.1%	232
Flow cytometric analysis	99.1%	232	98.3%	229
Conventional cytogenetics (karyotype)	96.2%	225	96.6%	225
Molecular testing	78.2%	183	54.9%	128

Genetic and Molecular Genetic Testing (Karyotype; Mutational studies)

Statement 15 Other ALL Testing

- B-ALL
 - Mutations panels that may include, but are not limited to
 - *PAX5*, *JAK1*, *JAK2*, and/or *IKZF1* R
 - Expression of CRLF2 R
- T-ALL
 - Mutations panels that may include, but are not limited to
 - *NOTCH1* and/or *FBXW7* R

R, recommendation

Genetic and Molecular Genetic Testing (Karyotype; Mutational studies)

Statements 16, 17, 19 Adult and Pediatric AML

- *FLT3*-ITD SR – all types
- *NPM1*, *CEBPA*, *RUNX1* SR – most types
- *KIT* SR (adult), R (peds) – Core binding factor AML
- Mutations panels that may include, but are not limited to
 - *IDH1*, *IDH2*, *TET2*, *WT1*, *DNMT3A*, and/or *TP53* R
 - (*ASXL1*)

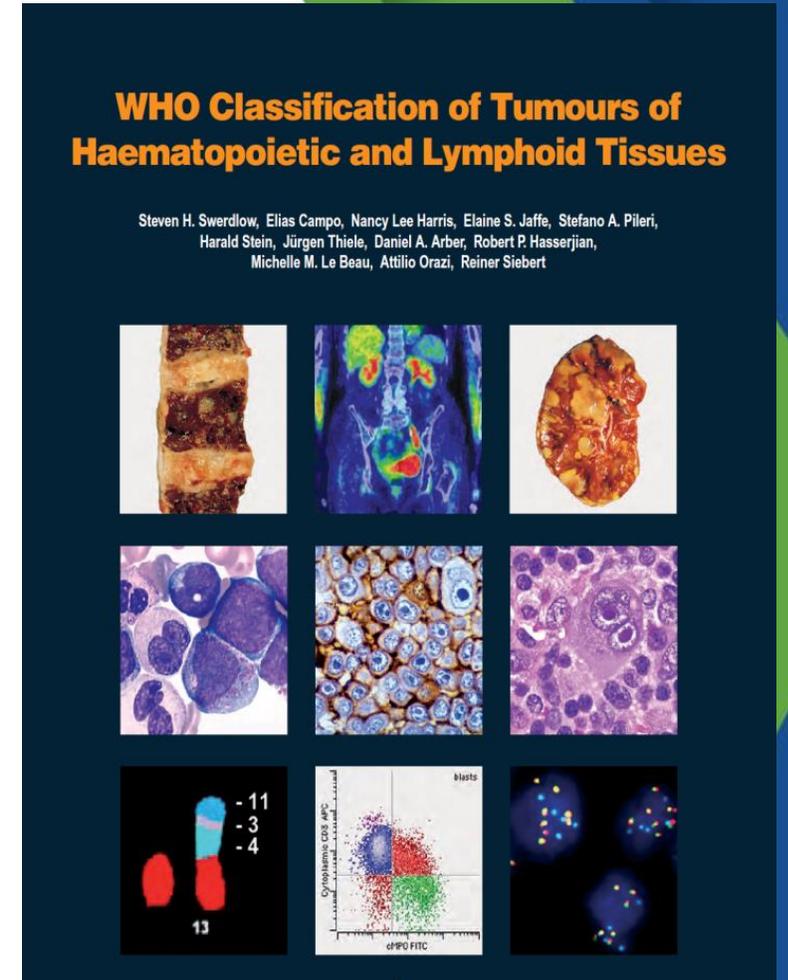
SR, strong recommendation; R, recommendation

Diagnostic tests for ALL/AML

ASCO/CAP/ASH recommended genetic tests (Adopted by SITC Guideline)	
Acute lymphoblastic leukemia	Acute myeloid leukemia
<i>t(9;22)(q34.1;q11.2); BCR-ABL1</i>	<i>FLT3-ITD</i>
<i>PAX5; CRLF2; JAK1; JAK2</i>	<i>NPM1; CEBPA; RUNX1</i>
CRLF2 overexpression (for B-ALL)	<i>IDH1; IDH2; TET2; WT1; DNMT3A; TP53</i>
<i>KMT2A (MLL); IKZF1</i> (for B-ALL)	<i>KIT</i> (for CBF AML)
<i>NOTCH1</i> and/or <i>FBXW7</i> (for T-ALL)	<i>RUNX1-RUNX1T1/CBFB-MYH1</i> (for CBF AML)
	<i>PML-RARA</i> for suspected APL

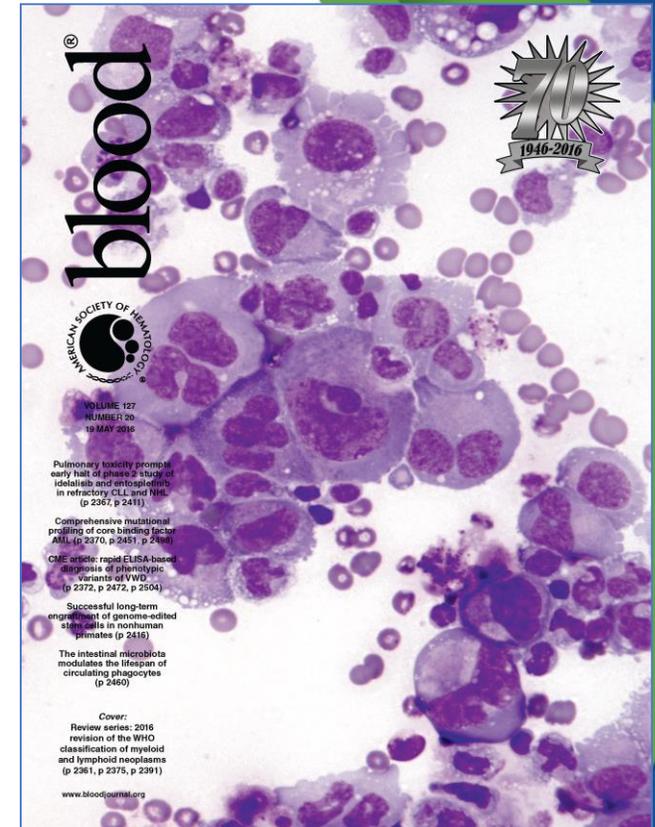
Classification

- Current Recommendation
 - Use the current WHO classification diagnostic terminology
- Future Direction
 - 5th edition WHO classification in the works with IARC
 - Expected publication date of late 2022
 - Have changed their process to not incorporate a clinical advisory committee, to exclude prior editors and to refuse input from hematopathology societies



