#### Outcomes of liver lesions categorized using Liver Imaging Reporting and Data System (LI-RADS) in patients who have undergone previous hepatic resection for Hepatocellular Carcinoma.

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# **Declaration:**

• The authors have nothing to declare.

# **Background (1)**

- Hepatocellular Carcinoma (HCC) is the third leading cancer deaths worldwide accounting for nearly 700,000 deaths annually.
- Current guidelines allow diagnosis of HCC non-invasively using cross-sectional imaging without the need for histological confirmation.
- However, characteristic contrast enhancement pattern is only present in around half of all HCC.

## **Background (2)**

- For lesions considered indeterminate, radiologists have no systematic way to convey diagnostic uncertainty.
- Liver Imaging Reporting and Data System (LI-RADS) was introduced to provide a stratified probability of the full spectrum of hepatic lesions and pseudo-lesions and characterised them into several categories.
- However, most studies thus far concentrate on treatment naïve patients and little is known about its diagnostic efficacy patients who had undergone previous resection.

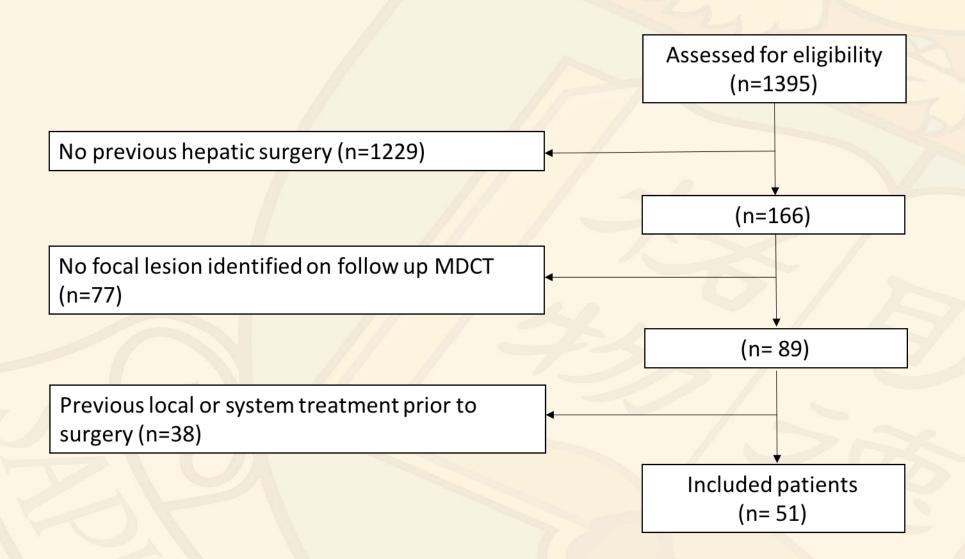
### Purpose

 To assess the inter-reader agreement and outcomes of focal liver lesions identified on computed tomography (MDCT) and characterized using LI-RADS verion 2017 in patients who had undergone previous hepatic resection for Hepatocellular Carcinoma (HCC).

### **Materials and Methods**

- Retrospective review of reports from radiological database between 2010-2015
- Patients undergone previous curative resection.
- Two abdominal radiologists independently reviewed lesions reported on follow-up CT scans using LI-RADS version 2017 categorization
- Compared with histological confirmation or long term follow-up.

