



# The Imaging Negative Hepatic Lesions: A Rare Case of Infiltrative Hepatocellular Carcinoma.



Onyinye Ugonabo MD<sup>1</sup>, Joseph Simmons MD<sup>1</sup>, Saba Altaranweh MD<sup>1</sup>, Philip R Jones DO<sup>1</sup>,  
Tejas Joshi MD<sup>1</sup>, Wesam Frandah MD<sup>1</sup>

<sup>1</sup> Marshall University, Joan C Edward School of Medicine, Huntington, West Virginia.

## Introduction

- Hepatocellular carcinoma (HCC) is the 6<sup>th</sup> most common cancer and the 4<sup>th</sup> leading cause of cancer related death world wide [1].
- The infiltrative type is most difficult to diagnose with imaging because of its inherently defined micronodules involving a segment or entire hepatic parenchyma without an identifiable mass.
- HCV remains the primary risk factor for HCC in the United States.
- Prognosis is poor and estimated at a 5 year survival of <20%.

## Case presentation

A 61 year old female with a past medical history of HCV cirrhosis with sustained virological response (SVR) presented with abdominal pain and worsening lower extremity edema.

**Examination findings:** distended abdomen, bilateral lower extremity pitting edema.

**Significant work up findings:** AST of 159unit/l (15-37), platelet of 109k/cmm (150-440), total bilirubin of 1.2mg/dl (0.2-1), AFP of >20,000ng/ml (0.5-8).

Right upper quadrant ultrasound, CT and MRI identified no hepatic mass, Fig A

Due to markedly elevated AFP, HCC was highly suspected. The patient proceeded to having an EGD-EUS fine needle guided aspiration done showing multiple infiltrative hepatic lesions, fig B and, biopsy was taken.

Biopsy report malignant cells positive for AFP with cells reactive for glypican-3 and negative for Hep-par1, supporting the diagnosis of HCC, fig C.

## Treatment and follow up

The patient was referred to oncology and a month later, she died.

