Comparison of Hepatic Artery Embolization and Selective Internal Radiation Therapy for Metastatic Neuroendocrine Tumors: A Single-Center Experience

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Background
• Neuroendocrine tumors have a predilection for metastasizing to the liver
• Liver metastases are major determinants of mortality and morbidity
• Liver-directed therapies can effectively ameliorate symptoms and reduce tumor burden
• The effect on survival is less well known
• Few studies have directly compared hepatic transarterial therapies
• Selective internal radiation therapy (SIRT, hepatic radioembolization) is increasingly being used but has not been compared to other modalities

Methods
• Retrospective single-center study (University of Iowa Hospitals and Clinics, UIW)
• The medical records of 42 patients with metastatic neuroendocrine tumor with hepatic metastases who received liver directed therapy were analyzed
• Patient demographics and survival were evaluated
• Chi-Square and Wilcoxon Rank-Sum tests were used to compare the different groups
• Time to progression (TTP) and overall survival (OS) were calculated using the Kaplan-Meier method

Conclusion
• In this cohort there is no significant difference in time to progression between patients treated with SIRT and patients treated with HAE or HACE
• On a subgroup analysis there seems to be a trend towards a longer time to disease progression in patients treated with chemoembolization compared to the other two modalities
• The overall survival after all three liver-directed therapies is relatively long

Summary
• Liver-directed therapies are beneficial in slowing tumor progression in patients who are not candidates for resection
• The optimal liver-directed therapy for metastatic neuroendocrine tumor remains unknown
• Hepatic artery embolization/chemoembolization does not appear to be inferior to selective internal radiation therapy in preventing tumor progression
• Chemoembolization may prolong tumor progression compared to bland embolization or selective internal radiation therapy
• Better understanding of how these therapies compare to one another is warranted

Results

Patient Characteristics
• 42 patients were identified
• All liver-directed therapies were performed at UIHC between 2001-2011
• The median age of the patients was 56.5 years (range 36-81 years)
• 21 women (50%), 21 men (50%)
• 10 patients had extra-hepatic metastases
• All patients were on somatostatin analogues at time of therapy

Table 1: Primary Tumor Location

<table>
<thead>
<tr>
<th>Location</th>
<th>No.</th>
<th>HAE</th>
<th>HACE</th>
<th>SIRT</th>
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<td>Unknown Primary</td>
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Initial Treatment (Fig. 1):
• Hepatic artery embolization (HAE): 31 patients (31%)
• Hepatic artery chemoembolization (HACE): 17 patients (40%)
• Selective internal radiation therapy (SIRT): 12 patients (29%)

Second Treatment (Fig. 2):
• 20 patients had a 2nd procedure
  • HAE: 9 patients (45%)
  • HACE: 5 patients (25%)
  • SIRT: 6 patients (30%)

Overall Survival (OS):
• The median overall survival for the entire group from the first intervention was 41.9 months (Fig. 3)
• The 5-year overall survival from the first intervention for the entire group was 45%
• There was no significant difference in median overall survival among patients in different treatment groups (HAE 21.3 months, HACE 68.7 months, SIRT 26.8 months) (p=0.300)

Time to Progression (TTP):
• The median TTP after the first intervention was 19.2 months (Fig. 4)
• There was no significant difference in time to progression after the first intervention between SIRT (15.1 months) and HAE or HACE (19.6 months) (p=0.968) (Fig. 5)
• There was a trend towards an increased time to progression in HACE (33.4 months), compared to HAE (12.1 months) or SIRT (15.1 months) (p=0.512) (Fig. 6)

Figure 1: Type of 1st Procedure

Figure 2: Type of 2nd Procedure

Figure 3: Overall Survival

Figure 4: Overall TTP

Figure 5: TTP: SIRT vs HACE

Figure 6: TTP: SIRT vs HACE with different solid lines.