#### **Results**



### **Patient Characteristics**

|  | Patients with HCC recurrence in     | Patients with no recurrence in    |  |  |
|--|-------------------------------------|-----------------------------------|--|--|
|  | indexed lesions (N=20)              | indexed lesions (N=31*)           |  |  |
| M:F  | 16:4                                | 25:6                              |  |  |
| Median Age (range)   | 58 (45-89)                          | 59 (46-78)                        |  |  |
| Cause of liver disease:  |                                     |                                   |  |  |
| • HBV  | 17 (85%)                            | 24 (77%)                          |  |  |
| • HVC  | 2 (10%)                             | 3 (10%)                           |  |  |
| Others   | 1 (5%)                              | 4 (13%)                           |  |  |
| Serum AFP  |                                     |                                   |  |  |
| • Median (ng/ml)   | 15 (2-26160)                        | 4 (2-212)                         |  |  |
| <ul> <li>Raised (&gt;20ng/ml)</li> </ul>                       | 9 (45%)                             | 4 (13%)                           |  |  |
| *6 patients had HCC recurrence subs<br>2-year overall survival | equently although the indexed lesic | ons were considered to be benign. |  |  |

### Per lesion comparison of LI-RADS Category, AASLD criteria and final diagnosis

| Consensus LI-RADS<br>categorization | No of lesions<br>(N=98) | HCC as per AASLD<br>criteria | HCC as final diagnosis |  |
|-------------------------------------|-------------------------|------------------------------|------------------------|--|
| 1                                   | 17                      | 0                            | 0                      |  |
| 2                                   | 5                       | 0                            | 0                      |  |
| 3                                   | 50                      | 0                            | 13                     |  |
| 4                                   | 18                      | 8                            | 14                     |  |
| 5 / TIV                             | 8                       | 8                            | 8                      |  |

### Number of Observations in Each LI-RADS category between 2 reviewers

|            |                      | Reviewer 2 |   |    |    |         |       |
|------------|----------------------|------------|---|----|----|---------|-------|
|            | LR<br>Categorisation | 1          | 2 | 3  | 4  | 5 / TIV | Total |
| Reviewer 1 | 1                    | 15         | 4 | 0  | 0  | 0       | 19    |
|            | 2                    | 0          | 2 | 0  | 0  | 0       | 2     |
|            | 3                    | 0          | 2 | 47 | 1  | 0       | 50    |
|            | 4                    | 0          | 0 | 6  | 13 | 0       | 19    |
|            | 5 / TIV              | 0          | 0 | 0  | 0  | 8       | 8     |
|            | Total                | 15         | 8 | 53 | 14 | 8       | 98    |

### **Per lesion analysis**

- 35 (35.7%) HCC and 63 (64.3%) non-malignant
- 17/35 (51.4%) HCC have arterial hypervascularity and contrast washout on portovenous or delayed imaging
- 9/35 (25.7%) HCCs <10mm, 14/35 (40.0%) between 10-20 mm and 11/35 (34.3%) >20mm
- Median axial diameters of HCC were larger than those of nonmalignant lesions (15 vs 8 mm p<0.0001).

### **Performance of LI-RADS**

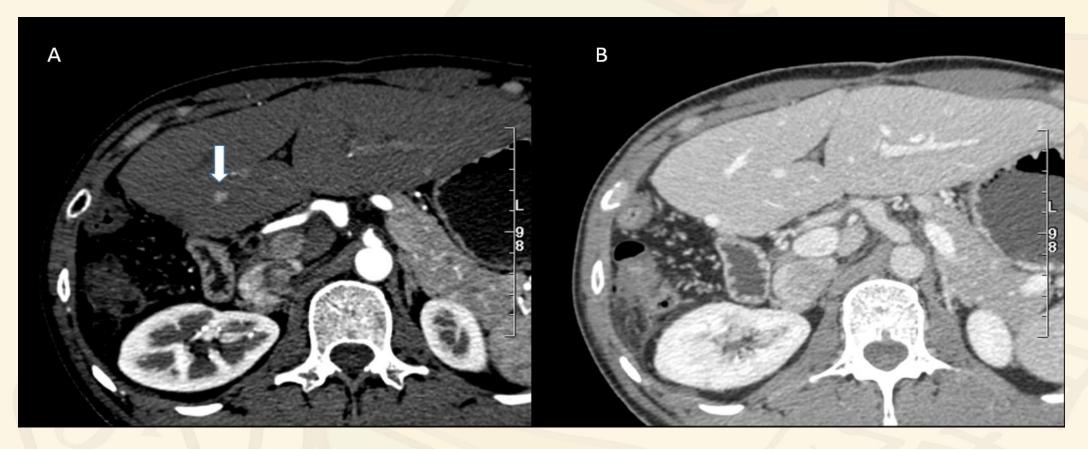
- All 8/8 (100%) LR-TIV & LR-5, 14/18 (77.8%) LR-4, and 13/50 (26.0%) LR-3 were HCC.
- All 22/22 (100%) LR-2 and LR-1 lesions were benign.
- All 13 LR-3 HCC were <20 mm.

