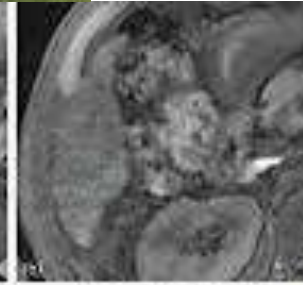
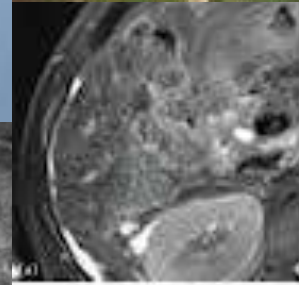
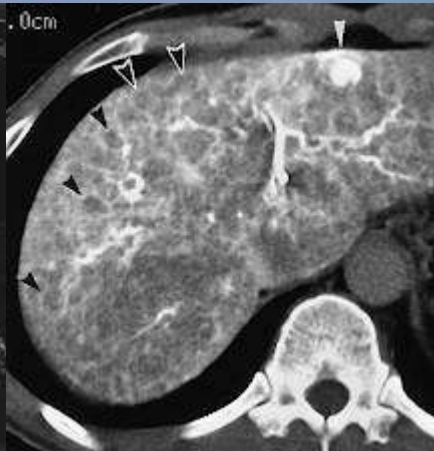
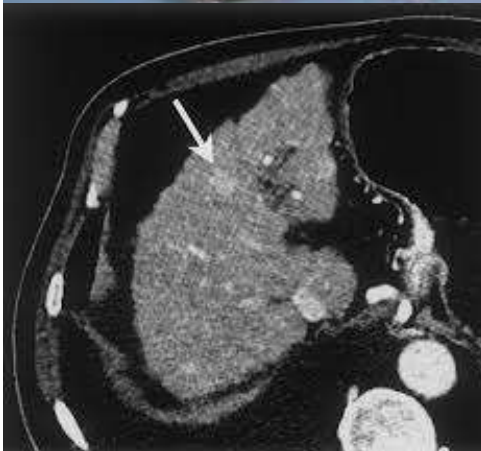


Challenges in diagnosis of hepatocellular carcinoma

Do Young Kim

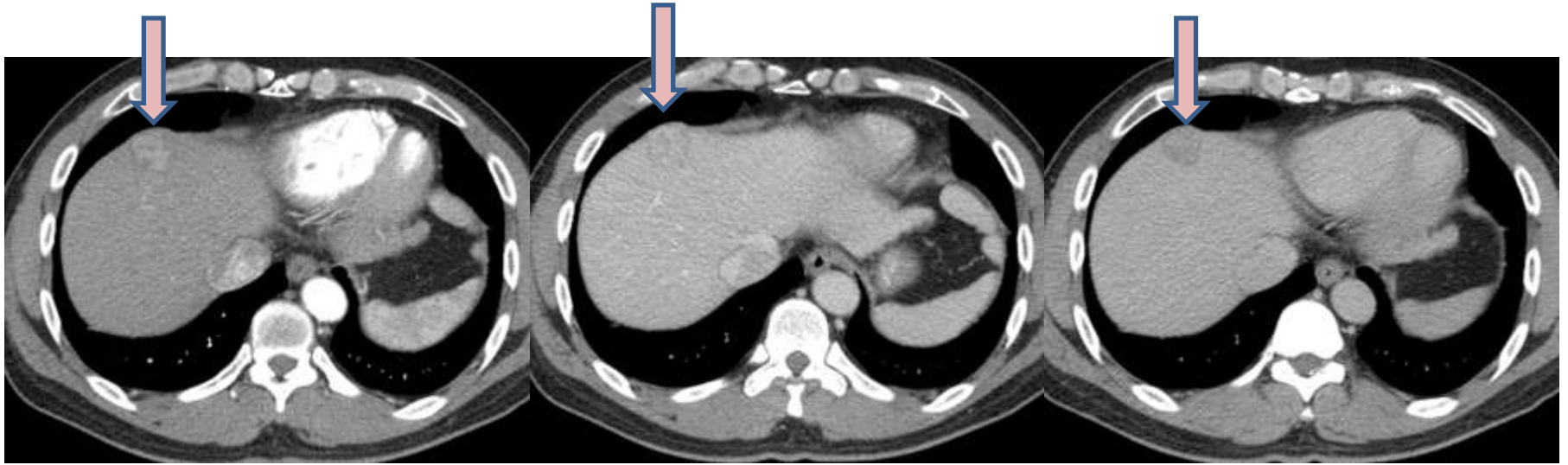
Department of Internal Medicine,
Yonsei University College of Medicine

