



An Exceptional Coexistence: Primary Hepatic Leiomyosarcoma and Basal Cell Carcinoma: A Case Report

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ABSTRACT

Background: Primary hepatic leiomyosarcoma (LMSH) is a rare nosological entity originating from the mesenchymal lineage. Its co-occurrence with basal cell carcinoma (BCC), a neoplasm of epithelial origin, constitutes an exceptionally rare and poorly documented association. We present the case of a 43-year-old patient whose clinical course was marked by the successive diagnosis of these two distinct pathologies.

Case Presentation: In 2015, a primary LMSH was managed with surgical resection followed by adjuvant chemotherapy (doxorubicin and ifosfamide). Subsequently, the development of locally advanced and unresectable BCC in 2018 required targeted therapy with vismodegib, a Hedgehog pathway inhibitor. This case highlights the therapeutic challenges posed by multiple primary neoplasms and underscores the importance of vigilant oncologic surveillance.

Conclusion: This rare coexistence emphasizes the need for multidisciplinary management and lifelong surveillance in patients with rare primary malignancies.

Keywords: Hepatic Leiomyosarcoma, Basal Cell Carcinoma, Multiple Primary Neoplasms, Vismodegib, Hedgehog Signaling Pathway.

Case Studies

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Introduction

Leiomyosarcoma, a malignant tumor arising from smooth muscle cells, constitutes 10–20% of all soft tissue sarcomas [1]. Its primary hepatic localization remains exceptionally rare, posing significant diagnostic and therapeutic challenges [2]. In contrast, basal cell carcinoma (BCC) is the most prevalent cutaneous malignancy in the general population, with increasing incidence [3]. The simultaneous occurrence of these neoplasms—distinct in embryological origin and biology—raises complex questions regarding diagnosis,

pathophysiology, and prognosis. While multiple primary neoplasms in a single patient are recognized, the coexistence of LMSH and BCC is exceptionally rare [4, 5]. We report this unique case to discuss management challenges and prognostic implications.

Case Presentation

A 43-year-old male with no significant medical history presented with two successive neoplasms.

- **2015:** Diagnosis of LMSH. Partial surgical resection (R2 margin) was performed, followed by adjuvant chemotherapy (doxorubicin and ifosfamide) for eight cycles.
- **2017:** Unresectable hepatic progression was treated with radiofrequency ablation. Second-line chemotherapy (ifosfamide) was initiated for nine cycles (2017–2022), achieving disease stability.
- **Late 2023:** Development of three frontal tumors. Surgical resection revealed infiltrated margins, with histopathology confirming BCC.
- **2024:** New lesions on the back and neck. MRI showed infiltrative cutaneous/subcutaneous involvement with osseous lysis. Histology confirmed recurrent, multifocal, nodular, keratinizing, and pigmented BCC. Given unresectability, vismodegib (Hedgehog inhibitor) was initiated in August 2024. The patient exhibited partial radiological response and clinical improvement.

Discussion

Diagnostic and Differential Diagnosis

- **LMSH Diagnosis:** Relies on correlation of clinical, radiological, and histopathological findings. Imaging suggests a heterogeneous mass with necrosis/hemorrhage, but histology (spindle cells, marked atypia, high mitotic rate) and immunohistochemistry (smooth muscle markers: actin, desmin, h-caldesmon; CD117/DOG1-negative) confirm the diagnosis [2] (Figure 1).

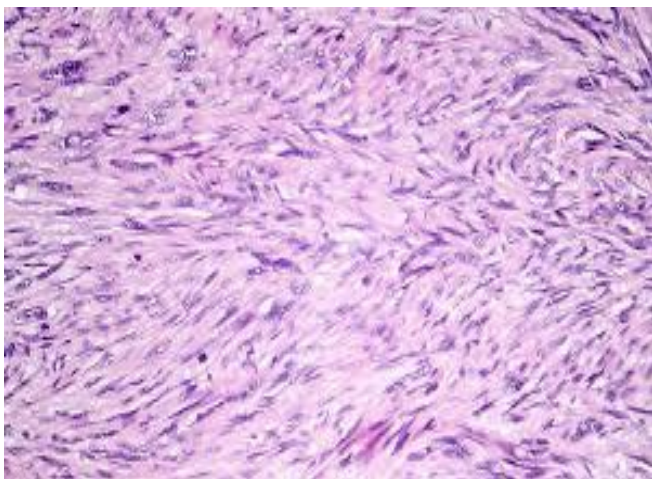


Figure 1: Immunohistochemistry of Hepatic Leiomyosarcoma

- **BCC Diagnosis:** Defined by basaloid nests with palisading nuclei and stromal retraction, confirmed by CK5/6 and BerEP4 positivity. Absence of metastatic LMSH supports independent primary tumors [3].
- **Contrast in Biology:** LMSH is diagnosed late with poor prognosis dependent on R0 resection, while BCC is typically early-stage with slow local progression and

rare metastasis. This contrast supports distinct primary origins (Figure 2).

