

# משוב הסדנה

- סרקו את הברקוד וענו על השאלות
- תודה על המשוב!



# Cases

Nov 2025

Ron Ram

# Case 1

- A 61-year-old with primary-refractory LBCL, LDH 2×ULN, bulky 9 cm mass, prior R-CHOP. PET shows no CNS uptake; CSF negative but high CNS-IPI. Which initial plan best aligns with current practice?
  - A. Immediate leukapheresis → no bridging
  - B. Leukopheresis → polatuzumab-BR bridging to control bulk
  - C. Pola-BR for 2 cycles, then apheresis if PR
  - D. Involved-site RT only → apheresis

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# Case 1

- Assuming commercial CAR-T access to axi-cel, liso-cel, tisa-cel, which choice best balances efficacy for high-risk disease with neurotoxicity awareness per recent real-world data?
  - A. Axi-cel (higher efficacy signal; higher ICANS risk)
  - B. Liso-cel (lower efficacy; lowest ICANS)
  - C. Tisa-cel (best PFS)
  - D. Any—no signal differences exist

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# Case 1

- Day +5: Grade 2 CRS; ICE 10/10. Best next step?
  - A. Start dexamethasone 10 mg q6h immediately
  - B. Tocilizumab now; hold steroids unless ICANS
  - C. Anakinra first-line for CRS
  - D. Observe only

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# Case 1

- Day +12: afebrile, ferritin 12,000, rising AST/ALT, coagulopathy, cytopenias, minimal/no CRS—concern for IEC-HS. Best initial management?
  - A. High-dose IVIG alone
  - B. IL-1 blockade
  - C. Repeat tocilizumab doses
  - D. High-dose steroids and ruxolitinib
  - E. Siltuximab if fibrinogen < 150

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## Case 2

A 68-year-old MCL progressed after RCHOP/RDHAP and then covalent BTKi. What strategy aligns with current practice?

- A. Allo-HCT upfront; CAR-T only if relapse post-HCT
- B. Venetoclax bridge to allo-HCT; avoid CAR-T at 68
- C. Bispecific preferable; avoid CAR-T if bulky disease
- D. Brexu-cel as preferred next step; allo-HCT reserved for post-CAR-T failure/selected fit pts

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# Case 2

- Pre-infusion disease control approach?
  - A. Multi-agent cytotoxic chemo to CR
  - B. Venetoclax +/- BTKi
  - C. Prolonged steroids
  - D. BTKi re-challenge

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# Case 2

- Received brexu-cel, achieved CR, and now relapses 9 months later with nodal and marrow involvement. ECOG 1, 10/10 MUD available. Which next management step is most appropriate according to recent data?
  - A. Proceed to allo-HCT if PR/CR achieved after short bridging
  - B. Repeat CAR-T infusion with liso-cel
  - C. Glofitamab for remission induction followed by maintenance rituximab only
  - D. R-DHAP followed by auto-HCT consolidation

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# Case 3

- A 72-year-old man with AML, adverse cytogenetics, achieves CR1 after 6 cycles of VEN-AZA, but remains MRD+. He has a 10/10 unrelated donor and HCT-CI score of 3. ECOG 0. Which approach most closely reflects current practice for patients like this?
  - A. Defer transplant and continue azacitidine + venetoclax until MRD negativity
  - B. Proceed with MAC regimen to overcome MRD positivity
  - C. Use RIC
  - D. Avoid transplant due to HCTCI > 2
  - E. Autologous HCT after salvage chemotherapy

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# Case 3

- 2 potential donors:

A 10/10 matched unrelated donor (MUD) offering a BM graft

A haploidentical son donor

Which donor is preferable?

- A. 10/10 MUD BM graft with only CNI as prophylaxis (no MTX)
- B. Cord-blood transplant as safest option in the elderly
- C. Haploidentical PBSC graft to maximize GVL
- D. 10/10 MUD BM graft with PTCy-based GVHD prophylaxis

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# Case 3

- Day +35: new hypertension, LDH 2× ULN, creatinine rising, proteinuria 1.2 mg/mg, schistocytes on smear. What is the most appropriate next step?
  - A. Observe—likely calcineurin toxicity – self limited
  - B. Start plasmapheresis immediately with maintenance rituximab
  - C. Stop CNI and initiate complement-directed therapy if high-risk
  - D. Begin high-dose steroids
  - E. Start ruxolitinib

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# Case 4

- A 55-year-old with steroid-refractory cGVHD (skin/fascia + mouth), prior calcineurin inhibitor. Next best systemic option?
  - A. Belumosudil or ruxolitinib
  - B. Acalabrutinib
  - C. Photopheresis mandatory before meds
  - D. High-dose cyclophosphamide