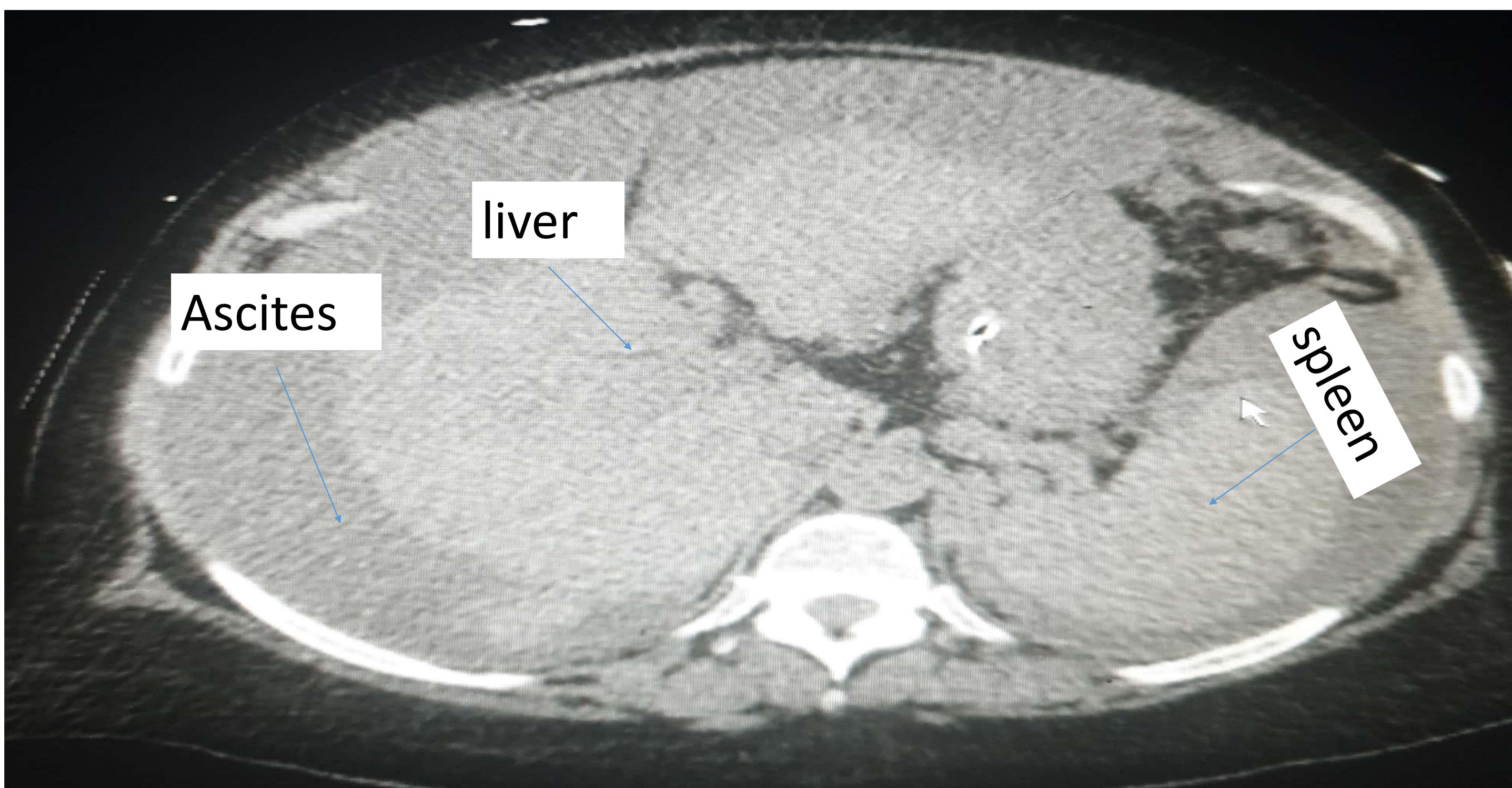


Fig A: CT abdomen showing cirrhotic liver with no defined hepatic mass.