Classification

- Current Recommendation
 - Use the current WHO classification terminology diagnostic terminology
- Future Direction
 - 5th edition WHO classification in the works with IARC
 - Expected publication date of late 2022
 - Have changed their process to not incorporate a clinical advisory committee, to exclude prior editors and to refuse input from hematopathology societies
 - Separate International Consensus Classification of Myeloid and Lymphoid Neoplasms
 - Clinical Advisory Committee meeting held in September 2021
 - Expected publication in 2022
 - Update of CAP/ASH guidelines
 - Planned to occur after new classifications are published

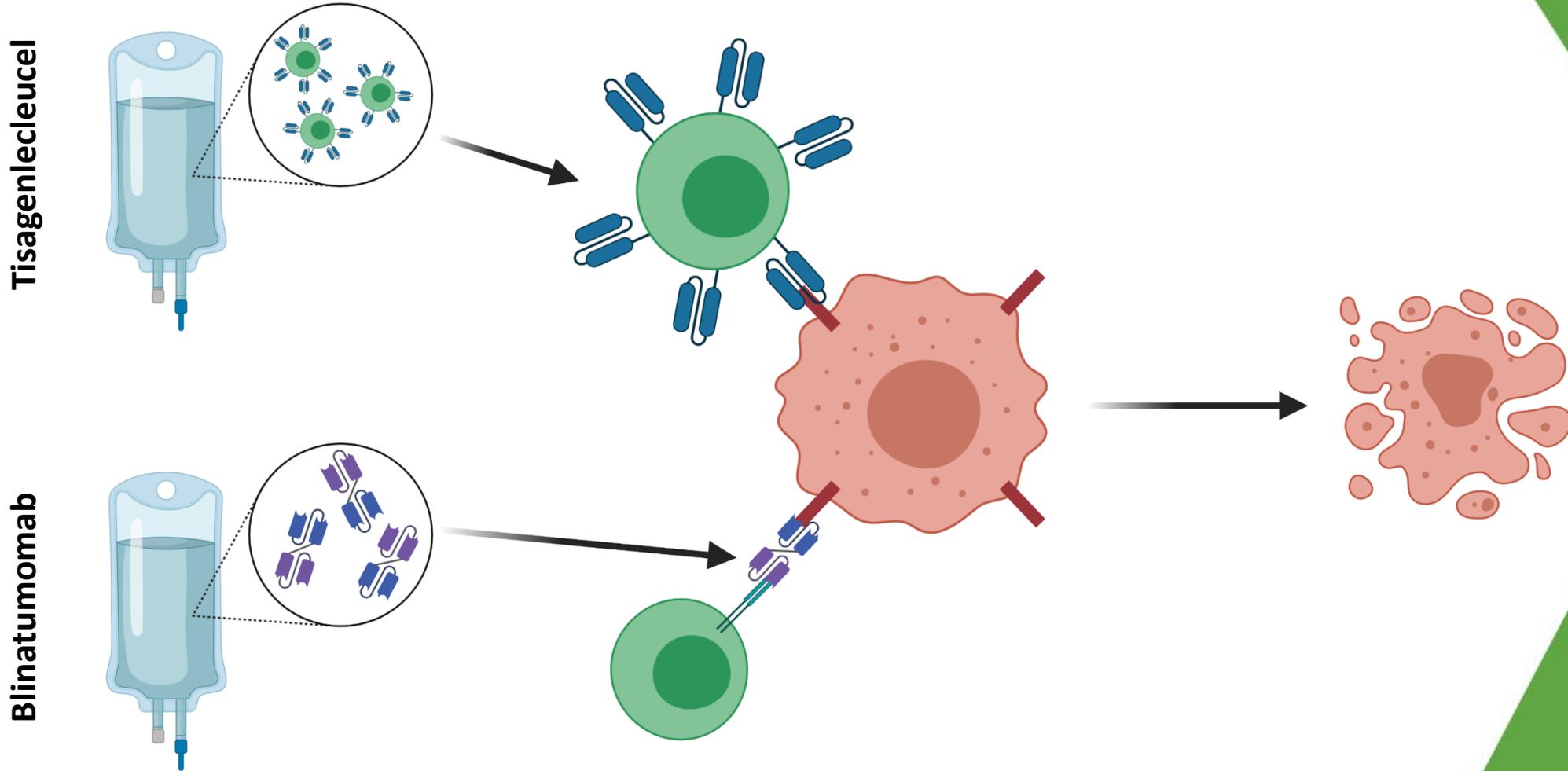


Blood 127:2391, 2016

Outline

- Diagnostic testing in AL
- **Toxicities associated with immunotherapy for acute leukemia**
- Sequencing therapies in acute leukemia

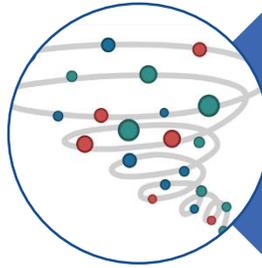
T cell engagers vs. CAR T therapy



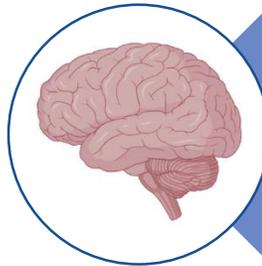
T cell engagers vs. CAR T therapy

	CAR T cells	T cell engagers
Structure	Synthetic gene construct encoding an scFv against tumor antigen linked to activation/costimulatory motifs	Recombinant protein with two specificities: one for tumor antigen and one for T cell antigen (usually CD3)
Effector cell types	Engineered CD8+ and CD4+ T cells	Endogenous CD8+ and CD4+ T cells
Immune synapse	Atypical	Typical
Serial killing	Yes	Yes
Killing mechanisms	Perforin and granzyme B, Fas-Fas-L, or TNF/TNF-R	Perforin and granzyme B
Trafficking	Active	Passive
Clinical applications	Pre-treatment lymphodepletion followed by a single infusion	No lymphodepletion; repeat administration and continuous infusions
Specificity	Manufactured for each patient	“Off-the-shelf”
Availability	Limited to REMS program facilities	Most cancer centers

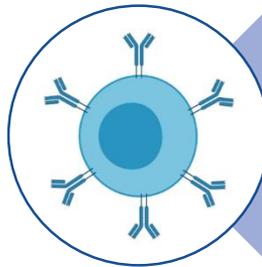
Common toxicities with tisagenlecleucel and blinatumomab