Liver nodule



Diagnosis of HCC: Typical imaging findings

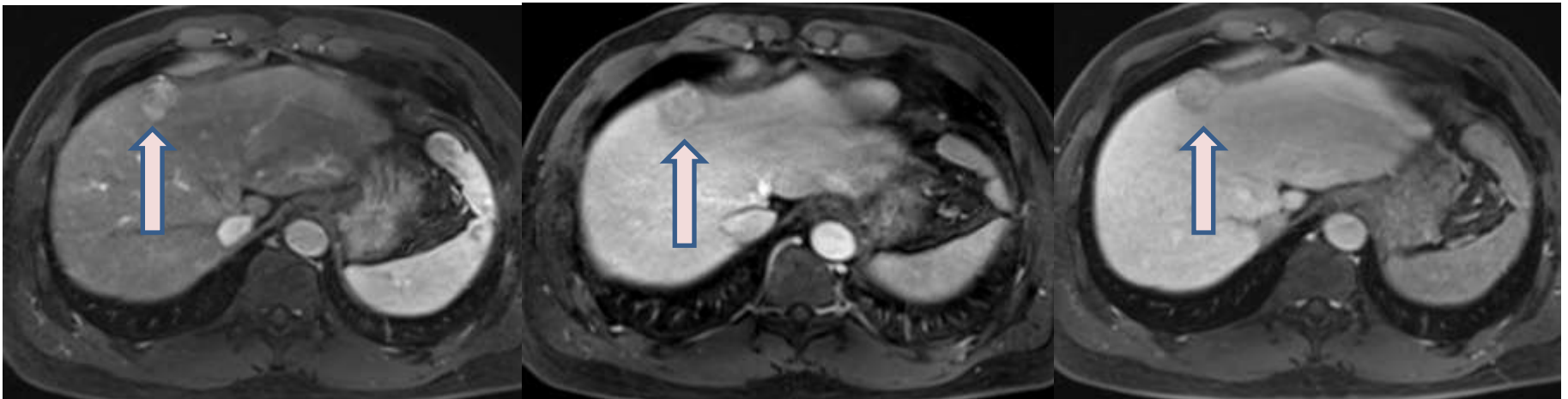
- Increased enhancement on arterial phase
- Decreased enhancement (**Washout**) on delayed or equilibrium phase