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# Case 4

- A 62-year-old man with severe cGVHD (skin + fascia) starts ruxolitinib 10 mg BID. After 6 weeks, platelets fall to  $55 \times 10^9/L$ , Hb 8.5 g/dL, ANC  $1.2 \times 10^9/L$ . What is the best next step?
  - A. Continue same dose — cytopenias usually transient
  - B. Hold ruxolitinib for 1 week then restart full dose
  - C. Switch to belumosudil immediately
  - D. Reduce ruxolitinib to 5 mg BID
  - E. Add G-CSF and continue 10 mg BID

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- According to the 2020 NIH/EBMT chronic GVHD risk model, which of the following factors most strongly predicts non-relapse mortality and overall survival?
  - A. Extent of skin involvement (>50% BSA)
  - B. Presence of oral GVHD alone
  - C. Thrombocytopenia ( $<100 \times 10^9/L$ )
  - D. Time from transplant >12 months
  - E. History of acute GVHD

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# Case 5

- A 63-year-old woman, day +25 after axi-cel, remains neutropenic (ANC  $0.6 \times 10^9$ /L) with IgG  $< 400$  mg/dL. Which of the following best reflects the recommended prophylaxis strategy after CAR-T therapy?
- A. Levofloxacin for 6 months
- B. No need for PJP prophylaxis if ANC  $> 0.5 \times 10^9$ /L and IgG  $> 200$  mg/dL
- C. Use IVIG only when IgG  $< 200$  mg/dL or symptomatic infections with high CD4/CD8
- D. Live vaccines may start at 3 months post-infusion
- E. Continue antiviral + PJP prophylaxis for  $\geq 6$ –12 months or until CD4  $> 200$

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# Case 5

- Same patient, 6 months post-CAR-T, clinically well, IgG > 500 mg/dL, no ongoing immunosuppression. When should **revaccination** begin?
- A. Immediately after lymphocyte recovery
- B. Inactivated vacc. at 3–6 months; live vacc.  $\geq$  12 months
- C. All vaccines delayed  $\geq$  24 months
- D. Live and inactivated vaccines both at 6 months
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- At day +90, patient's IgG remains  $< 300$  mg/dL despite no infection and slow B-cell recovery. What is the best approach?
- A. Give a single IVIG dose and stop prophylaxis
- B. Begin routine monthly IVIG for 12 months regardless of infections
- C. Start IVIG only if febrile neutropenia develops
- D. Add broad-spectrum antibiotics instead of IVIG
- E. initiate IVIG replacement

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# Case 6

- A 62-year-old AML patient in CR1 has A 1-Ag mismatched unrelated donor (MMUD) PBSC graft planned with RIC Flu/Bu2. Which GVHD-prophylaxis platform reflects modern best practice?
- A. PTCy + tacrolimus + MMF
- B. Ruxolitinib with MMF
- C. Tacrolimus + MTX only
- D. ATG + sirolimus

# Case 6

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# Case 6

- Day +25 post-HCT the patient develops grade II skin + GI GVHD while on PTCy–Tac–MMF. What is the recommended first-line treatment?
  - A. Methylprednisolone 2 mg/kg/day
  - B. ECP with Tac levels above 20 ng/ml
  - C. ruxolitinib
  - D. ATG bolus
  - E. Withdraw tacrolimus to boost GVL

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# Case 6

- Developed grade II acute skin + lower GI GVHD on day +25. After 7 days of methylprednisolone 2 mg/kg/day, there is no clinical improvement. What is the most appropriate next management step?
  - A. Increase steroid dose to 4 mg/kg/day
  - B. Add ATG
  - C. Ruxolitinib 10 mg BID
  - D. Switch immunosuppression to sirolimus monotherapy
  - E. Start ibtutinib

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# Case 7

- A 58-year-old man with DLBCL relapsed 8 months after R-CHOP. He received 2 cycles of R-DHAP, achieving PR on PET (Deauville 4). According to current evidence and guidelines, which statement best reflects modern practice?
  - A. Autologous HCT.
  - B. CAR-T.
  - C. CAR-T if double-hit cytogenetics in PR.
  - D. High dose melphalan and autologous HCT only if good performance status.
  - E. Observation and treat only if progression documented.

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# Case 7

- The same patient's salvage PET shows residual 3 cm uptake (SUV 6). The transplant team plans BEAM conditioning. Which statement is most accurate regarding conditioning and disease control before auto-HCT?
  - A. TBI-based conditioning is superior to BEAM in DLBCL.
  - B. Patients in PR after salvage can proceed to BEAM auto-HCT, but CR is better than PR.
  - C. BEAM intensity should be increased for PET-positive cases.
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- The patient relapses 6 months post-auto-HCT with bulky nodal disease, LDH ↑, ECOG 1. He receives bridging pola-BR before **axi-cel** infusion. Which pre-CAR-T factor best predicts **ICANS** and **non-response**?
  - A. CR prior to CART is associated with no expansion and response
  - B. Bendamustine as bridging prior to infusion is associated with decreased incidence of ICANS
  - C. Patient age > 60 y independently increases severe ICANS risk
  - D. Elevated ferritin at infusion correlates with higher CRS/ICANS risk and lower response
  - E. Prior auto-HCT is associated with decrease response

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