

Expanding Horizons: Evolving Paradigms in Locoregional Therapy for Recurrent and Residual Hepatocellular Carcinoma



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Introduction

- Recurrence rates of Hepatocellular Carcinoma (HCC) remain up to 70% within 5 years following resection¹ and 62% within 3 years following locoregional therapies (LRT) such as transarterial chemoembolization (TACE).²
- Long-term prognosis remains unsatisfactory, directing research to identify effective approaches for salvage treatment.
- Although recurrent HCC is often managed similarly to de novo disease, retreatment strategies must address the added complexity of prior interventions and altered hepatic reserve.
- This exhibit reviews salvage strategies using LRTs, including TACE, transarterial radioembolization (TARE), and ablation, for patients with residual or recurrent HCC after initial intervention.

Imaging & Assessment

Residual Disease³

- AASLD post-LRT imaging guidelines: Post-TACE – 6 weeks; Post-TARE- 12 weeks; Post-ablation: 4 weeks.
- Persistent enhancement with washout at prior treatment site; use subtraction.
- Treatment: Repeat LRT.
- Typically, there are minimal additional considerations.

Recurrent Disease³

- Long-term imaging every 3-6 months after 2 years.
- Persistent nodular enhancement with washout near or at different site; prior imaging shows nonviable disease.
- Treatment: Reassessment.
- Consider distorted vasculature, prior LRT, liver function.