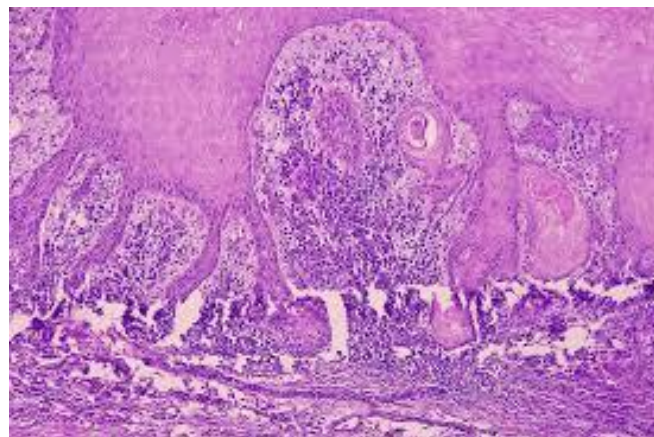


Figure 2: Immunohistochemistry of Basal Cell Carcinoma

Pathophysiological Hypotheses

1. **Coincidental Association:** High BCC incidence may statistically co-occur with rare neoplasms without shared biology.
2. **Genetic Susceptibility:** LMSH involves TP53/RB1/PTEN alterations; BCC is driven by Hedgehog pathway mutations (PTCH1/SMO). A global DNA repair defect could theoretically predispose to both [8, 9].
3. **Environmental/Immunological Factors:** UV exposure is a major BCC risk. Immunosuppression (e.g., UV-induced or systemic) may facilitate multiple primary cancers [3].

Molecular Biology: Insights and Limitations

Comparative genomic analysis (e.g., NGS) could clarify clonal vs. independent origins but is lacking in this case. Future studies should integrate molecular profiling to elucidate shared mechanisms [10].

Clinical Implications and Management

- **Diagnostic Challenge:** New lesions in rare-cancer patients require thorough pathology to exclude metastasis.
- **Multidisciplinary Approach:** Collaboration among hepatologists, oncologists, surgeons, dermatologists, and pathologists is critical.
- **Lifelong Surveillance:** First atypical cancers increase risk of subsequent primaries, warranting lifelong monitoring [5].

Limitations and Perspectives

- **Retrospective, Single Case:** Limits causal inference and generalizability. Lack of molecular data hampers biological insights.
- **Value of Case Reports:** Documenting rare cases enriches literature, stimulates international registries, and generates testable hypotheses.

Conclusion

The coexistence of LMSH and BCC exemplifies the diagnostic and therapeutic challenges of multiple primary neoplasms. Successful management hinges on multidisciplinary collaboration and molecular integration. Future multicentric studies with genomic profiling are essential to transform isolated observations into robust biological insights.

Conflict of Interest

The authors declare that they have no links of interest.

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Declarations

This study used anonymized patient data. As the research involved no direct intervention or modification of standard patient care, formal approval from an ethics committee was not required in accordance with institutional and national guidelines for observational studies. All patient data were anonymized prior to analysis to protect confidentiality and treated according to the Algerian national guidelines

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Authors' Contributions

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Formal Analysis: Sihem BENSALÉM; **Investigation:** Abdelaziz AMMARI, Sihem BENSALÉM, Assia BENSALÉM; **Resources:** Abdelaziz AMMARI; **Data Curation:** Abdelaziz AMMARI, Sihem BENSALÉM, Assia BENSALÉM; **Writing - Original Draft Preparation:** Abdelaziz AMMARI, Sihem BENSALÉM; **Writing - Review & Editing:** Abdelaziz AMMARI, Sihem BENSALÉM, Assia BENSALÉM; **Visualization:** Sihem BENSALÉM; **Supervision:** Abdelaziz AMMARI, Sihem BENSALÉM; **Project Administration:** Assia BENSALÉM; **Funding Acquisition:** Not applicable.

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