# Avascular Transformation of Hepatocellular Carcinoma with Systemic Immunotherapy



Karmanos

CANCER INSTITUTE

Wavne State University

Mithil Gudi BS<sup>1</sup>, Alexander Eskandarian BS<sup>1</sup>, Ali Yalcintepe MD<sup>1,2</sup>, Karan T. Singh MD<sup>1,3</sup>, Jeffrey Critchfield MD<sup>1,2,4</sup>

Wayne State University School of Medicine<sup>1</sup>, Detroit Medical Center<sup>2</sup>, John. D Dingell VA Medical Center<sup>3</sup>, Karmanos Cancer Institute<sup>4</sup>

## Background

- The combination of local embolization and immunotherapy for hepatocellular carcinoma (HCC) is an emerging approach.
- While atezolizumab and bevacizumab therapy is considered the gold standard, the STRIDE regimen, a dual immunotherapy therapy consisting of a priming dose of tremelimumab (anti-CTLA-4) with monthly durvalumab (anti-PD-L1), has grown in use.
- We present a unique case of a patient who was unintentionally initiated on STRIDE regimen between mapping angiography and embolization, which seemingly led to rapid avascular tumor transformation.

# **Case Summary**

- A 74-year-old male with HCC, HFrEF w/ICD, HTN, CAD (s/p LAD stent) underwent mapping angiography, revealing hypervascularity of the target HCC from numerous vascular hepatic segmental, gastroepiploic, and inferior phrenic branches.
- Given the extensive vascular supply and high lung shunt fraction (12%), the plan shifted from Y-90 to transarterial chemoembolization (TACE) embolization with 300–500-micron drug-eluting doxorubicin beads.
- Repeat embolization angiography, performed 6 weeks later due to follow-up difficulties, showed substantially decreased tumor vascularity in the hepatic flexure branch of the gastroepiploic artery, inferior phrenic artery, and the segment 4 and 5 right hepatic artery branches.
- The inferior phrenic, hepatic segment 4, and hepatic segment 5 arteries were chemoembolized. Afterward, discussion with the patient's oncologist revealed that the STRIDE regimen had been initiated under the assumption that embolization had already occurred, with the last dose 7 days prior.

# Case Images



Figure 1: Axial CT image of target hepatocellular carcinoma

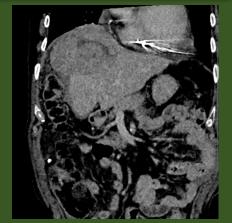


Figure 2: Coronal CT image of target hepatocellular carcinoma

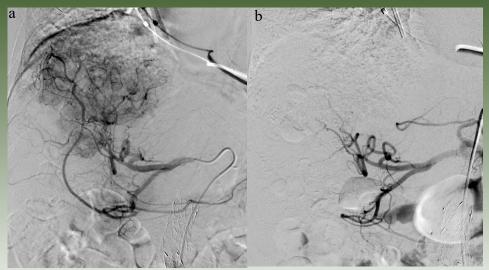


Figure 3: a) DSA from celiac artery showing tumor mass and vasculature during mapping procedure; b) DSA run from celiac artery when patient returned for TACE embolization

## Discussion

- The substantially decreased vascularity of the tumor compared to initial mapping is possibly attributed to the initiation of systemic treatment.
- The patient continued immunotherapy with complete mRECIST response was identified on 1 and 3 month imaging.
  - Unfortunately, the patient passed away 3 months later due to unrelated endocarditis and septic shock.
- The unanticipated reduction in HCC vascularity within a 6-week period is interesting given that the therapeutic mechanism of tremelimumab and durvalumab is not rooted in anti-angiogenic properties.
- Durvalumab, while not generally classified as a vesicant, is a platinum compound
  - Platinum compounds such as cisplatin and oxaliplatin are known to have some vesicant properties leading to tissue damage and microthombosis
- The avascular transformation could also be attributed to a strong, local immune response due to the immune checkpoint inhibitor properties of the STRIDE regimen

#### Conclusion

- With the growing use of immunotherapy treatments, their impact on tumor vascularity and sequencing with local embolotherapy warrants further investigation.
- This finding highlights the importance of timing and communication among multidisciplinary teams in delivering oncologic care.

### References

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