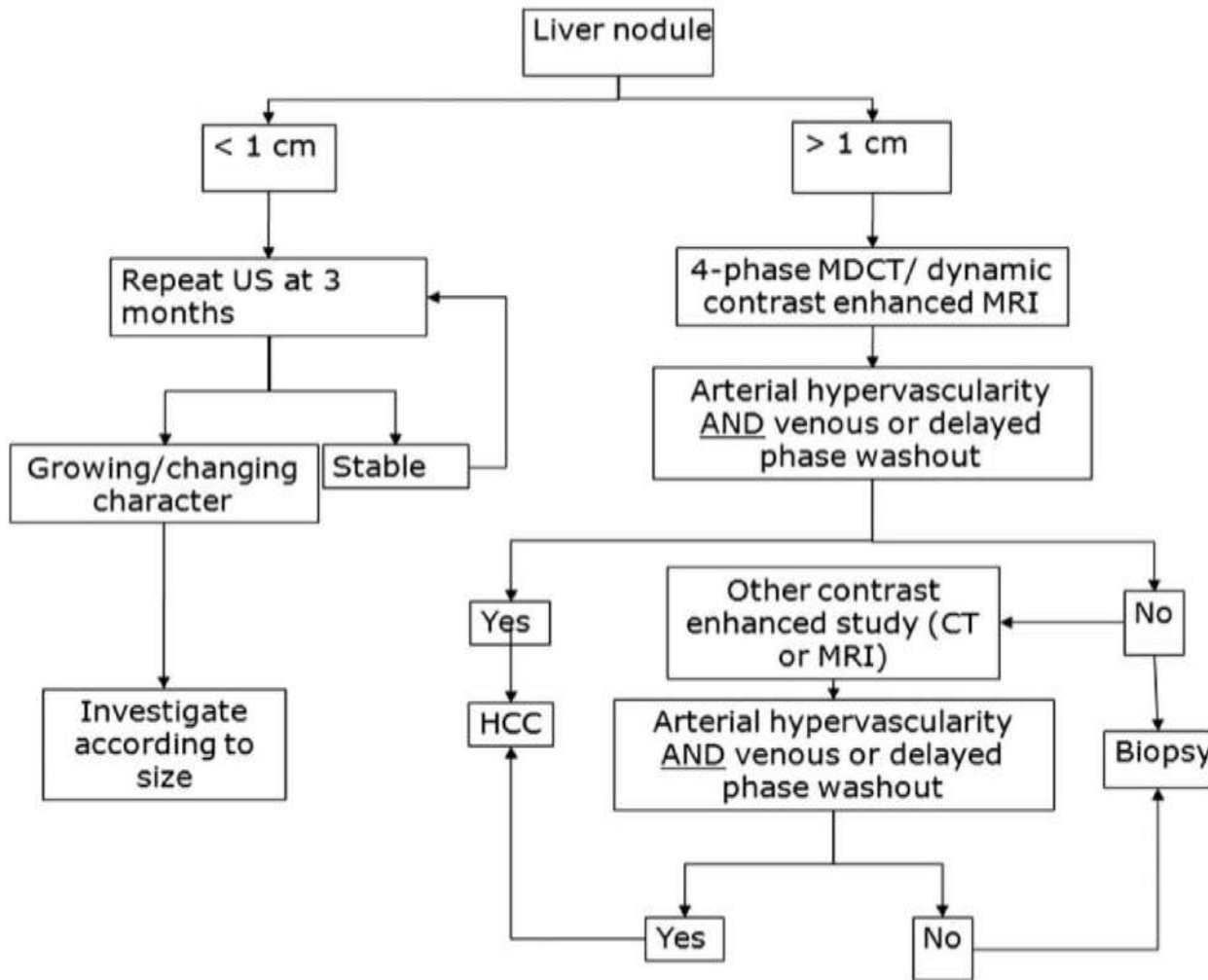
Arterial Phase

Portal Phase

Delayed Phase

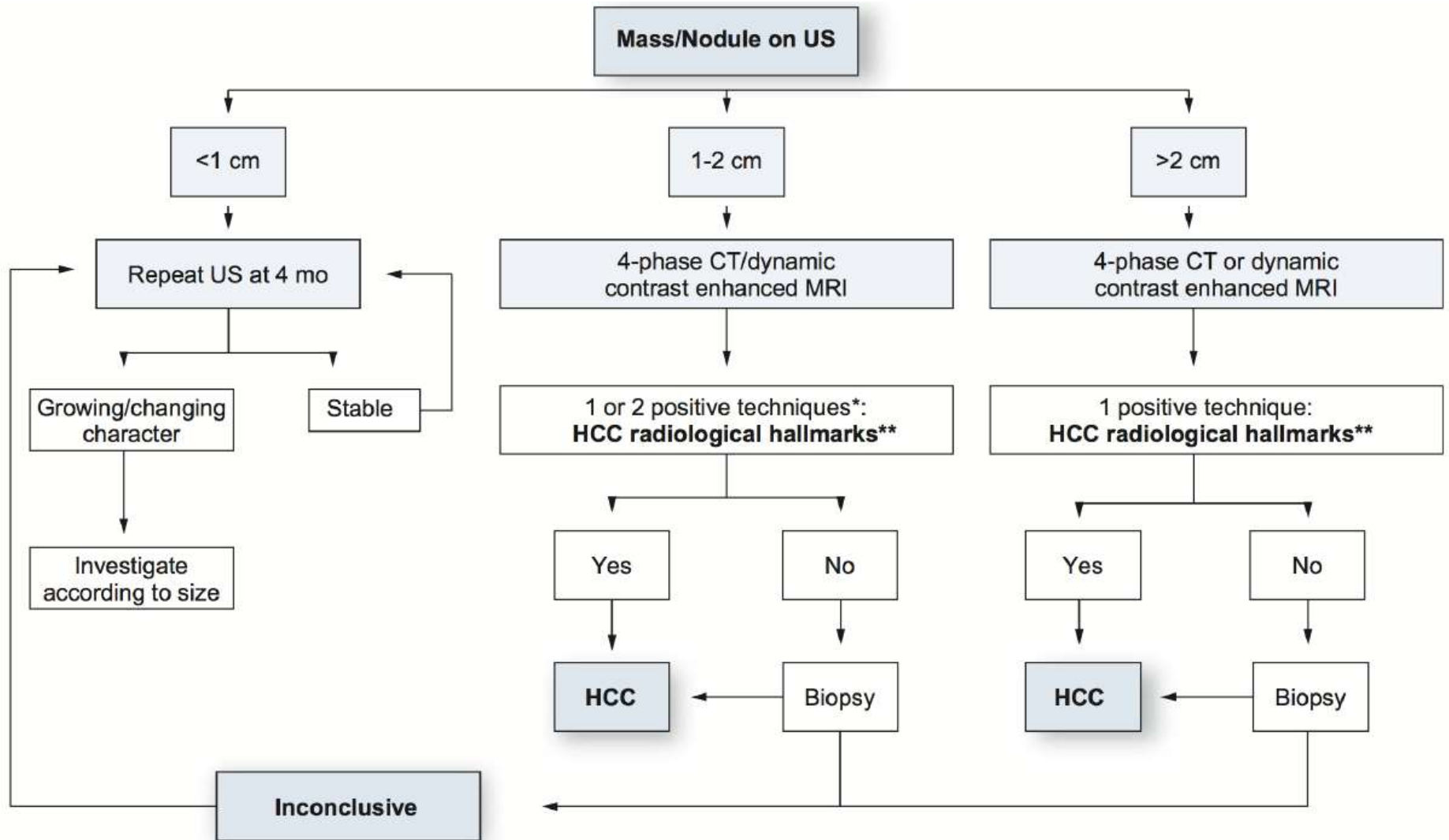


AASLD 2010 Update



