DISCHARGE SUMMARY

PATIENT IDENTIFICATION

Admission 04/06/2025 - 04/14/2025

Attending Physician: Dr. Emma Richardson, Medical Oncology Consulting Services: Interventional Radiology, Thoracic Surgery

DIAGNOSIS AT DISCHARGE

Metastatic KRAS G12D-positive non-small cell lung cancer with progression on second-line therapy

- 1. Status post CT-guided biopsy of new liver lesion for molecular profiling
- 2. Managed pleural effusion via PleurX catheter placement

DETAILED ONCOLOGICAL HISTORY

Initial Diagnosis (December 2022)

Patient initially presented with persistent cough and right-sided chest pain. Chest CT revealed a 3.8 cm right upper lobe mass with right pleural effusion and a 2.5 cm right adrenal mass. Thoracentesis performed on 12/28/2022 yielded malignant cells.

Histopathology (Initial Diagnosis)

CT-guided Biopsy of Right Upper Lobe Mass (12/31/2022):

- **Gross Description:** Three cores of tan-gray lung tissue, each measuring 0.8-1.2 cm in length
- Microscopic Description:
 - o Infiltrating adenocarcinoma with predominant acinar pattern (60%)
 - Solid component (30%)
 - Micropapillary features (10%)
 - o Moderate nuclear pleomorphism with prominent nucleoli
 - Abundant eosinophilic cytoplasm
 - Occasional intracytoplasmic vacuoles
 - Mitotic rate: 8 per 10 HPF
 - Surrounding desmoplastic stroma with moderate lymphocytic infiltration
 - No lepidic growth pattern identified
 - Surrounding lung parenchyma with anthracotic pigment
- Immunohistochemical Profile:
 - o TTF-1: Strongly positive (90% of tumor cells)

- Napsin A: Positive (diffuse, moderate intensity)
- CK7: PositiveCK20: Negative
- p40: NegativeCDX2: Negative
- o GATA3: Negative
- Synaptophysin: Focal weak positivity (5% of cells)
- o Chromogranin: Negative
- Ki-67: 40% proliferation index

• Molecular Testing:

- Next-Generation Sequencing Panel:
 - KRAS G12D mutation (allelic frequency 35%)
 - TP53 R273H mutation (allelic frequency 42%)
 - STK11 frameshift deletion (p.P281fs*6)
 - No other actionable alterations (EGFR, ALK, ROS1, BRAF, MET, RET, NTRK all wild-type)
- o PD-L1 (22C3): TPS <1%, CPS 5, IC 3%
- o TMB: Intermediate (8 mutations/Mb)
- o MSI Status: Stable

Pleural Fluid Cytology (12/28/2022):

- Malignant cells present consistent with adenocarcinoma
- Immunocytochemistry positive for TTF-1 and CK7

Staging

Clinical stage IVA (cT2bN1M1a) with right adrenal metastasis and malignant pleural effusion.

Treatment Course

1. First-Line Therapy:

- Carboplatin AUC 5 + Pemetrexed 500 mg/m² + Pembrolizumab 200 mg IV q3 weeks
- Initiated 01/23/2023
- Completed 4 cycles with partial response
- Continued pembrolizumab + pemetrexed maintenance
- Disease progression documented after 11 months (12/2023)

2. Second-Line Therapy:

- Docetaxel 75 mg/m² + Ramucirumab 10 mg/kg IV q3 weeks
- o Initiated 01/08/2024
- Initial stable disease for 6 months
- o Disease progression documented on 03/25/2025 scan

3. Procedures:

- o Therapeutic thoracentesis (12/2022, 01/2023)
- o Talc pleurodesis (02/2023) initially successful
- Repeat thoracentesis for recurrent effusion (03/2025)

CURRENT ADMISSION DETAILS

Reason for Admission

Patient was admitted for management of progressive dyspnea due to recurrent pleural effusion and to undergo CT-guided biopsy of newly identified liver lesion for comprehensive resistance mechanism profiling.

