Radiological Evaluation of Hepatocellular Carcinoma

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Abstract
Introduction: Hepatocellular carcinoma is the fifth most common cancer worldwide and the second leading cause of cancer-related death. More than 7 lakh people die of HCC each year1,2. Imaging and histopathology plays a very important role in the diagnosis of HCC.

Materials & Methods: This is an observational study conducted over a period of one year, from September 2016 to October 2017 on 36 patients aged between 20-80yrs with sonologically detected hepatic mass lesions suspicious for HCC. They were evaluated with 16 slice CECT and 1.5T MRI. After taking informed consent, USG guided Biopsy/FNAC was done from these lesions and only those lesions that showed HCC were included in the study. The lesion characteristics are retrospectively analysed for the presence of specific features on US, CECT, MRI.

Results: Majority of the cases occurred in the age group of 51-60 years (38.88%), and there is a male preponderance (83.33%). Half the patients are cirrhotic (52.78%). Hepatomegaly is observed in 55.56% cases. 50% of the cases measured 5-10 cm in largest dimension. Majority of the lesions are hyperechoic and show intralosomal vascularity. Majority of the lesions showed enhancement in the arterial phase of the CECT, followed by washout in the venous phase. Majority of them showed a well-defined capsule around the lesion, are hypointense on T1 and Hyperintense on T2, showing a mosaic architecture. Two thirds of the patients tested positive for Hepatitis B, deranged liver function tests and increased Alpha feto protein levels.

Conclusion: HCC can occur in a wide range of age groups. Though majority of the lesions show similar characteristics, HCC can have a wide spectrum of radiological appearances, and hence no single feature can be pathognomonic of the diagnosis. Hence, a tissue diagnosis is always desirable, whenever there is high index of suspicion, in lesions showing atypical pattern of imaging features.

Introduction
Hepatocellular carcinoma (HCC) is the fifth most common cancer worldwide and the second leading cause of cancer-related death. More than 7 lakh people die of HCC each year1,2. It occurs primarily in subjects who have chronic liver disease or liver cirrhosis3 and is the primary cause of death among this group. The high-risk group includes patients testing positive for Hepatitis B, Hepatitis C, or those suffering with cirrhosis4. HCC usually occurs in the setting of cirrhosis3, although this is not always the case, with the classic example being HCC associated with hepatitis B in the noncirrhotic liver. Several imaging diagnostic systems have been proposed for the interpretation of liver lesions in at-risk patients for HCC. Although these imaging-based diagnostic systems represent important advances, they have limitations and they are not perfectly consistent with each other5. Unfortunately, despite numerous
technological developments and improvements in recent years, the diagnostic accuracy of non-invasive imaging in patients with cirrhosis is still relatively low, ranging respectively between 0.60 and 0.72 for USG, between 0.74 and 0.83 for MDCT and between 0.71 and 0.87 for MRI.\textsuperscript{1-13} The relatively poor diagnostic performance for the detection of HCC in cirrhotic liver is due principally to overlapping imaging features and, thus, difficulties in differentiating dysplastic nodules from small HCC, and to problems associated with diagnosing arterially enhancing nodules smaller than 2 cm in diameter. For these reasons patients generally undergo regular follow-up with CT or MRI.

**Aims and Objectives**
1) To study the radiological behaviour of HCC
2) To study the incidence of specific imaging features on Ultrasound, MDCT and MRI in pathologically proven cases of Hepatocellular carcinoma.

**Materials and Methods**

**Study Design:** Observational study
**Study Period:** One year from September 2016 to October 2017, at department of Radiodiagnosis, Andhra Medical College, Visakhapatnam.

**Inclusion Criteria**
The study group included patients with normal renal function (serum creatinine<1.4mg/dl) with
1. Clinical suspicion of focal liver lesions.
2. Previous imaging studies depicted focal liver lesions with nonspecific appearance.

**Exclusion Criteria**
1. Patients with imaging features characteristic of benign hepatic lesion
2. Pregnancy
3. Acute or chronic renal failure
4. History of allergy to contrast media
5. Inability to tolerate sufficient breath hold.
6. Patients with pace makers or implants. Sample size: 36 cases.