Cytokine release
syndrome

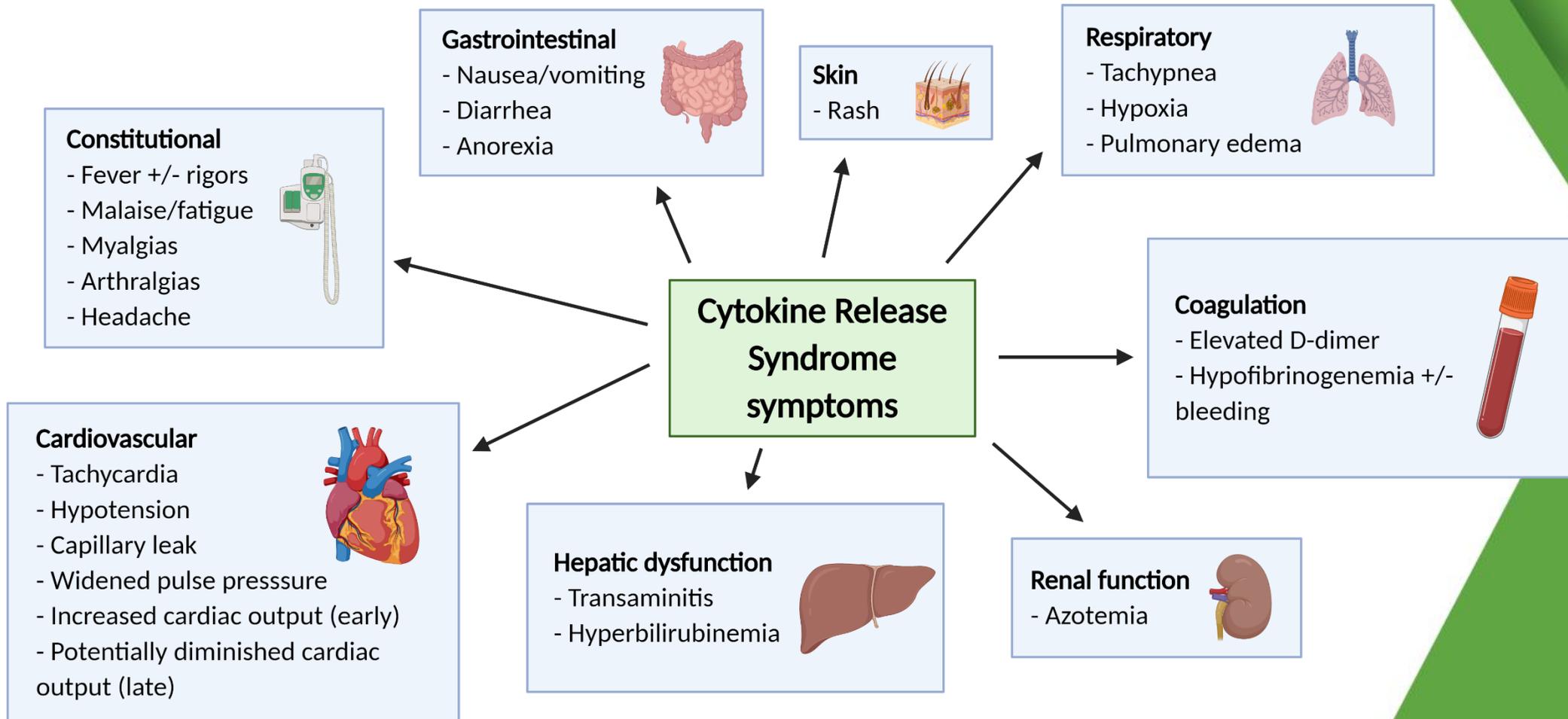


Neurotoxicity



B cell aplasia

Cytokine release syndrome



ASTCT CRS grading

CRS parameter	Grade 1	Grade 2	Grade 3	Grade 4
Fever	≥ 38°C	≥ 38°C	≥ 38°C	≥ 38°C
with				
Hypotension	None	Not requiring vasopressors	Requiring a vasopressor with or without vasopressin	Requiring multiple vasopressors (excluding vasopressin)
and/or				
Hypoxia	None	Requiring low-flow nasal cannula or blow-by	Requiring high-flow nasal cannula, face mask, non-rebreather mask or venturi mask	Requiring positive pressure (e.g. CPAP, BiPAP, intubation and mechanical ventilation)

Monitoring for CRS

- Events requiring physician notification include:
 - Deviations from baseline systolic blood pressure
 - Heart rate >120 or <60 bpm
 - Arrhythmia
 - Respiratory rate >25 or <12 breaths/minute
 - Arterial oxygen saturation <92% on room air
 - Upward trend in blood creatinine or liver function tests
 - Tremors or jerky movements in extremities
 - Altered mental status
 - Temperature $\geq 38^{\circ}\text{C}$

Management of CRS with CAR T

Grade 1	Grade 2	Grade 3	Grade 4	Tocilizumab-unresponsive	Tocilizumab + steroids-unresponsive
Close monitoring and supportive care	Consider tocilizumab	Tocilizumab	Tocilizumab + steroids	If CRS does not respond to 1 dose of tocilizumab, combine steroids + tocilizumab	Options include: Anakinra, siltuximab, HD methylprednisone

- For **elderly patients or those with significant co-morbidities**, tocilizumab should be considered earlier in the treatment course.
- If CRS does not improve after tocilizumab + steroids, **infections** should be considered and managed appropriately.
- If steroids are used, a **rapid taper** should be employed once symptoms begin to improve.

Management of CRS with blinatumomab

CRS grade	Patients weighing 45 kg or more	Patients weighing less than 45 kg
3	<ul style="list-style-type: none">• Interrupt blinatumomab.• Administer dexamethasone 8 mg Q8h intravenously or orally for up to 3 days and taper thereafter over 4 days• When CRS is resolved, restart blinatumomab at 9 mcg/d, and escalate to 28 mcg/d after 7 days if CRS does not recur.	<ul style="list-style-type: none">• Interrupt blinatumomab.• Administer dexamethasone 5 mg/m² Q8h intravenously or orally for up to 3 days and taper thereafter over 4 days• When CRS is resolved, restart blinatumomab at 5 mcg/m²/d, and escalate to 15 mcg/m²/d after 7 days if CRS does not recur.
4	Discontinue blinatumomab permanently. Administer dexamethasone as instructed for Grade 3 CRS.	

