

AMSER Rad Path Case of the Month:



28-year-old nulliparous female with liver cirrhosis presents for follow-up after abnormal lab results

Chintan Rajput, MS-IV, MBS

Rowan University School of Osteopathic Medicine

Linda White Nunes, MD, MPH

University of Pennsylvania Penn Medicine – Department of Radiology

Rashmi Tondon, MD

Department of Pathology and Laboratory Medicine University of Pennsylvania Perelman School of Medicine



Patient Presentation

28-year-old GOPO female presenting for evaluation of abnormal lab results

- <u>PSHx</u>: Fontan, serial liver biopsies
- <u>PMHx</u>: HLHS, ASD, Tricuspid insufficiency, SVT, Asthma, Protein losing enteropathy, *Liver cirrhosis*
- <u>Medications</u>: Aspirin, Cholecalciferol, Enalapril, Furosemide, Adenosine, Metoprolol, Spironolactone, Warfarin
- <u>Allergies</u>: NKDA

Pertinent Labs

Date	AFP (Normal: 0 - 8.3 ng/mL)
7/21/2005	13.6
1/8/2016	16.3
1/5/2017	36.7
4/19/2018	920.5
6/21/2018	1060.0

What Imaging Should We Order?





ACR Appropriateness Criteria¹

Variant 2:

Penn Medicine

Chronic liver disease. Screening and surveillance for hepatocellular carcinoma (HCC). No prior diagnosis of HCC.

Procedure	Appropriateness Category	Relative Radiation Level	
MRI abdomen without and with IV contrast	Usually Appropriate	0	
MRI abdomen without and with hepatobiliary contrast	Usually Appropriate	0	
US abdomen	Usually Appropriate	0	
CT abdomen with IV contrast multiphase	Usually Appropriate	ବବବବ	
MRI abdomen without IV contrast	May Be Appropriate	0	
MR elastography abdomen	May Be Appropriate	0	
US elastography ARFI abdomen	May Be Appropriate	0	
CT abdomen without IV contrast	Usually Not Appropriate	***	
FDG-PET/CT skull base to mid-thigh	Usually Not Appropriate	****	
1D transient elastography abdomen	Usually Not Appropriate	0	
CT abdomen without and with IV contrast	Usually Not Appropriate	****	

This imaging modality was ordered by the gastroenterologist





Findings (unlabeled)





Findings (unlabeled)



Findings: (labeled)



Impression: New 2.2 cm segment 7 lesion demonstrates arterial phase enhancement and delayed washout with enhancing pseudo-capsule





Additional MRI Characteristics



T1 In phase - Isointense to liver

Diffusion Restriction in ADC, which favors malignancy²



Additional MRI Characteristics





T2 - Isointense to spleen, which suggests a solid lesion²

T2 Out of phase - loses signal, (unusual) which suggests microscopic fat²



DDX (based on imaging)⁷

Figure 2. Differential diagnosis of a solid mass



Micro Path



Penn Medicine

Image of lesion demonstrates thickened trabeculae surrounded by endothelial cells. Individual tumor cells are polygonal with high nuclear cytoplasmic ratio, distinct cell membranes and granular eosinophilic cytoplasm.

Image courtesy of Dr. Rashmi Tondon Department of Pathology and Laboratory Medicine University of Pennsylvania Perelman School of Medicine

(H&E image, 10X magnification)

LI-RADS[®] v2018³

CT/MRI Diagnostic Table

Arterial phase hyperenhancement (APHE)		No APHE		Nonrim APHE		
Observation size (mm)		< 20	≥ 20	< 10	10-19	≥ 20
Count additional major features: Enhancing "capsule" Nonperipheral "washout" • Threshold growth	None	LR-3	LR-3	LR-3	LR-3	LR-4
	One	LR-3	LR-4	LR-4	LR-4 LR-5	LR-5
	≥ Two	LR-4	LR-4	LR-4	LR-5	LR-5



Penn Medicine

Observations in this cell are categorized based on one additional major feature:

• LR-4 – if enhancing "capsule"

• LR-5 – if nonperipheral "washout" **OR** threshold growth



Final Dx:

<u>Hepatocellular Carcinoma (HCC)</u> with subsequent pathology concordance



Likely secondary to cirrhosis from post-Fontan physiology⁴



Case Discussion

- Hepatocellular carcinoma⁵
 - Annual incidence of HCC in the US: 6 per 100,000
 - 4th leading cause of cancer-related death in the world
 - 3.7 times more likely to occur in men than women

Hepatocarcinogenesis:



Portal venous flow

OATP Expression



OATP: Organic anionic transporting polypeptides⁶

RMSER



Case Resolution

Patient underwent Trans-Catheter Arterial Chemoembolization (TACE) in 7/2018



Impression Stable size of the treatment zone in segment 7 with no findings of viable HCC. No new suspicious liver lesion.

Penn Medicine

Date	AFP (Normal: 0-8.3 ng/mL)
6/2018	1060.0
8/2018	412.0
1/2019	19.2
6/2019	18.9
8/2019	20.9

 Currently placed on combined heart (UNOS Status 4) and liver transplant list (with HCC exception).



References:

- Horowitz JM, Kamel IR, Arif-Tiwari H, Asrani SK, Hindman NM, Kaur H, McNamara MM, Noto RB, Qayyum A, Lalani T. ACR Appropriateness Criteria® Chronic Liver Disease. J Am Coll Radiol. 2017 May;14(5S):S103-S117. doi: 10.1016/j.jacr.2017.02.011. Review. PubMed PMID: 28473066.
- 2. Vu, L., Morelli, J., & Szklaruk, J. (2018). Basic MRI for the liver oncologists and surgeons. Journal of Hepatocellular Carcinoma, Volume 5, 37–50. https://doi.org/10.2147/jhc.s154321
- Chernyak, V., Fowler, K. J., Kamaya, A., Kielar, A. Z., Elsayes, K. M., Bashir, M. R., ... Sirlin, C. B. (2018). Liver Imaging Reporting and Data System (LI-RADS) Version 2018: Imaging of Hepatocellular Carcinoma in At-Risk Patients. Radiology, 289(3), 816–830. https://doi.org/10.1148/radiol.2018181494
- Munsterman, I. D., Duijnhouwer, A. L., Kendall, T. J., Bronkhorst, C. M., Ronot, M., ... van Wettere, M. (2018). The clinical spectrum of Fontan-associated liver disease: results from a prospective multimodality screening cohort. European Heart Journal, 40(13), 1057–1068. https://doi.org/10.1093/eurheartj/ehy620
- Global Burden of Disease Liver Cancer Collaboration, Akinyemiju T, Abera S, et al. The Burden of Primary Liver Cancer and Underlying Etiologies From 1990 to 2015 at the Global, Regional, and National Level: Results From the Global Burden of Disease Study 2015. JAMA Oncol. 2017;3(12):1683–1691. doi:10.1001/jamaoncol.2017.3055
- Choi JY, Lee JM, Sirlin CB. CT and MR imaging diagnosis and staging of hepatocellular carcinoma: part I. Development, growth, and spread: key pathologic and imaging aspects. Radiology. 2014 Sep;272(3):635-54. doi: 10.1148/radiol.14132361. Review. PubMed PMID: 25153274; PubMed Central PMCID: PMC4263631.
- 7. Befeler AS. Evaluation of a p;atient with a liver mass. Cancer therapy advisor. https://www.cancertherapyadvisor.com/home/decision-support-in-medicine/gastroenterologyhepatology/evaluation-of-a-patient-with-a-liver-mass/

Penn Medicine