Hospital Course

1. Pleural Effusion Management:

- Diagnostic thoracentesis performed on admission confirmed malignant cells
- PleurX catheter placed by Interventional Radiology on 04/07/2025
- Initial drainage of 1,250 mL of serosanguineous fluid with symptomatic improvement
- o Patient and family educated on home management of PleurX catheter

2. Liver Biopsy:

- CT-guided biopsy of segment VII liver lesion performed on 04/08/2025
- o Procedure well-tolerated without complications
- Preliminary pathology confirmed metastatic adenocarcinoma consistent with lung primary
- Tissue submitted for comprehensive molecular profiling including RNA sequencing and immune profiling

3. Comprehensive Restaging:

- o CT chest/abdomen/pelvis with contrast (04/10/2025) showed:
 - Increase in size of primary RUL mass (4.2 cm from 3.6 cm)
 - Interval development of liver metastases (largest 2.1 cm in segment VII)
 - Stable right adrenal metastasis
 - Moderate right pleural effusion
 - No other new sites of disease

4. Multidisciplinary Tumor Board Discussion (04/13/2025):

- Reviewed complete clinical history and pathology
- Noted co-mutations in KRAS, TP53, and STK11 ("KPS" phenotype) associated with primary resistance to immunotherapy
- Discussed therapeutic options including:
 - Docetaxel monotherapy (lowered efficacy expectations given ramucirumab failure)
 - Gemcitabine-based chemotherapy
 - Clinical trial options: KRAS G12D-directed degrader (Phase I trial at our institution)
 - Expanded molecular and immune profiling may identify additional therapeutic targets

Histopathology (Current Admission)

CT-guided Biopsy of Liver Lesion (04/08/2025):

• **Gross Description:** Four cores of tan-brown liver tissue, each measuring 0.6-1.0 cm in length

Microscopic Description:

- Metastatic adenocarcinoma involving liver parenchyma
- Predominantly solid growth pattern (80%)
- o Reduced acinar component (20%) compared to primary tumor
- o Increased nuclear pleomorphism compared to primary tumor
- Higher mitotic rate (15 per 10 HPF)
- Focal tumor necrosis (approximately 10% of tumor volume)
- o Surrounding hepatic parenchyma uninvolved

Immunohistochemical Profile:

- TTF-1: Remains positive (75% of tumor cells)
- Napsin A: Weakly positive (reduced from primary)
- CK7: Positive
- Ki-67: 60% proliferation index (increased from 40% in primary)
- Additional markers being processed

Preliminary Molecular Testing:

- KRAS G12D mutation confirmed (allelic frequency 42%)
- TP53 R273H mutation confirmed
- STK11 frameshift deletion confirmed
- Additional comprehensive molecular profiling pending (results expected in 10-14 days)

Pleural Fluid Cytology (04/06/2025):

- Malignant cells present consistent with known adenocarcinoma
- Higher nuclear grade compared to previous cytology specimens
- Increased N:C ratio and nuclear irregularity

DISCHARGE PLAN

Medications:

- 1. Current home medications to continue
- 2. Dexamethasone 4 mg PO daily for 5 days, then 2 mg daily for 5 days, then discontinue
- 3. Hydrocodone/acetaminophen 5/325 mg PO q6h PRN pain
- 4. Ondansetron 8 mg PO q8h PRN nausea

Follow-up Appointments:

- 1. Medical Oncology (Dr. Richardson): 04/22/2025 at 10:00 AM
 - Review final molecular profiling results
 - Discuss third-line therapy options
 - Consider clinical trial referral

2. Interventional Radiology: 04/21/2025 at 2:00 PM

PleurX catheter check

3. Palliative Care: 04/23/2025 at 11:00 AM

- Symptom management optimization
- Goals of care discussion

Home Care Instructions:

- 1. PleurX catheter drainage:
 - Drain 500-1000 mL every other day or PRN symptoms
 - o Record drainage amounts for review at follow-up
 - Call office for increased pain, redness around catheter site, or fever >38.0°C
- 2. Monitoring:
 - o Daily weight
 - Track temperatures twice daily
 - Record pain levels and medication use

Additional Plans:

- 1. Complete molecular profiling results anticipated by 04/24/2025
- 2. Phase I clinical trial screening visit pending molecular results
- 3. Optimization of symptom management plan at oncology follow-up

CONDITION AT DISCHARGE

Patient is hemodynamically stable with improved respiratory status following PleurX catheter placement. Ambulating with minimal assistance. Pain controlled on current regimen. ECOG Performance Status 2 (baseline 1 prior to recent progression).

Electronically signed by: Emma Richardson, MD Medical Oncology 04/14/2025 16:30