ASTCT ICANS grading - adults

ICANS: Immune cell associated neurotoxicity syndrome

Neurotoxicity domain	Grade 1	Grade 2	Grade 3	Grade 4
ICE score	7–9	3–6	0–2	0 (patient is unarousable)
Depressed level of consciousness	Awakens spontaneously	Awakens to voice	Awakens only to tactile stimulus	Patient is unarousable or requires vigorous or repetitive tactile stimuli to arouse; stupor or coma
Seizure	N/A	N/A	Any clinical seizure focal or generalized that resolves rapidly or non-convulsive seizures on EEG that resolve with intervention	Life-threatening prolonged seizure (>5 min), repetitive clinical or electrical seizures without return to baseline in between
Motor findings	N/A	N/A	N/A	Deep focal motor weakness such as hemiparesis or paraparesis
Elevated ICP/cerebral edema	N/A	N/A	Focal/local edema on neuroimaging	Diffuse cerebral edema on neuroimaging, decerebrate or decorticate posturing, cranial nerve VI palsy, papilledema, or Cushing's triad

ASTCT ICANS grading - pediatric

Neurotoxicity domain	Grade 1	Grade 2	Grade 3	Grade 4
ICE score (age ≥12 years)	7–9	3–6	0–2	0 (patient is unarousable)
CAPD score (age <12 years)	1–8	1–8	≥9	Unable to perform CAPD
Depressed level of consciousness	Awakens spontaneously	Awakens to voice	Awakens only to tactile stimulus	Unarousable or requires vigorous or repetitive tactile stimuli to arouse
Seizure (any age)	N/A	N/A	Any clinical seizure focal or generalized that resolves rapidly or non-convulsive seizures on EEG that resolve with intervention	Life-threatening prolonged seizure (>5 min), repetitive clinical or electrical seizures without return to baseline in between
Motor weakness (any age)	N/A	N/A	N/A	Deep focal motor weakness such as hemiparesis or paraparesis
Elevated ICP/cerebral edema (any age)	N/A	N/A	Focal/local edema on neuroimaging	Diffuse cerebral edema on neuroimaging; decerebrate or decorticate posturing, cranial nerve VI palsy, papilledema, or Cushing's triad

Immune effector cell-associated encephalopathy (ICE) score

- **Orientation:** Orientation to year, month, city, hospital: 4 points (1 point each)
- **Naming:** Name 3 objects (e.g., clock, pen, button): 3 points (1 point each)
- **Following commands:** (e.g., Show me 2 fingers or close your eyes and stick out your tongue): 1 point
- **Writing:** Ability to write a standard sentence (e.g., Our national bird is the bald eagle): 1 point
- **Attention:** Count backwards from 100 by 10: 1 point
- **Total scale:** 0-10

Monitoring for ICANS

- Altered mental status defines the onset of ICANS
- Work-up should include:
 - CRP
 - CBC
 - CMP
 - Fibrinogen
 - Prothrombin time test
 - PT/INR
- Head CT, EEG, and brain MRI may be considered

Management of ICANS with CAR T

- **4-1BB** CAR T agents: consider steroids at grade 2 ICANS; administer steroids for grades 3-4 ICANS
- Management of neurotoxicity **may take precedence** over low-grade CRS, due to possibility of tocilizumab worsening ICANS
 - For example: in the case of a patient with concomitant grade 1 CRS (fever) and grade 2 ICANS, steroids should be given. This does not apply to higher-grade CRS.
- If **steroids** are used, administer at least two doses and employ a fast taper
- **Levetiracetam** is recommended for management of seizures

Management of ICANS with blinatumomab

ICANS grade	Patients weighing more than 45 kg	Patients weighing less than 45 kg
3	Withhold blinatumomab until no more than Grade 1 and for at least 3 days, then restart blinatumomab at 9 mcg/d. Escalate to 28 mcg/d after 7 days if symptoms do not recur. If ICANS occurred at 9 mcg/d or takes more than 7 days to resolve, discontinue permanently.	Withhold blinatumomab until no more than Grade 1 and for at least 3 days, then restart blinatumomab at 5 mcg/m ² /d. Escalate to 15 mcg/m ² /d after 7 days if symptoms do not recur. If ICANS occurred at 5 mcg/m ² /d or takes more than 7 days to resolve, discontinue permanently.
4	Discontinue blinatumomab permanently.	

B cell aplasia

- Due to current clinical agents targeting CD19, which is expressed by both normal and neoplastic B cells
- Short- or long-term
- May result in hypogammaglobulinemia
- Increased risk of infection – prophylaxis required
- Managed through administration of intravenous immunoglobulin
- Might indicate persistence of CAR T cells

Outline

- Diagnostic testing in AL
- Toxicities associated with immunotherapy for acute leukemia
- Sequencing therapies in acute leukemia

First Results of an Open Label Phase II Study to Evaluate the Efficacy and Safety of Inotuzumab Ozogamicin for Induction Therapy followed by a Conventional Chemotherapy Based Consolidation and Maintenance Therapy in Patients Aged 56 Years and Older with Acute Lymphoblastic Leukemia (INITIAL-1 trial)