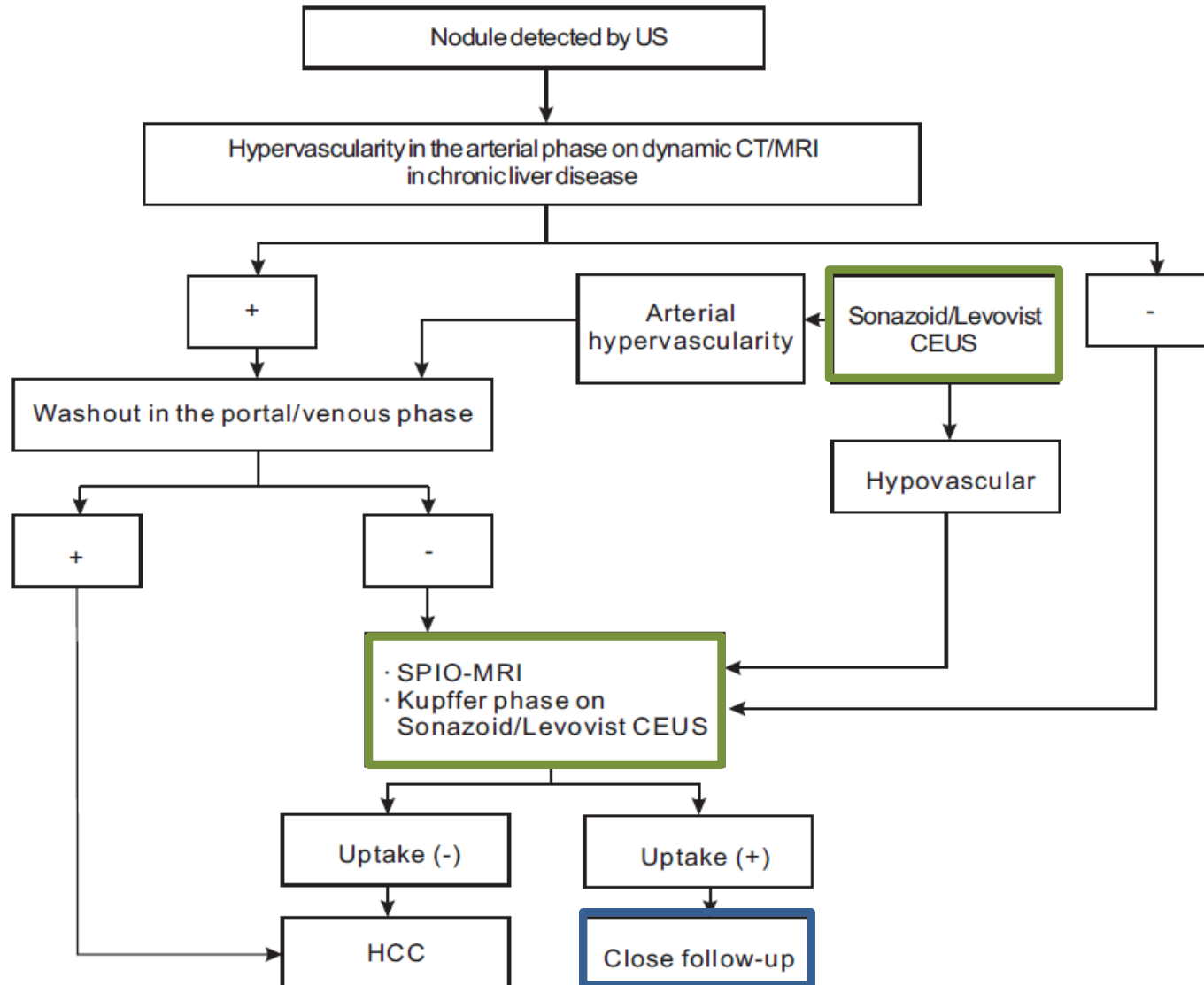
- **Typical pattern(<2cm)**
→ **61.7% sensitivity**
96.6% specificity
- **HCC washout**
→ **< 1 cm: 27%**
>1, <1.5 cm: 39%
>1.5,<2 cm: 50%
>2cm: 82%