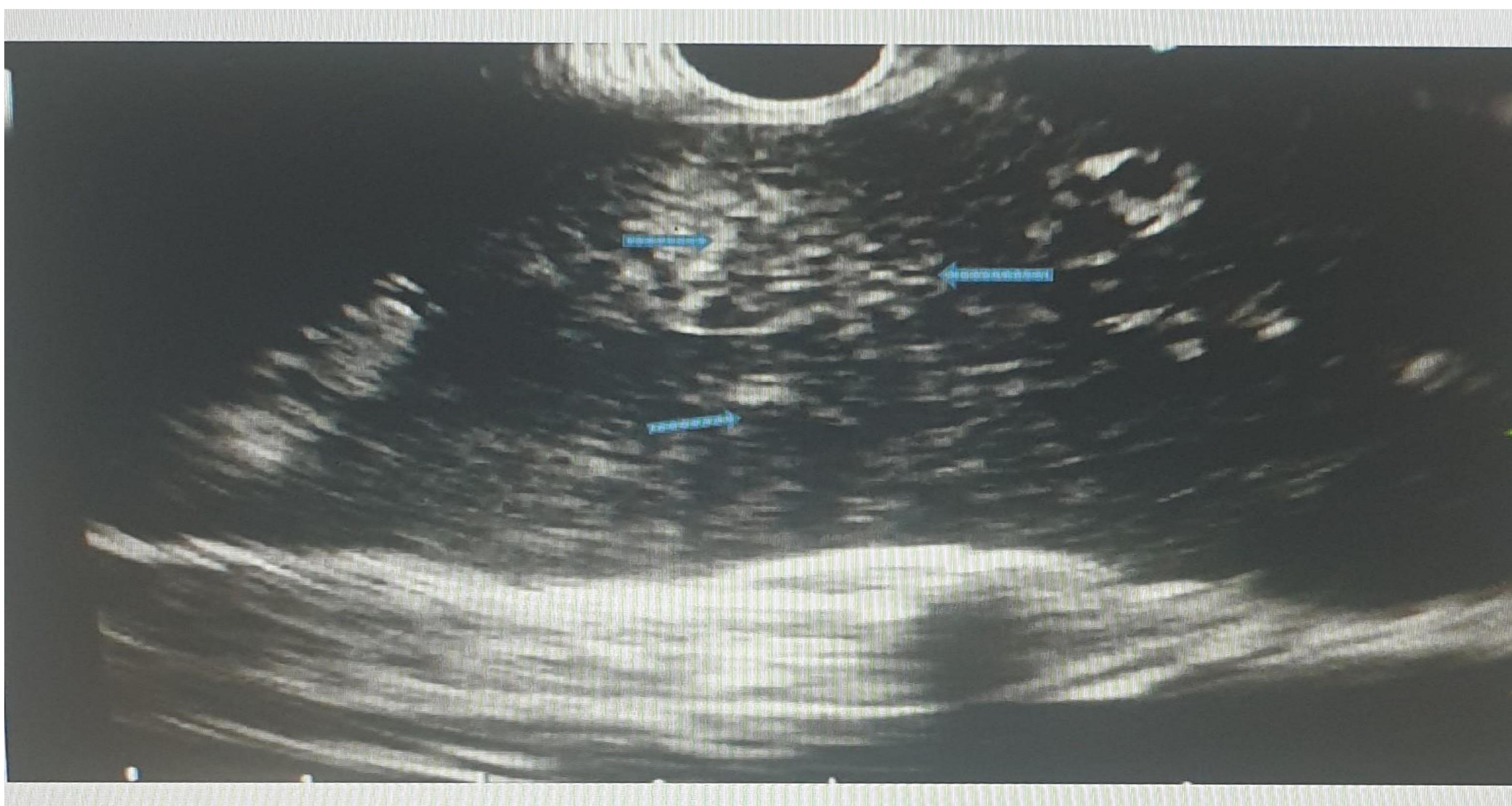


Fig B: EGD-EUS showing multiple infiltrative hepatic lesions.