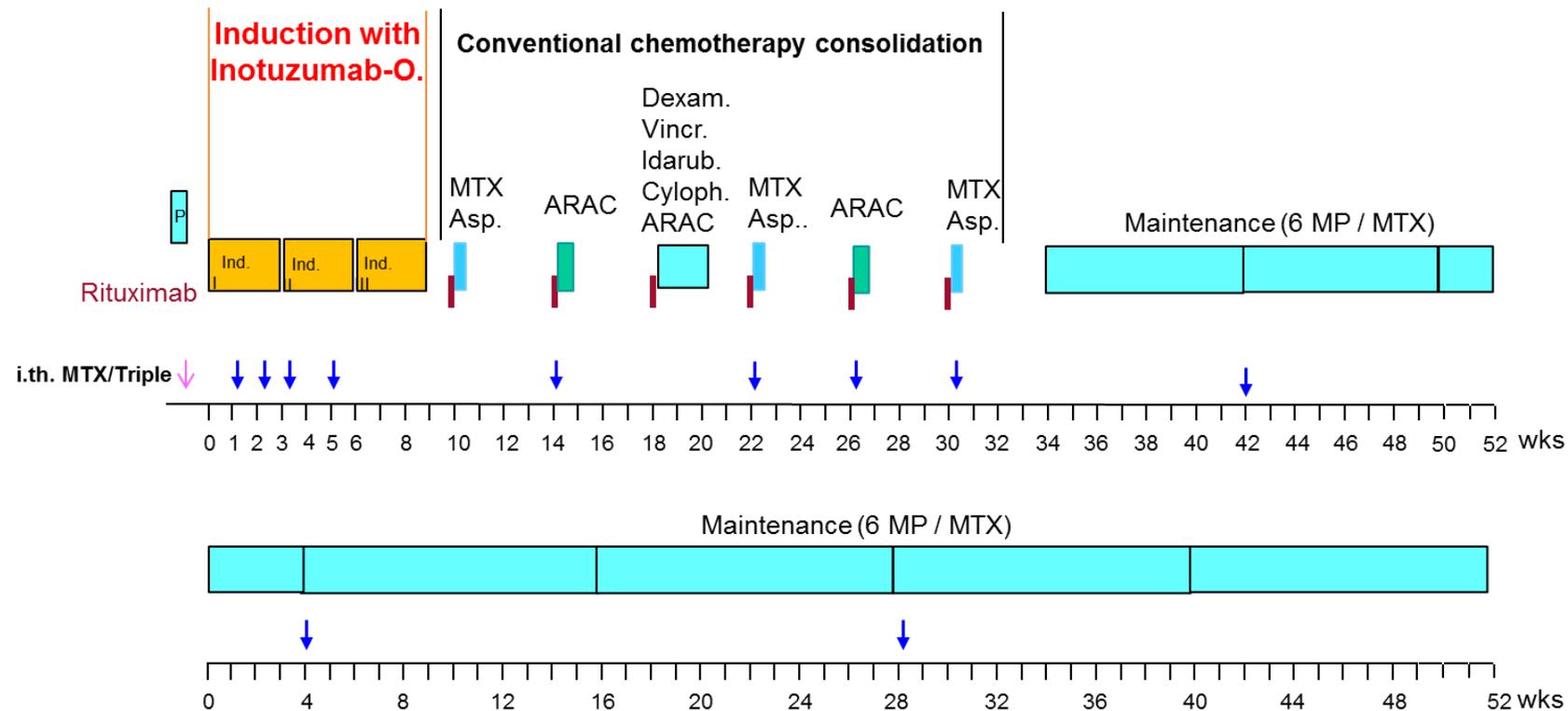
Matthias Stelljes

University of Muenster / Germany

for the GMALL study group

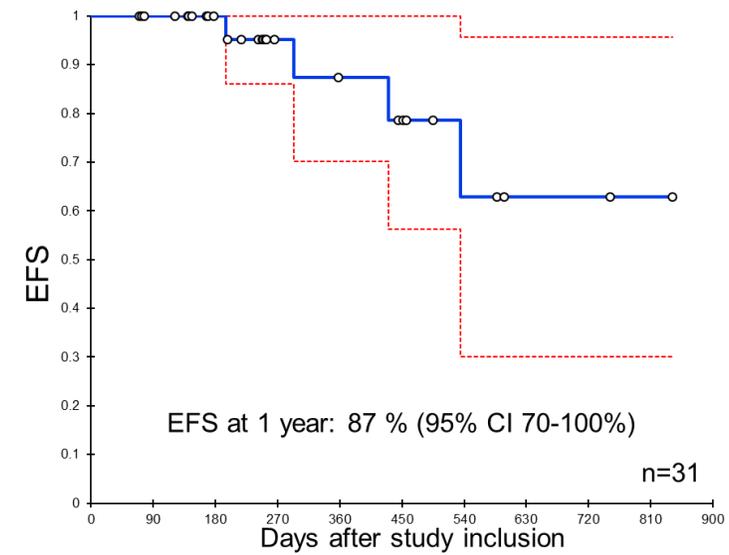
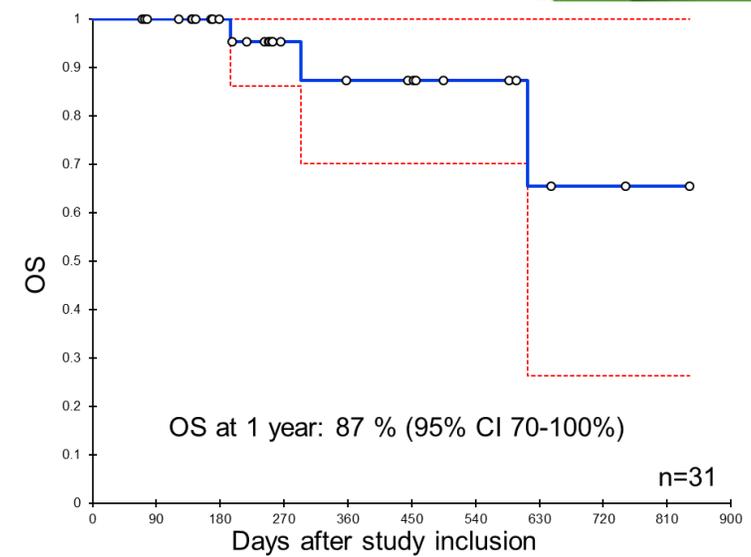
INITIAL-1 Trial: Treatment Schedule

An open label phase II study to evaluate the efficacy and safety of **I**notuzumab Ozogamicin for **I**nduction **T**herapy followed by a conventional chemotherapy-based consolidation and maintenance therapy **I**n patients aged 55 years and older with **A**cute **L**ymphoblastic leukemia (ALL)