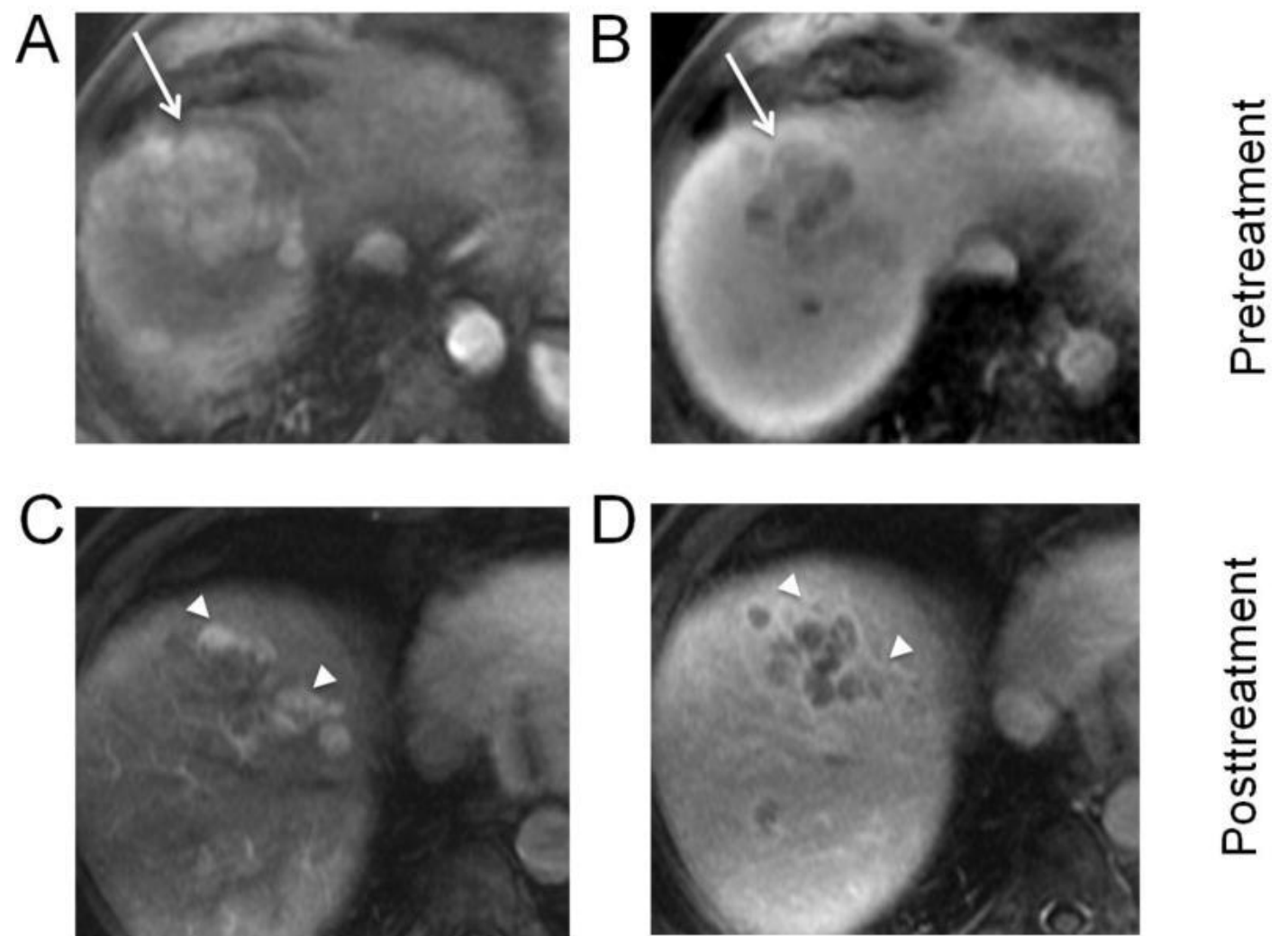


Figure 1: Imaging characteristics on MRI – (A) –Pre-TACE hyperenhancement. (B) – washout. (C) – Post-TACE hyperenhancement. (D) - washout⁴