EASL-EORTC 2011



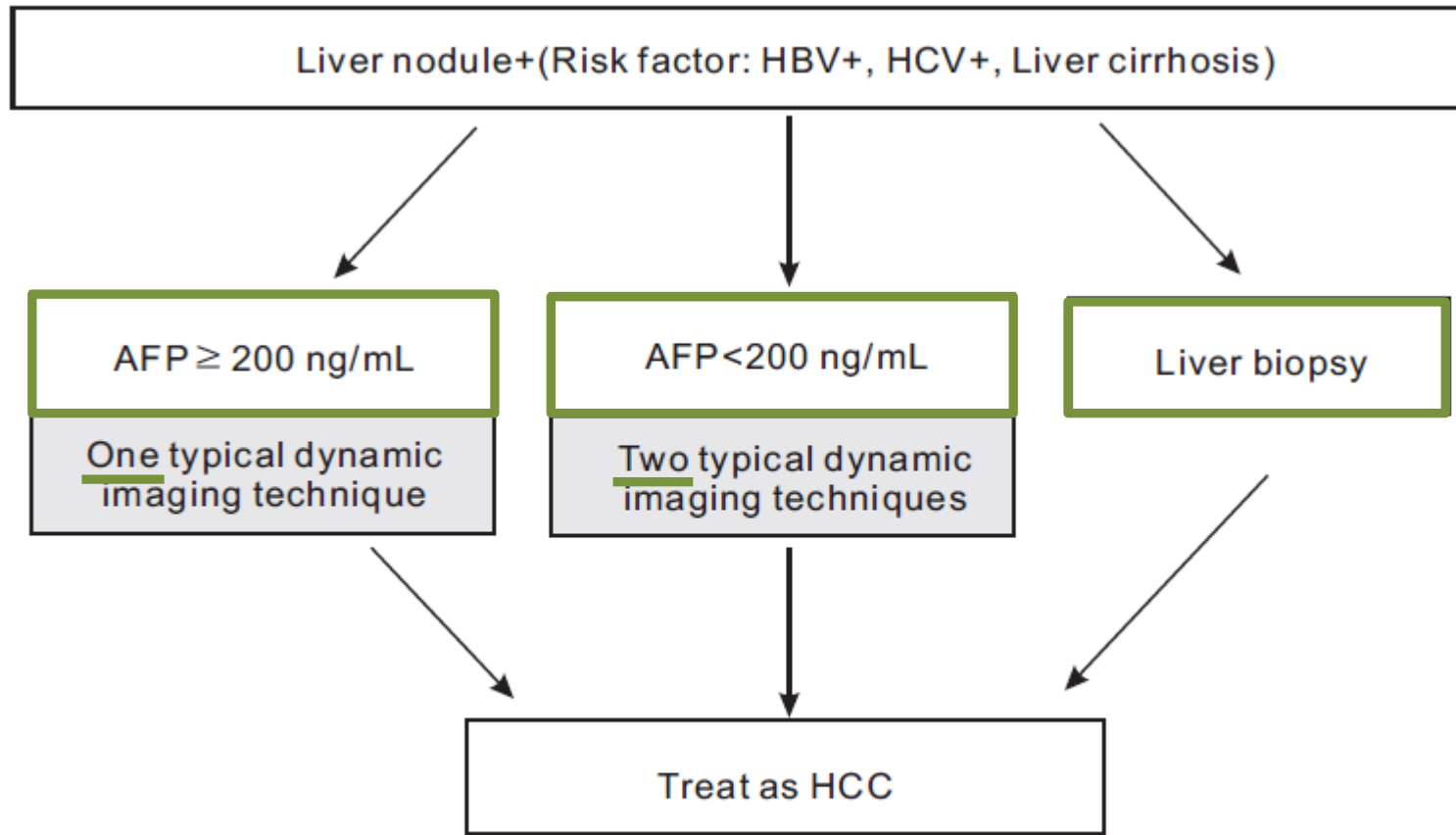
APASL guideline in 2010

(Asian Pacific Association for the Study of the Liver)



KLCSG guideline in 2009

(Korean Liver Cancer Study Group)



If liver cirrhosis patients have a tumor ≥ 2 cm, typical characteristic finding of HCC in either one of dynamic contrast enhancement CT or MRI, regardless of serum AFP level

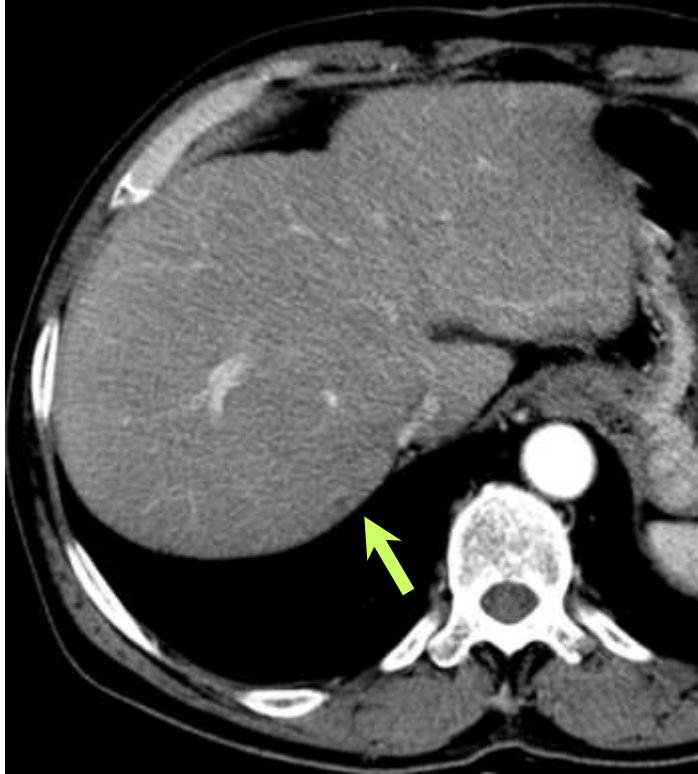
Discordance among the guidelines

- Nodules <1cm showing typical CE pattern
 - AASLD, EASL: US F/U
 - APASL, JSH: regardless of size
- Hypervascularity
 - AASLD, EASL, Korean : CT, MRI
 - APASL, JSH: + CEUS (CTAP)

Discordance among the guidelines

- Hypervascular nodule /s WO
 - AASLD, EASL : Bx
 - APASL, JSH: further evaluation with CEUS, SPIO-MRI or EOB-MRI
 - some ICC, adenoma, FNH-like nodule
- Hypovascular nodule
 - AASLD, EASL, Korean: US F/U
 - APASL: defect on SPIO-MRI or CEUS → HCC
 - HGDN
 - JSH : defect on EOB-MRI & CEUS →HCC, one of them →Bx