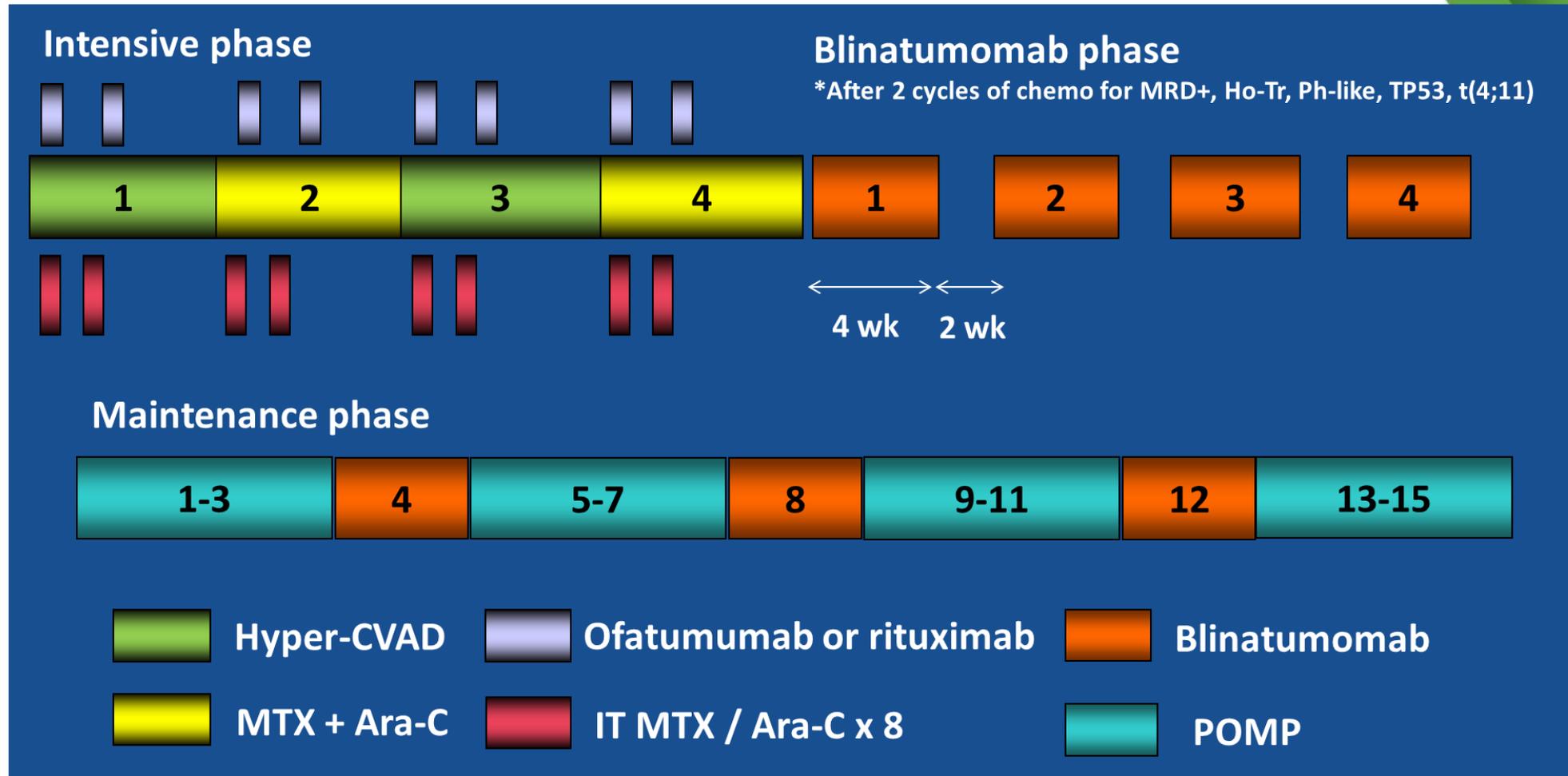
INITIAL-1 Trial: Results

Evaluable for hematological remission, n=31	
CR / CRi after at least 1 induction	31 pts (100%)
Patients receiving 3 cycles inotuzumab	29 pts (94%)
Early deaths within the first 3 months	0
Evaluable for MRD (by PCR), n=27	
MRD negative remission as best response	21 pts (78%)
Hematological / molecular relapse	2 / 1 pts
Allogeneic HSCT in remission / after relapse	3 / 1 pts



Median FU: 249 (70-842) days

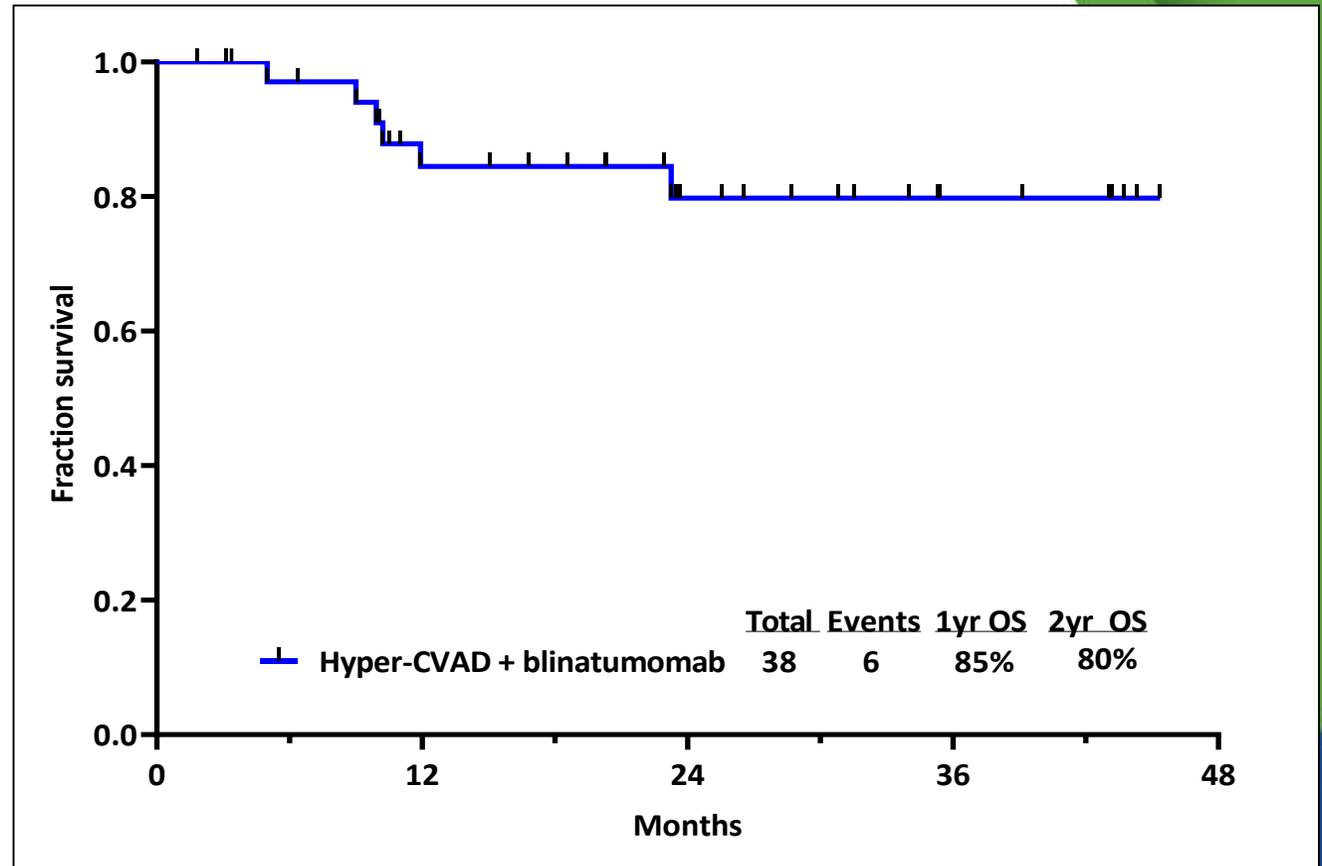
Hyper-CVAD + blinatumomab in newly-diagnosed B-ALL