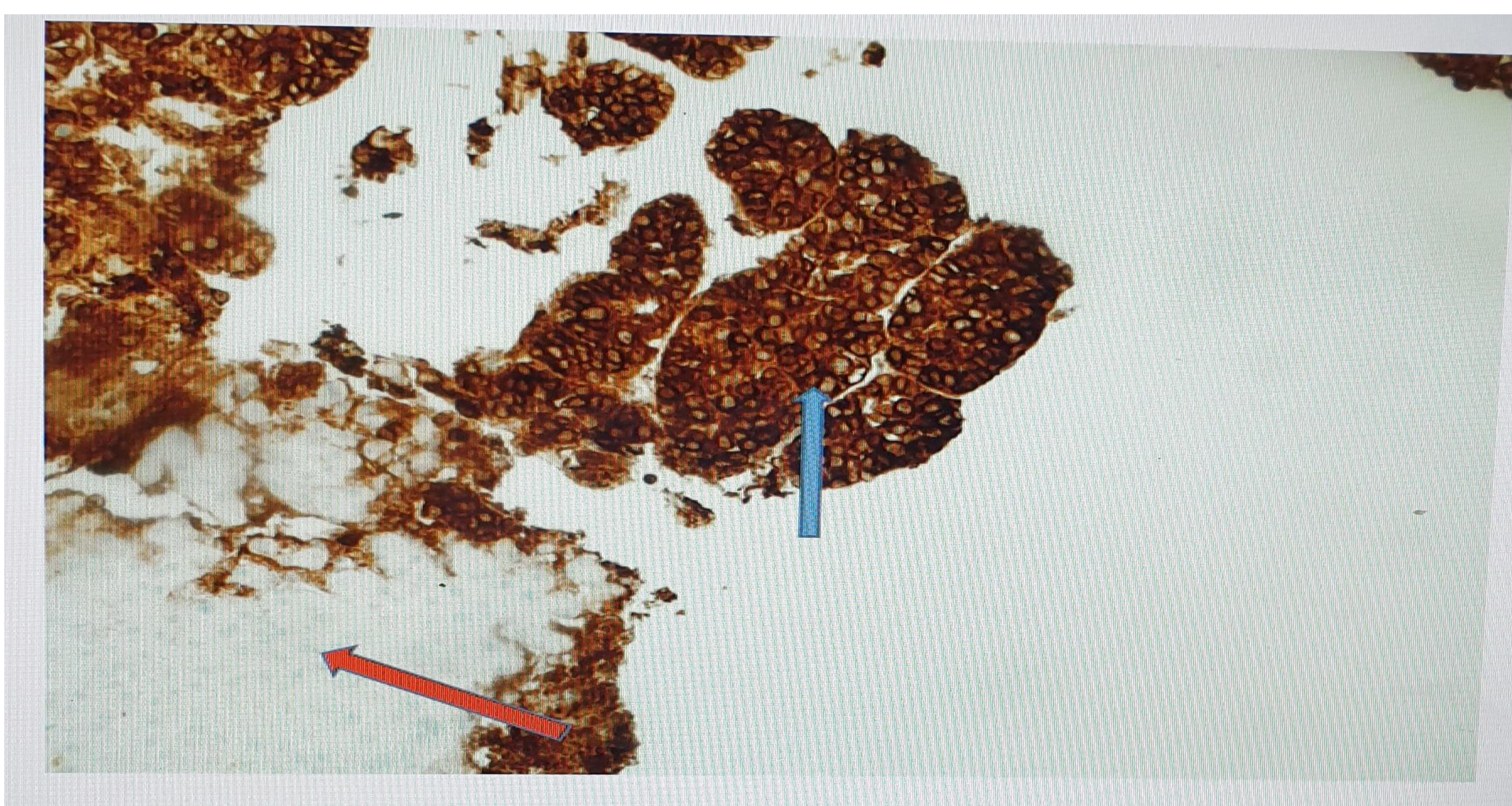


Fig C; glypican-3 immunostain with strong diffuse staining (blue arrow), background non neoplastic liver (red arrow).

## Discussion

- SVR is associated with decreased risk of HCC.
- Cirrhosis directly increases risk of developing HCC
- Diagnosis of HCC can be achieved non invasively using abdominal ultrasound, CT, MRI and EUS-FNA.
- Abdominal ultrasound is the initial modality recommended for HCC surveillance, with a sensitivity for detecting early HCC about 47% [2].
- AFP cut off level of >20ng/ml has a sensitivity of 60% with low specificity
- A level of > 400ng/ml is diagnostic of HCC with a specificity of almost 100%. Incremental changes in AFP is associated with increased mortality rate.
- The median survival rate of infiltrative HCC with AFP of > 400 is estimated to be 5 months
- EUS is superior to CT in detecting small hepatic lesions, with a sensitivity of 100% compared to 71% of CT [3]
- The data on the treatment of infiltrative HCC is still under review. Intra-arterial therapy and chemotherapy like sorafenib have shown a survival benefit
- We recommend that EUS be considered an integral modality while investigating HCC.

## References

1. Llovet, J.M., Kelley, R.K., Villanueva, A. *et al.* Hepatocellular carcinoma. *Nat Rev Dis Primers* **7**, 6 (2021). <https://doi.org/10.1038/s41572-020-00240-3>
2. Lim J, Singal AG. Surveillance and Diagnosis of Hepatocellular Carcinoma. *Clin Liver Dis (Hoboken)*. 2019;13(1):2-5. doi:10.1002/cld.761
3. Girotra M, Soota K, Dhaliwal AS, Abraham RR, Garcia-Saenz-de-Sicilia M, Tharian B. Utility of endoscopic ultrasound and endoscopy in diagnosis and management of hepatocellular carcinoma and its complications: What does endoscopic ultrasonography offer above and beyond conventional cross-sectional imaging?. *World J Gastrointest Endosc*. 2018;10(2):56-68. doi:10.4253/wjge.v10.i2.56