Nodule (<1cm) showing hypervascularity /s WO

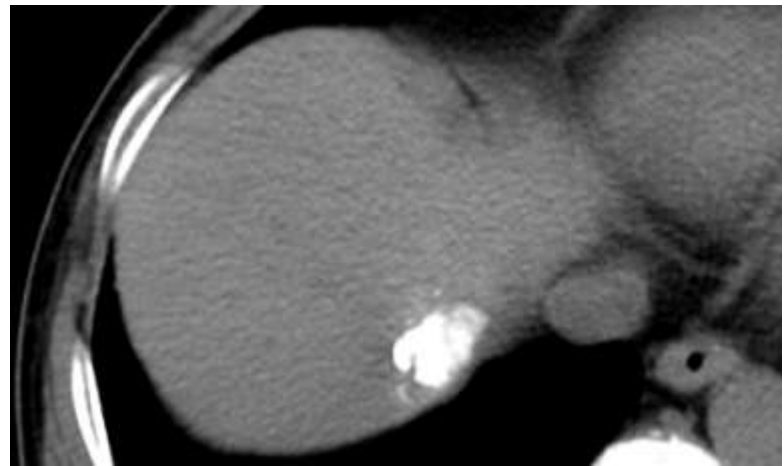
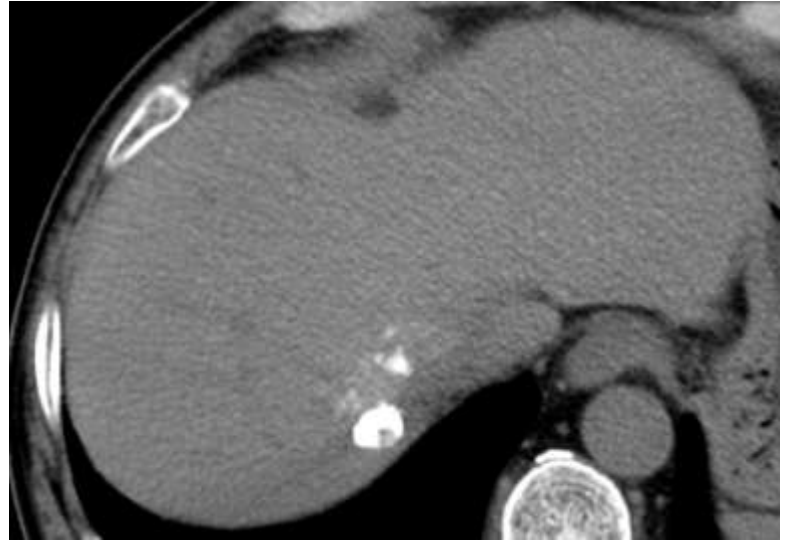
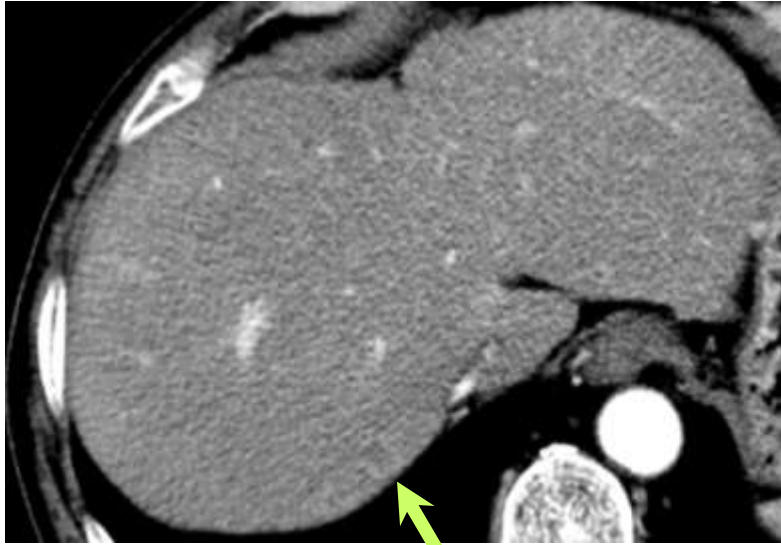