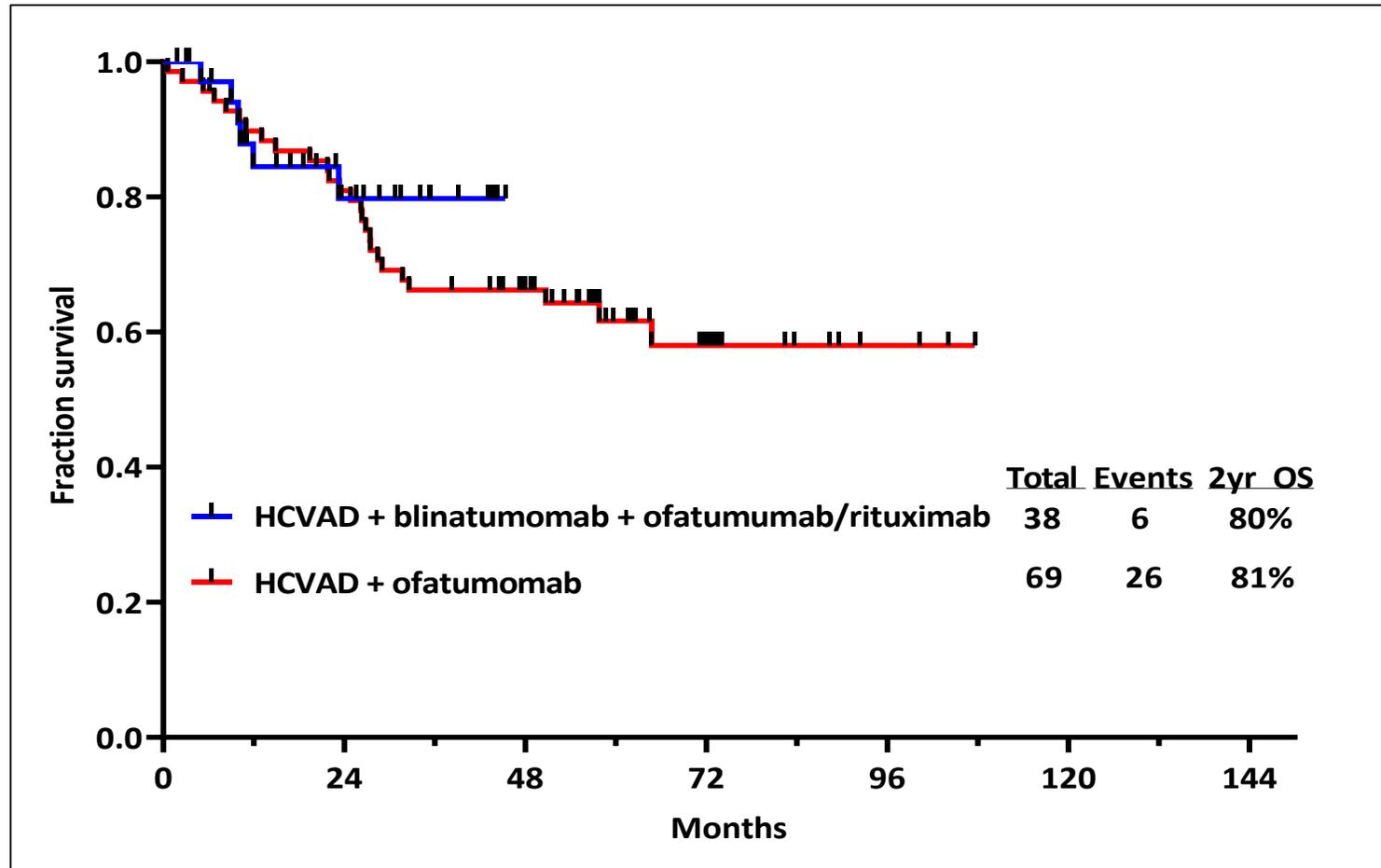
Hyper-CVAD + blinatumomab in newly-diagnosed B-ALL

Response*	n/N (%)
CR after induction	26/32 (81)
CR at any time	32/32 (100)
MRD negativity after induction	24/34 (71)
MRD negativity at anytime	33/34 (97)
30-day mortality	0