### **LR-4 benign lesion**



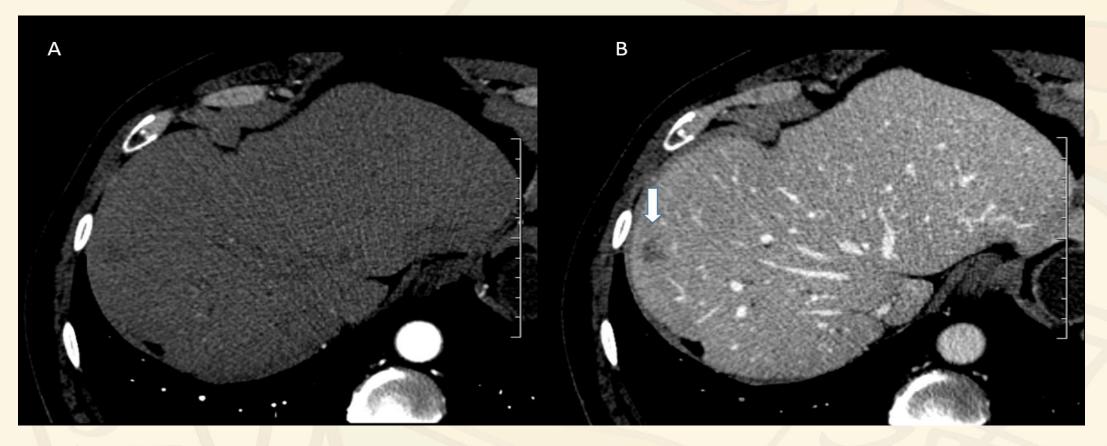
69 year-old woman with hepatitis B who underwent right hepatectomy for HCC. (A) Arterially hyperenhancing observation (40 mm) is seen on arterial phase CT (arrow), but is not noted to washout or have capsule on portovenous (B) or delayed phase CT. Both reviewers considered this lesion to be LR-4 but did not fulfil the AASLD criteria for HCC. (C) This lesion was not seen on follow-up imaging in 12 months.

## **LR-3 HCC (1)**



43 year-old man with hepatitis B and previous right hepatectomy for HCC. (A) Arterially hyperenhancing observation (6 mm) is seen on arterial phase CT (arrow), with no contrast washout on portovenous (B) and delayed phase CT. The lesion was resected and histologically confirmed as a HCC. Both reviewers assigned this lesion as LR-3 and non-diagnostic of HCC based on AASLD criteria.

### **LR-3 HCC (2)**



59 year-old man with hepatitis B and previous wedge resection for HCC. (A) The lesions (14 mm) is isodense / faintly hypodense on arterial phase CT, but is noted to have contrast washout on portovenous (B) and delayed phase CT. The lesion was considered to be LR-3 and not diagnostic of HCC by AASLD criteria. The lesion was resected and histologically confirmed to be a HCC.

#### **Inter-rater agreement**

- Very good agreement between the two reviewers using the AASLD criteria, k = 0.851 (95%CI 0.708-0.994 p<0.0001) and good agreement using LI-RADS categorization, k = 0.799 (95%CI 0.699-0.899 p<0.0001).</li>
- 13 discordance using LI-RADS categorization, 4 between LR-1 and 2, 2 between LR-2 and 3, 7 between LR-3 and 4. None were more than one categorization.