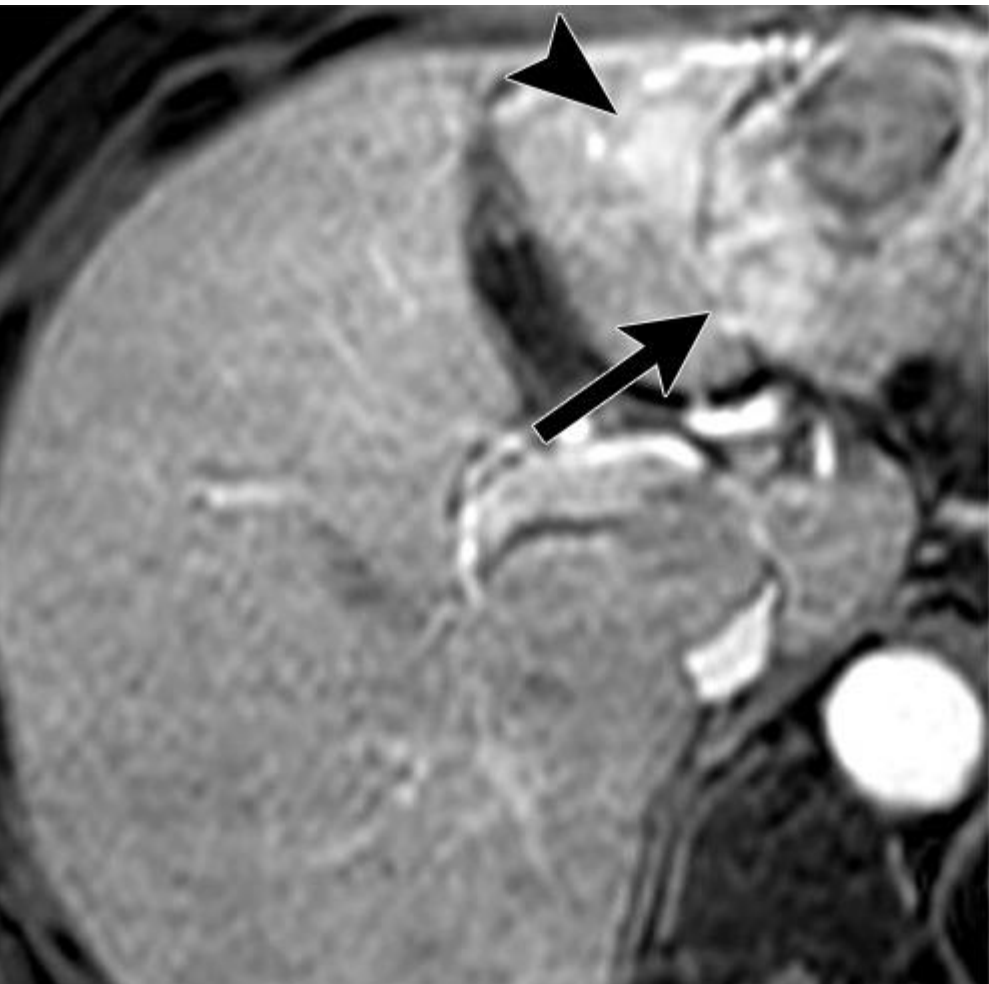


Figure 2: 8 months post-TARE – new washout (not shown) with increased size.⁵

Initial Treatment Selection

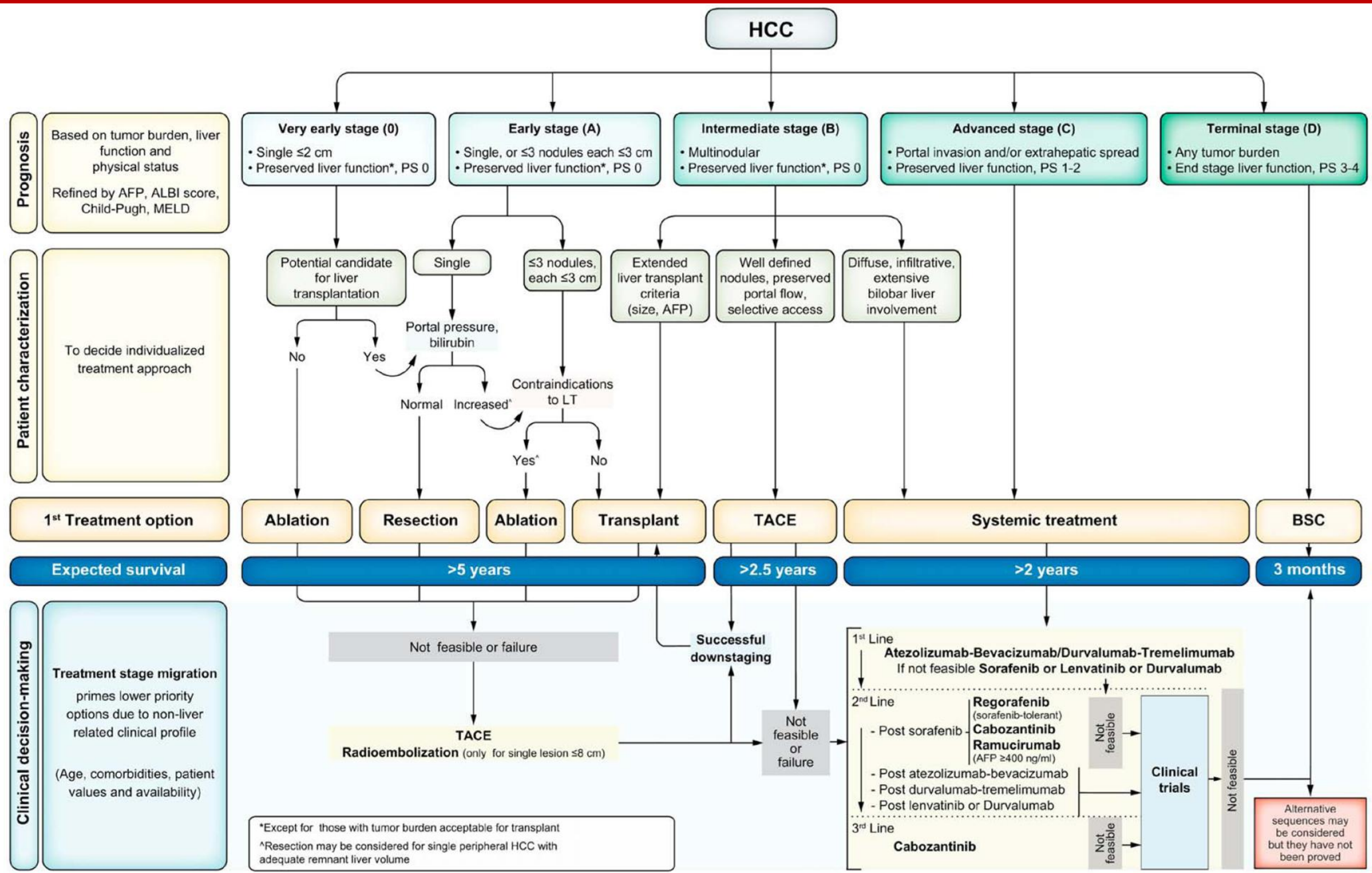


Figure 3: AASLD Treatment Hierarchy detailing initial de novo treatment selection of therapy given classification of HCC tumor.³