AP

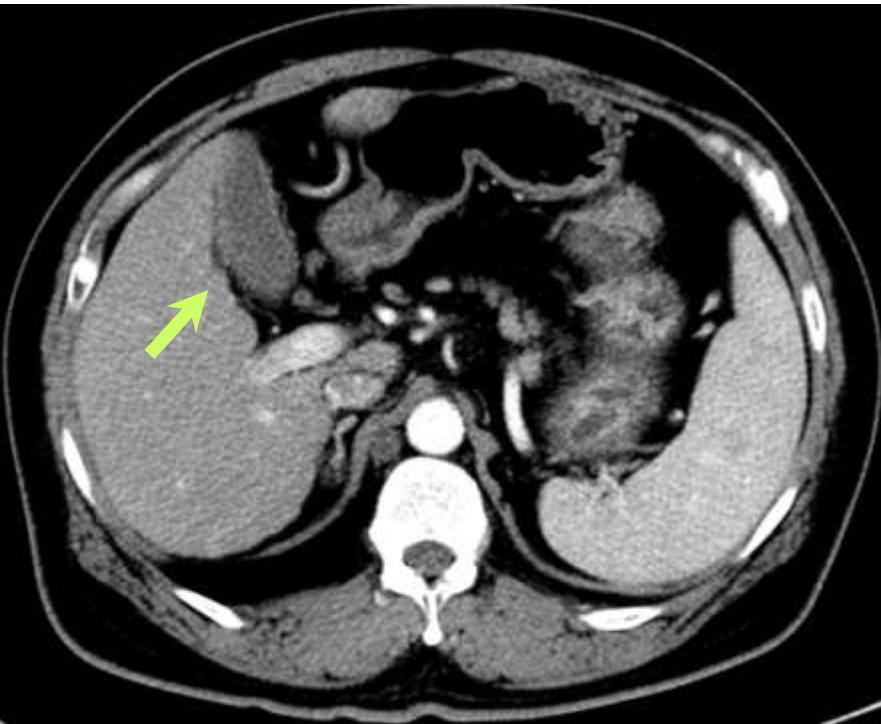


EP

→ 14m later, Dx of HCC & TACE

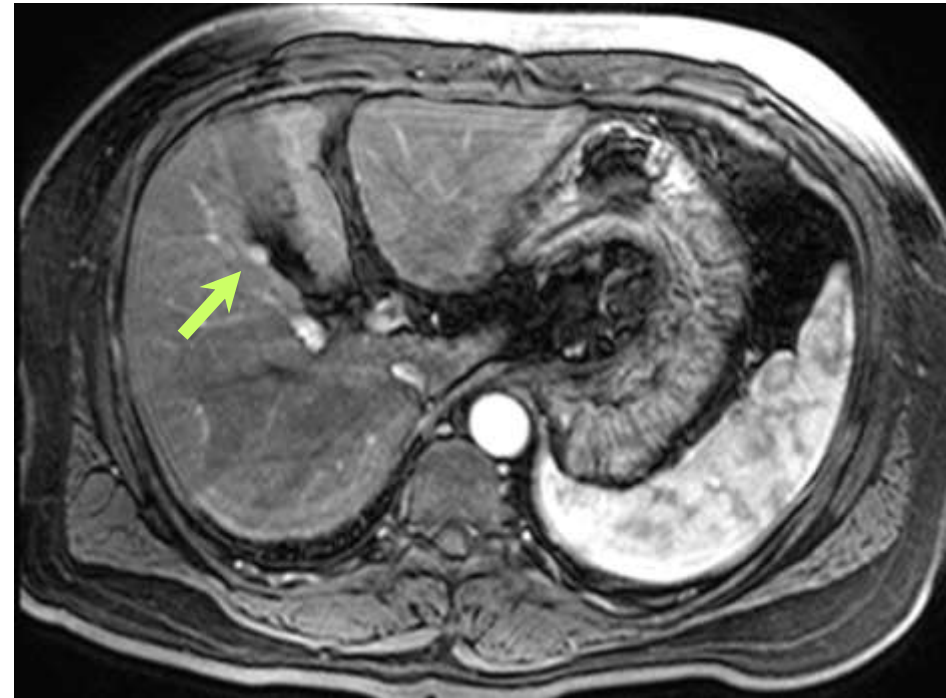


Nodule (<1cm) showing hypervascularity /s WO



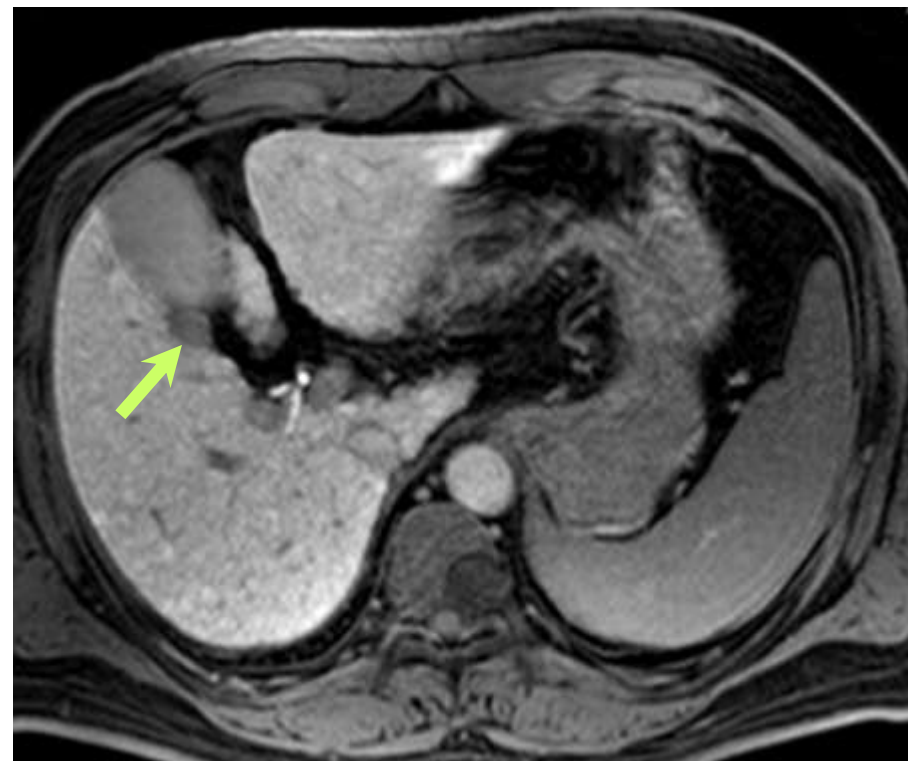
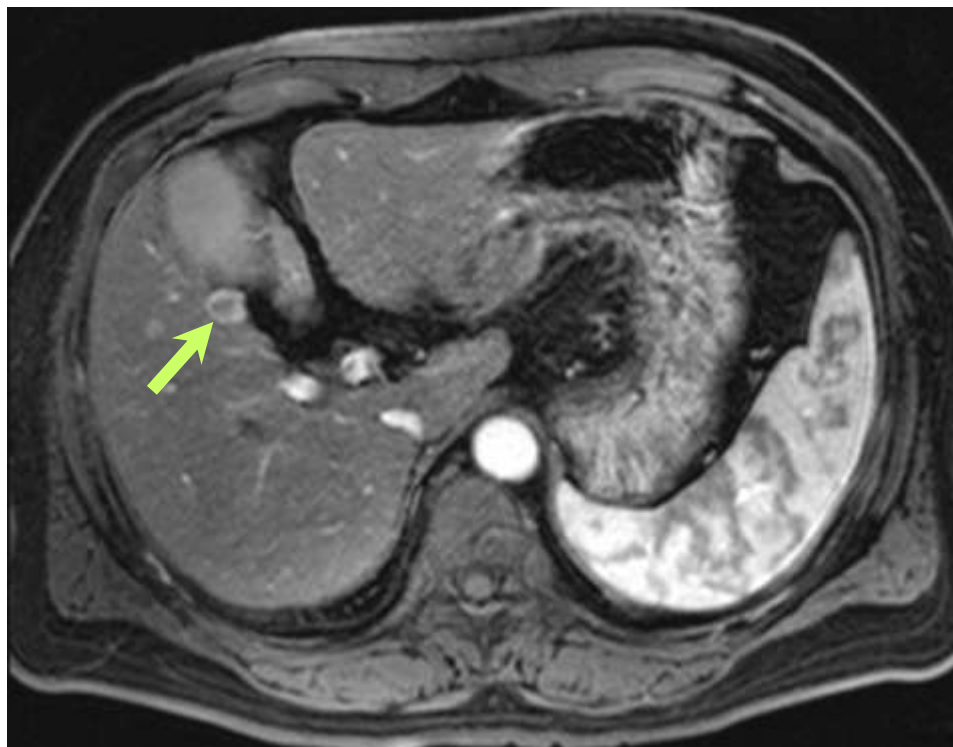
AP