## **Discussion (1)**

- Our results shows that both the AASLD criteria and LR-5 categorization have high specificity in diagnosing HCC
- However, the sensitivities of both were low due to their restricting diagnostic criteria.
- By not distinguishing LR-4 and LR-5, significantly higher sensitivities for diagnosing HCC can be achieved while maintaining excellent specificities

# **Discussion (2)**

- 26% (13/50) LR-3 lesions in our cohort were HCC, higher than other published studies. This could be due to the very high risks patient cohort in our study compared to the general population.
- 31% (13/42) LR-3 lesions in our cohort were not consistently visualized on follow-up imaging (psuedolesions).

### Conclusion

- LI-RADS categorization has good inter-reader agreement although LR-5 alone is too restrictive for diagnosis of HCC.
- Sensitivity can be significantly increased when LR-4 and LR-5 categorization were combined.
- A significant proportion of LR-3 lesions were HCC and accelerated follow-up for these lesions is necessary to distinguish them from benign aetiology.

### References

- J. Ferlay, H.R. Shin, F. Bray, D. Forman, C. Mathers, D.M. Parkin, Estimates of worldwide burden of cancer in 2008: GLOBOCAN 2008, Int J Cancer 127(12) (2010) 2893-917.
- L. European Association For The Study Of The, R. European Organisation For, C. Treatment Of, EASL-EORTC clinical practice guidelines: management of hepatocellular carcinoma, J Hepatol 56(4) (2012) 908-43.
- A.C.o. Radiology, Liver Imaging Reporting and Data System version 2017., 2017. http://www.acr.org/Quality-Safety/Resources/LIRADS. (Accessed 1/8/2017.
- D.G. Mitchell, M.R. Bashir, C.B. Sirlin, Management implications and outcomes of LI-RADS-2, -3, -4, and -M category observations, Abdom Radiol (NY) (2017).
- A. Darnell, A. Forner, J. Rimola, M. Reig, A. Garcia-Criado, C. Ayuso, J. Bruix, Liver Imaging Reporting and Data System with MR Imaging: Evaluation in Nodules 20 mm or Smaller Detected in Cirrhosis at Screening US, Radiology 275(3) (2015) 698-707.
- D.G. Mitchell, J. Bruix, M. Sherman, C.B. Sirlin, LI-RADS (Liver Imaging Reporting and Data System): summary, discussion, and consensus of the LI-RADS Management Working Group and future directions, Hepatology 61(3) (2015) 1056-65.
- I. Joo, J.M. Lee, D.H. Lee, S.J. Ahn, E.S. Lee, J.K. Han, Liver imaging reporting and data system v2014 categorization of hepatocellular carcinoma on gadoxetic acid-enhanced MRI: Comparison with multiphasic multidetector computed tomography, J Magn Reson Imaging 45(3) (2017) 731-740.
- M. Tanabe, A. Kanki, T. Wolfson, E.A.C. Costa, A. Mamidipalli, M.P.F.D. Ferreira, C. Santillan, M.S. Middleton, A.C. Gamst, Y. Kono, A. Kuo, C.B. Sirlin, Imaging Outcomes of Liver Imaging Reporting and Data System Version 2014 Category 2, 3, and 4 Observations Detected at CT and MR Imaging, Radiology 281(1) (2016) 129-139.
- M.T. Corwin, G. Fananapazir, M. Jin, R. Lamba, M.R. Bashir, Differences in Liver Imaging and Reporting Data System Categorization Between MRI and CT, AJR Am J Roentgenol 206(2) (2016) 307-12.
- B.K. Barth, O.F. Donati, M.A. Fischer, E.J. Ulbrich, C.A. Karlo, A. Becker, B. Seifert, C.S. Reiner, Reliability, Validity, and Reader Acceptance of LI-RADS-An In-depth Analysis, Acad Radiol 23(9) (2016) 1145-53.