Salvage Treatment Selection

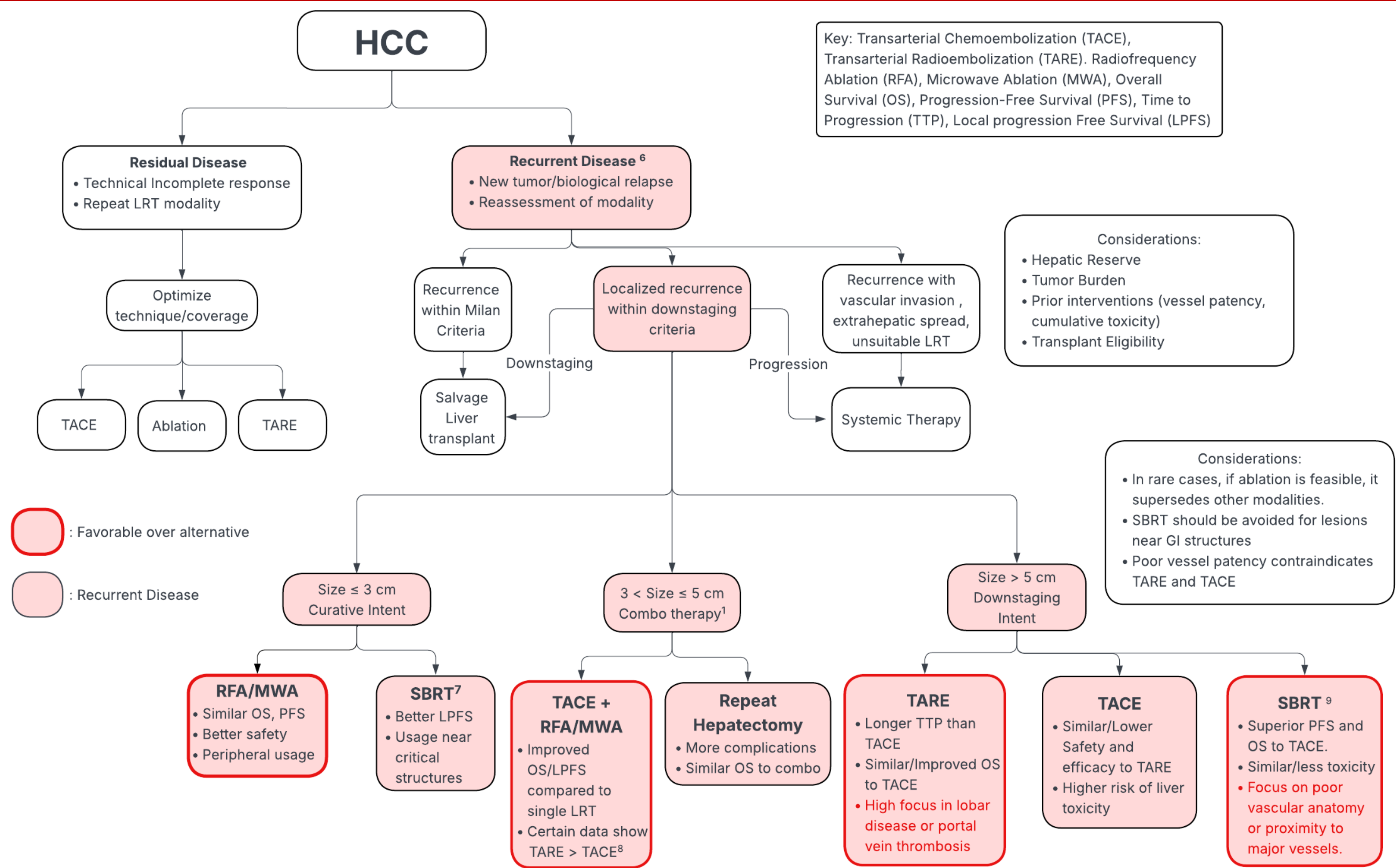


Figure 4: Algorithm for treatment selection in residual vs. recurrent HCC following LRT.

