Patient Information

Name: Yi Zhu ID: SYN228 Date of Birth: 04/08/1963 Admit Date: 04/02/2025 Discharge Date: 04/14/2025 Length of Stay: 12 days Attending: Dr. Alexandra Lee

Assessment and Plan (Brief Overview)

62yo female with high PD-L1 wild-type NSCLC on pembrolizumab (cycle #41) with exceptional and durable response (37 months) now with Grade 3 immune-related myocarditis and Grade 2 hepatitis requiring permanent discontinuation of immunotherapy.

What happened: Presented with progressive dyspnea, chest discomfort, elevated troponin, and liver enzymes. Echo showed reduced LVEF 40% (baseline 60%). Cardiac MRI and biopsy confirmed immune-mediated myocarditis. Treated with high-dose steroids with significant clinical and laboratory improvement.

On discharge: Symptoms resolved. Cardiac function improved (LVEF 50%). Liver enzymes normalizing. Systemic disease remains in near-complete response.

Follow-up: Cardio-oncology in 1 week, Oncology in 2 weeks. Will not resume immunotherapy due to life-threatening toxicity.

Hospital Course (Detailed)

Ms. SYN228 is a 62-year-old female with stage IVB NSCLC (diagnosed 02/25/2022) with liver and contralateral pulmonary metastases and no actionable mutations but high PD-L1 expression (80% TPS). She has had exceptional response to pembrolizumab monotherapy since 03/18/2022 with >90% reduction in primary tumor and liver metastases, continuing to cycle #41 (last dose 03/22/2025).

Patient presented with 5 days of progressive shortness of breath, chest discomfort, and generalized weakness. Initial evaluation revealed troponin elevation (5.2 ng/mL), ECG changes, and transaminitis.

Cardiac workup included transthoracic echocardiogram showing reduced LVEF (40% from baseline 60%) with global hypokinesis, coronary angiography showing no obstructive disease, and cardiac MRI demonstrating diffuse myocardial edema with non-vascular pattern of late gadolinium enhancement. Endomyocardial biopsy confirmed lymphocytic infiltrate with CD8+ T-cell predominance consistent with immune-mediated myocarditis.

Hepatic evaluation revealed elevated transaminases (ALT 185 U/L, AST 142 U/L) with normal bilirubin and alkaline phosphatase. Other causes of hepatitis were excluded, confirming immune-related etiology.

Treatment consisted of pembrolizumab discontinuation, high-dose methylprednisolone (1000 mg IV daily for 3 days) followed by oral prednisone (1 mg/kg/day) with slow taper plan. Heart failure medications were initiated per cardio-oncology protocol. Telemetry monitoring showed no significant arrhythmias throughout hospitalization.

Clinical response was excellent with troponin trending down, LVEF improving to 50%, and liver enzymes normalizing. No other organ systems showed evidence of immune-related toxicity.

A formal case review was conducted on April 10, 2025, with medical oncology, cardiology, and cardio-oncology specialists. Given the severity of myocarditis (Grade 3) and excellent prior response, the consensus recommendation was permanent discontinuation of pembrolizumab with close surveillance.

Laboratory Data

Test	On Admission	Peak/ Nadir	Discharg e	Reference Range
Troponir I	5.2	8.4	0.8	<0.04 ng/mL
CK-MB	36	48	12	0-7 ng/mL
BNP	865	1100	350	<100 pg/mL
ALT	148	185	65	7-56 U/L
AST	122	142	48	8-48 U/L
Alk Phos	115	115	96	45-115 U/L

Test	On	Peak/	Discharg	Reference
	Admission	Nadir	е	Range
T. Bili	1.2	1.2	0.8	0.2-1.2 mg/dL

Diagnostic Studies

- 1. Echocardiogram (04/02/2025):
 - LVEF 40% (decreased from 65% baseline)
 - Global hypokinesis
 - Small pericardial effusion
- 2. Cardiac MRI (04/04/2025):
 - o Diffuse myocardial edema on T2-weighted images
 - o Patchy late gadolinium enhancement in non-vascular distribution
 - Findings consistent with myocarditis
- 3. Endomyocardial biopsy (04/05/2025):
 - o Lymphocytic infiltrate with CD8+ predominance
 - Scattered eosinophils
 - Myocyte necrosis
 - Consistent with immune-mediated myocarditis
- 4. Repeat Echocardiogram (04/12/2025):
 - LVEF improved to 50%
 - Mild global hypokinesis
 - Stable small pericardial effusion

Discharge Medications

- 1. Prednisone 60 mg PO daily × 2 weeks, then taper by 10 mg every week
- 2. Metoprolol succinate 25 mg PO daily
- 3. Lisinopril 2.5 mg PO daily
- 4. Famotidine 20 mg PO BID
- 5. Levothyroxine 112 mcg PO daily (home medication for prior immune-related hypothyroidism)
- 6. Calcium/Vitamin D supplementation

Follow-up Appointments

- 1. Cardio-oncology: April 21, 2025 (1 week)
- 2. Medical oncology: April 28, 2025 (2 weeks)

- 3. Cardiology: May 12, 2025 (4 weeks)
- 4. Lab monitoring: Weekly troponin, BNP, CMP × 4 weeks
- 5. Imaging: Repeat echocardiogram in 2 weeks, CT chest/abdomen/pelvis in 8 weeks

Special Instructions

- 1. Report immediately:
 - Worsening shortness of breath
 - Chest pain or pressure
 - Irregular heartbeat or palpitations
 - Dizziness or lightheadedness
 - Weight gain >2 kg in 3 days
- 2. Activity restrictions:
 - No heavy lifting (>10 pounds) for 2 weeks
 - No strenuous exercise until cardiology clearance
 - Maintain light physical activity as tolerated
 - Daily weight monitoring
- 3. Medication precautions:
 - Follow steroid taper exactly as prescribed
 - o Take all heart medications at the same time daily
 - Avoid NSAIDs and decongestants

Condition at Discharge

Stable with significant improvement in symptoms. Able to ambulate without dyspnea. No chest pain. Oxygen saturation 98% on room air. Vitals stable. LVEF improved to 50%. Liver enzymes normalizing. ECOG performance status 1 (baseline 0).

Adverse Event Reporting

This case has been submitted to the FDA Adverse Event Reporting System (FAERS) and added to the Immune-Related Adverse Events (irAE) Registry for long-term follow-up.

Alexandra Lee, MD, PhD Medical Oncology April 14, 2025 16:45