**Results**
The youngest patient diagnosed with HCC in our study is 25 years and the oldest is 81 years. The average age of the patient at diagnosis is 54 years. Majority of the cases occurred in the age group of 51-60 years (38.88%), followed by 61-70 years (25%) and 41-50 years (17.94%) in that order, of which 30 persons (83.33%) are male and the rest female. Cirrhosis is noted in 19 (52.78%) of the 36 patients, of which 2 are females and 17 are males. The youngest person with cirrhosis, diagnosed with HCC, is 40 years. Hepatomegaly is found in 20 people (55.56%), more so with larger lesions. Of all the lesions studied, no lesion showed intralesional calcification. Majority of the lesions studied measure between 5-10 cm in largest dimension, shown by 18 lesions, corresponding to 50%. 8 lesions (22.22%) measured < 5 cm and 7 lesions (19.44%) measured 10-15 cm. The average size of the lesion is 7.986 cm in largest dimension. Of the 36 lesions studied, 25 lesions, corresponding to 69.44% are hyperechoic, 9 lesions, corresponding to 25% are mixed echogenic and 2 lesions, corresponding to 5.56% are predominantly hypoechoic. Intralesional vascularity is seen in 30 persons, accounting for 83.33% and Peri-lesional halo could be demonstrated in 26 cases, corresponding to 72.22%. Intralesional necrosis is seen in 15 cases (41.67%). 7 persons (19.44%) have portal vein thrombosis. In this study, 35 lesions (97.22%) showed enhancement in arterial phase and one lesion showed no enhancement (2.78%). Venous phase washout was seen in 32 of the 35 lesions that showed enhancement in the arterial phase (91.43%). While 3 lesions (8.57%) did not show wash out in the venous phase. A well-defined delayed enhancing capsule could be demonstrated in 28 lesions (77.78%). 34 lesions (94.44%) showed T1 isointense signal, one (2.77%) showed T1 Hyperintense signal and one (2.77%) remained isointense. 35 lesions (97.22%) showed T2 Hyperintense signal while one lesion (2.77%) showed hypo intensity on T2. 28 lesions (77.78%) showed mosaic architecture while 8 (22.22%) lesions did not.
Table 1: Frequency of Cirrhosis

<table>
<thead>
<tr>
<th></th>
<th>Male</th>
<th>Female</th>
<th>Total</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cirrhotic</td>
<td>17</td>
<td>2</td>
<td>19</td>
<td>52.78%</td>
</tr>
<tr>
<td>Non-Cirrhotic</td>
<td>13</td>
<td>4</td>
<td>17</td>
<td>47.22%</td>
</tr>
<tr>
<td>Total</td>
<td>30</td>
<td>6</td>
<td>36</td>
<td>100%</td>
</tr>
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</table>

Table 2: Pattern of Enhancement

<table>
<thead>
<tr>
<th>Pattern of Enhancement</th>
<th>Frequency</th>
<th>%</th>
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</thead>
<tbody>
<tr>
<td>None</td>
<td>1</td>
<td>2.78%</td>
</tr>
<tr>
<td>Arterial</td>
<td>35</td>
<td>97.22%</td>
</tr>
<tr>
<td>Total</td>
<td>36</td>
<td>100%</td>
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</table>

Table 3: Wash out Pattern

<table>
<thead>
<tr>
<th>Wash out pattern</th>
<th>Frequency</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Venous phase</td>
<td>32</td>
<td>91.43%</td>
</tr>
<tr>
<td>NA</td>
<td>3</td>
<td>8.57%</td>
</tr>
<tr>
<td>Total</td>
<td>35</td>
<td>100%</td>
</tr>
</tbody>
</table>

Table 4: T1 & T2 Signal Characteristics

<table>
<thead>
<tr>
<th>Signal</th>
<th>T1</th>
<th>T2</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hypo intense</td>
<td>34</td>
<td>1</td>
</tr>
<tr>
<td>Hyperintense</td>
<td>1</td>
<td>35</td>
</tr>
<tr>
<td>Isointense</td>
<td>1</td>
<td>0</td>
</tr>
<tr>
<td>Total</td>
<td>36</td>
<td>36</td>
</tr>
</tbody>
</table>

Figure 1: Ultrasound examination shows a well-defined hypoechoic perilesional halo in this 51-year-old male.
Figure 2: CECT in a 70-year-old man shows arterial phase enhancement of the lesion with subsequent washout in the venous phase.