Additional Considerations

- The algorithm assumes that the listed LRTs are technically feasible and equally clinically indicated.
- Absolute contraindications, such as poor hepatic reserve or portal vein thrombosis, should be considered before selecting a modality along with lesion-specific factors, such as tumor location and vascular proximity.
- Therapy should be individualized to maximize tumor control, preserve liver function, and maintain or downstage patients for transplant eligibility.
- Figure 4 serves as a general guideline when multiple locoregional therapy options are feasible and individual patient factors must guide final selection.

Discussion

- Residual disease often warrants repeating the same modality while recurrent disease requires reevaluation and may involve combination therapy, driven by patient and tumor characteristics.
- Ideal treatment strategies for recurrent HCC remain uncertain but LRTs continue to play a central role in tumor control and transplant bridging.
- Multimodal therapies show promise for intermediate lesions and warrant further clinical trials to guide integration into routine practice and optimize protocols.¹
- Integration with systemic therapies and advances in imaging or treatment planning may further improve outcomes.

References

- Criss CR, Makary MS. Salvage locoregional therapies for recurrent hepatocellular carcinoma. *World J Gastroenterol.* 2023;29(3):413-424. doi:10.3748/wjg.v29.i3.413
- Thakurdesai A, Ingawale S, Liu D, et al. Rates and risk factors for recurrence of hepatocellular carcinoma after initial complete remission in response to transcatheter arterial chemoembolization: A systematic review and meta-analysis. *J Clin Oncol.* 2025;43(4_suppl):612-612. doi:10.1200/JCO.2025.43.4_suppl.612
- Singal AG, Llovet JM, Yarrowan M, et al. AASLD Practice Guidance on prevention, diagnosis, and treatment of hepatocellular carcinoma [published correction appears in *Hepatology.* 2023 Dec 1;78(6):E105. doi: 10.1097/HEP.0000000000000621]. *Hepatology.* 2023;78(6):1922-1965. doi:10.1097/HEP.0000000000000466
- Kieler A, Fowler KJ, Lewis S, et al. Locoregional therapies for hepatocellular carcinoma and the new LI-RADS treatment response algorithm. *Abdom Radiol (NY).* 2018;43(1):218-230. doi:10.1007/s00261-017-1281-6
- Mendiratta-Lala M, Masch WR, Shampain K, et al. MRI Assessment of Hepatocellular Carcinoma after Local-Regional Therapy: A Comprehensive Review. *Radiol Imaging Cancer.* 2020;2(1):e190024. Published 2020 Jan 31. doi:10.1148/rycan.2020190024
- Taddei TH, Brown DB, Yarrowan M, Mendiratta-Lala M, Llovet JM. Critical Update: AASLD Practice Guidance on prevention, diagnosis, and treatment of hepatocellular carcinoma. *Hepatology.* 2025;82(1):272-274. doi:10.1097/HEP.0000000000001269
- Xi M, Yang Z, Hu L, et al. Radiofrequency Ablation Versus Stereotactic Body Radiotherapy for Recurrent Small Hepatocellular Carcinoma: A Randomized, Open-Label, Controlled Trial. *J Clin Oncol.* 2025;43(9):1073-1082. doi:10.1200/JCO.24-01532
- de Alcântara JPTL, Götz GWX da R. Transarterial Radioembolization with Yttrium-90 and SIRT Versus Conventional Transarterial Chemoembolization for Hepatocellular Carcinoma: A Systematic Review and Meta-analysis. *Academic Radiology.* Published online June 27, 2025. doi:10.1016/j.acra.2025.06.008
- Su K, Wang F, Li X, et al. Effect of external beam radiation therapy versus transcatheter arterial chemoembolization for non-diffuse hepatocellular carcinoma (≥ 5 cm): a multicenter experience over a ten-year period. *Front Immunol.* 2023;14:1265959. Published 2023 Sep 25. doi:10.3389/fimmu.2023.1265959