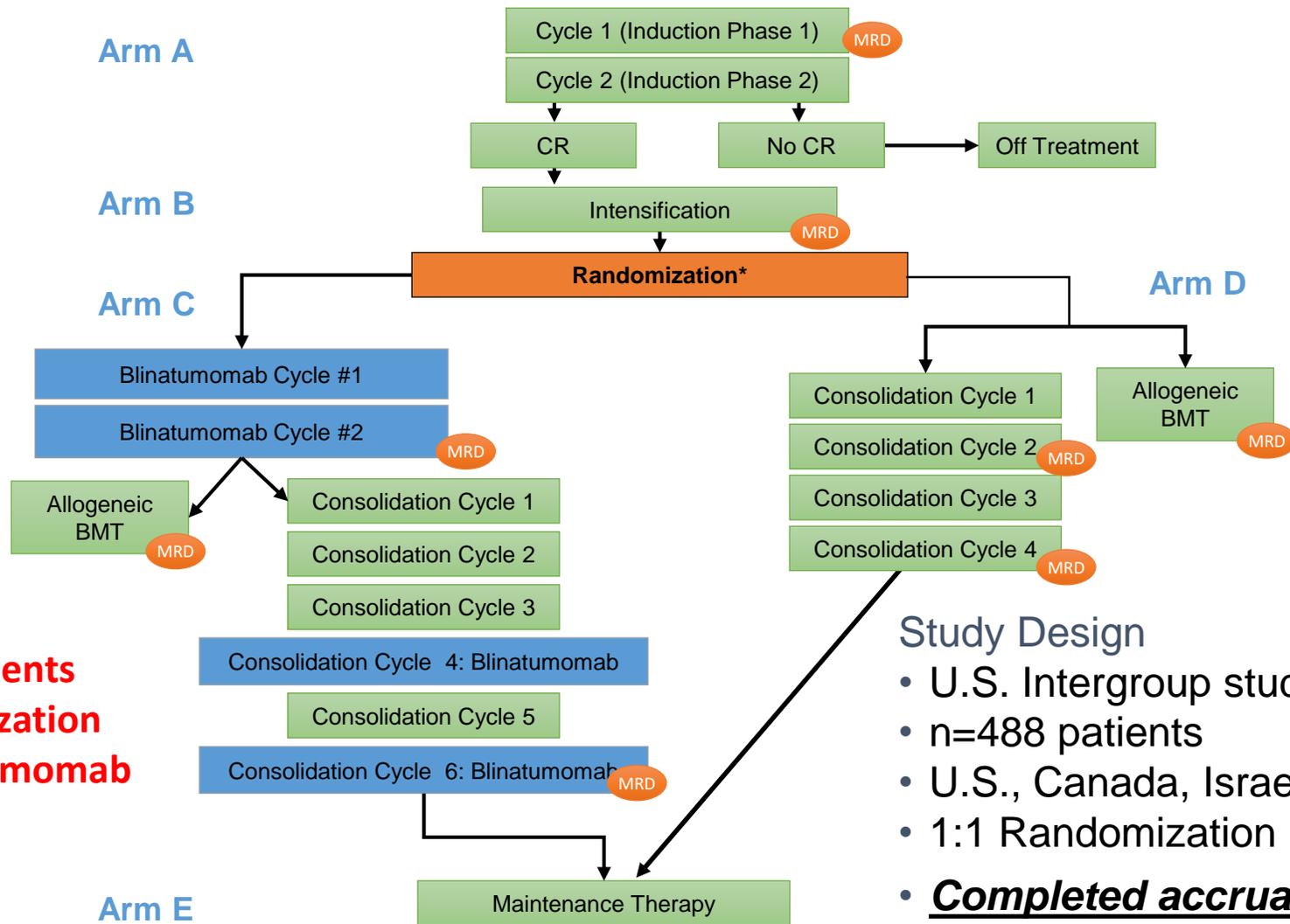
* 6 pts in CR at start; 4 pts MRD negative at start



Hyper-CVAD + blinatumomab in B-ALL: Historical comparison



E1910: Randomized Ph III Adult Frontline ALL



***MRD positive patients at time of randomization assigned to blinatumomab**

- Study Design**
- U.S. Intergroup study
 - n=488 patients
 - U.S., Canada, Israel
 - 1:1 Randomization
 - **Completed accrual October, 2019**

Approved immunotherapies for ALL

Drug	Type	Mechanism	Approval	Indications
Blinatumomab	Bispecific T cell engager (BiTE)	CD3 x CD19 bispecific	March 2018	Adult and pediatric patients with B-cell precursor ALL in first or second complete remission with MRD $\geq 0.1\%$
			July 2017	Relapsed or refractory B-cell precursor ALL in adults and children
			December 2014	Philadelphia chromosome-negative relapsed or refractory B-cell precursor ALL
Inotuzumab ozogamicin	Anti-CD22 antibody–drug conjugate	Antibody-drug conjugate, CD22 antibody + calicheamicin	August 2017	Adults with relapsed or refractory B-cell precursor ALL
Tisagenlecleucel	CAR T cell therapy	CD19 CAR T cells	August 2017	Patients up to 25 years of age with B-cell precursor ALL that is refractory or in second or later relapse

New CAR-T cell therapy just approved

- Brexucabtagene autoleucel (BA) approved on October 1 for adult patients with relapsed/refractory B cell precursor ALL
- Similar structure to axicabtagene ciloleucel, the approved CAR-T product for NHL (somewhat different manufacturing)
- Retroviral vector vs lentiviral vector for tisagenlecleucel (T)
- Signaling domain CD28z for BA, 41BBz for (T)
- In Zuma-3 phase 2 trial CR rate by 3 months 28/54 (52%); 97% MRD negative
- \geq grade 3 CRS and neurotoxicity 24 and 25%, respectively

Role of HCT and immunotherapy

- There is some concern of increased GVHD with combination of checkpoint inhibitors and HCT
- ELIANA and TOWER trials treated patients with prior allo-HCT, apparently safely
- In one study of CD19 CAR T treatment:
 - 17 patients with CR after CAR T proceeded to transplantation
 - 5 were alive with a complete remission
 - 6 had a relapse
 - 6 died from transplant-related toxic effects

Role of immunotherapy post-CAR T

- Patients may relapse after CAR T treatment due to lack or loss of target antigen, or other reasons.
- Potential treatment options include CAR T therapy targeting different antigens, blinatumomab, allo-HCT or salvage chemotherapy.
- Clinical trial enrollment should be strongly considered.

Thank you!

Learn more and register at:

<https://www.sitcancer.org/CPG-webinars>

Case Studies in Immunotherapy for the Treatment of Acute Leukemia

November 15, 2021, 11:30 a.m. – 12:30 p.m. ET

Immunotherapy for the Treatment of Breast Cancer

October 29, 2021, 1 – 2 p.m. ET

Case Studies in Immunotherapy for the Treatment of Urothelial Cancer

November 5, 2021, 5:30 – 6:30 p.m. ET

Targets for Cancer Immunotherapy: A Deep Dive Seminar Series

Eight online seminars will address key questions in the field of cancer immunotherapy **drug development**

SEMINAR 6: THE 4-1BB PATHWAY – October 21, 2021, 3:30 - 5:30 p.m. ET

SEMINAR 7: T CELL FUNCTIONAL STATES –
November 18, 2021, 4:30 – 6:30 p.m. ET

Learn more and register at:

<https://www.sitcancer.org/education/deepdive>



A Focus on Genitourinary Cancers

October 27, 2021, 12 – 4 p.m. ET

CME-, CPE-, CNE-, MOC-certified

Learn more and register at:

<https://www.sitcancer.org/education/aci>



Journal for ImmunoTherapy of Cancer

Earn CME Credit as a *JITC* Reviewer

JITC also cooperates with reviewer recognition services (such as Publons) to confirm participation without compromising reviewer anonymity or journal peer review processes, giving reviewers the ability to safely share their involvement in the journal.

Learn how to become a reviewer at
sitcancer.org/jitc



Society for Immunotherapy of Cancer

Cancer Immunotherapy Clinical Practice Guidelines Mobile App

sitcancer.org/CPG-app

   #SITCGuidelines



Thank you for attending the webinar!

Questions or comments: connectED@sitcancer.org