Figure 3: CECT in a 30-year-old male shows a well-defined capsule showing enhancement in the delayed phase.

Discussion
HCC can occur in a wide range of age groups including young patients. This is in contrast to the general opinion that HCC is a disease of the adults and elderly. The youngest patient in our study is 25 years old. The average age at diagnosis is 54 years. Cirrhosis is a known risk factor for the development of HCC, which is reflected in our study with 52.78% of the study group being cirrhotic. A male preponderance is noted with the disease, probably due to other risk factors like alcoholism, that predispose to Cirrhosis and eventually HCC. Hepatomegaly was observed with 55.56% of the cases, which doesn’t correlate with the size of the tumour or with the no. of lesions. Majority of the lesions studied measure
between 5-10 cm in largest dimension, shown by 18 lesions, corresponding to 50%. 8 lesions (22.22%) measured < 5 cm and 7 lesions (19.44%) measured 10-15 cm. The average size of the lesion is 7.986 cm in largest dimension. Of the 36 lesions studied, 25 lesions (69.44%) are hyperechoic, 9 lesions (25%) are mixed echogenic and 2 lesions (5.56%) are predominantly hypoechoic. Intralesional vascularity is seen in 83.33%, as detected by color doppler. Intralesional necrosis is noted in 41.67% of the cases which occurs when the tumour outgrows its vascular supply. Karahan O14 showed that the portal vein thrombosis was more frequently associated with lesions larger than 10 cm, whereas in our study, portal vein thrombosis was seen in the tumour sizes ranging from 4.4 cm to 10 cm. Peri-lesional halo could be demonstrated in 26 cases, corresponding to 72.22%. Of all the lesions studied, no lesion showed intralesional calcification, which is in contradiction to several studies, which found an incidence of intralesional calcifications, ranging from 3-25%15,16. In this study, 35 lesions (97.22%) showed enhancement in arterial phase and one lesion showed no enhancement (2.78%). Venous phase washout was seen in 32 of the 35 lesions that showed enhancement in the arterial phase, corresponding to 91.43%. While 3 lesions, corresponding to 8.57% did not show washout in the venous phase. These findings, are in close agreement with the findings made by K.H.Y. Lee (2003)17, who have found that hypervascular component was seen in 96% of the cases, 86% of the lesions showed arterial phase enhancement and 76% of the lesions showed washout in the venous phase. Baron et al14 in their study on 66 patients found that 39% of the lesions are hyper attenuating in the arterial phase, another 39% are showing mixed attenuation and 21% are hypo attenuating. A well-defined delayed enhancing capsule could be demonstrated in 28 lesions (77.78%). 34 lesions (94.44%) showed T1 isointense signal, one (2.77%) showed T1 Hyperintense signal and one (2.77%) remained isoointense. 35 lesions (97.22%) showed T2 Hyperintense signal while one lesion (2.77%) showed hypo intensity on T2.28 lesions (77.78%) showed mosaic architecture while 8 (22.22%) lesions did not. SGOT levels are raised in 100% of the patients while SGPT levels are raised in 83.33%. Raised AFP values are noted in 35 cases (97.22%), of which, 33 cases (91.66%) showed a value >1000 ng/ml. Normal value could be observed in only one case.

Conclusion
HCC can present over a wide range of age groups and can have a wide spectrum of radiological appearances. Hence, no single feature can be pathognomonic of the diagnosis. However, the most consistent findings in our study include:

**USG:** hyperechoic mass lesion with intralesional vascularity and a peripheral halo.

**CECT:** lesion showing enhancement in the arterial phase and washout in the venous phase with a well-defined capsule that enhances in the delayed phase and showing mosaic architecture.

**MRI:** T1 hypo intensity, T2 hyperintensity, with mosaic architecture. Biochemistry: Elevated AFP with deranged LFTs.

However, a tissue diagnosis is always desirable, whenever there is high index of suspicion, in lesions showing atypical pattern of imaging features.

**References**