→ 4m later



AP

→12m later,
HCC with microvascular invasion



Hypervascular Small HCC ($\leq 1\text{cm}$) on Gd-EOB-DTPA MRI & DWI

TABLE 3: Results of Multivariate Analysis for Diagnosis of Hepatocellular Carcinoma 1 cm and Smaller

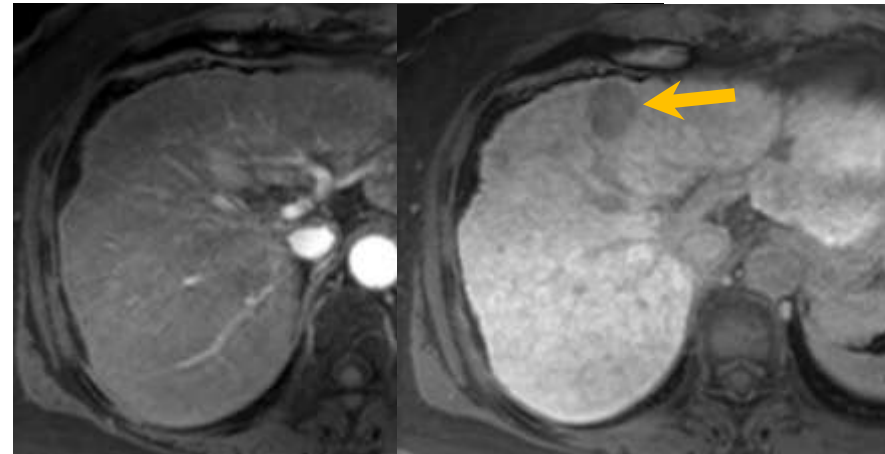
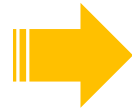
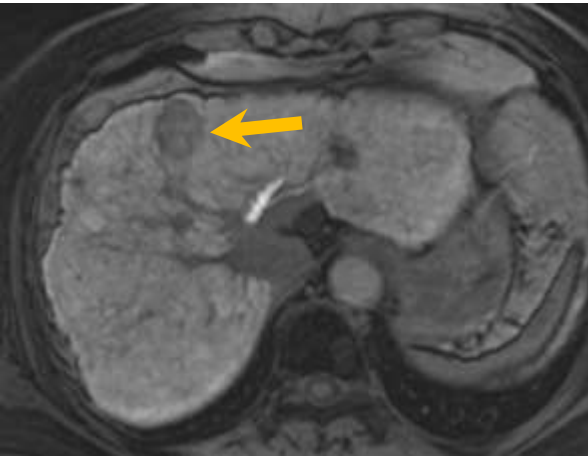
| MRI Finding | Odds Ratio | 95% CI | <i>p</i> |
|--|------------|----------|----------|
| <u>Hyperintensity on T2-weighted images</u> | 16.1 | 4.7–55.1 | <0.0001 |
| <u>Hyperintensity on diffusion-weighted images</u> | 5.7 | 1.6–20.5 | 0.0081 |
| Hypointensity on hepatobiliary phase images | 3.4 | 0.8–14.7 | 0.7537 |
| Washout on portal venous or 3-minute late phase images | 0.8 | 0.2–3.4 | 0.1063 |

TABLE 4: Sensitivity and Specificity for Diagnosis of Hepatocellular Carcinoma Measuring 1 cm or Smaller

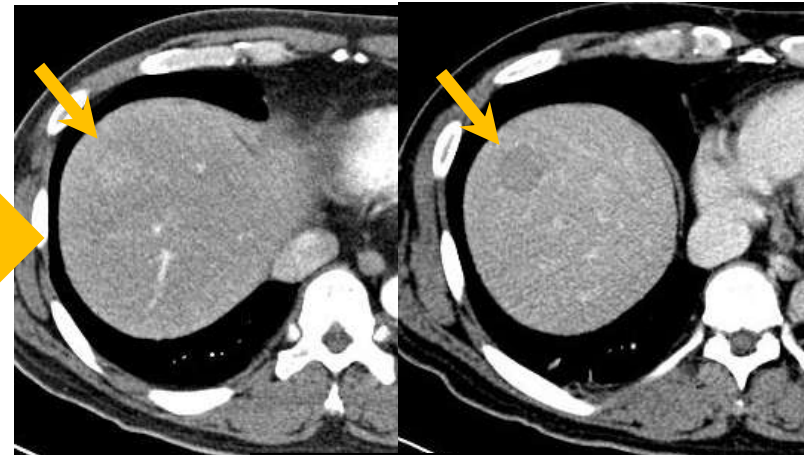
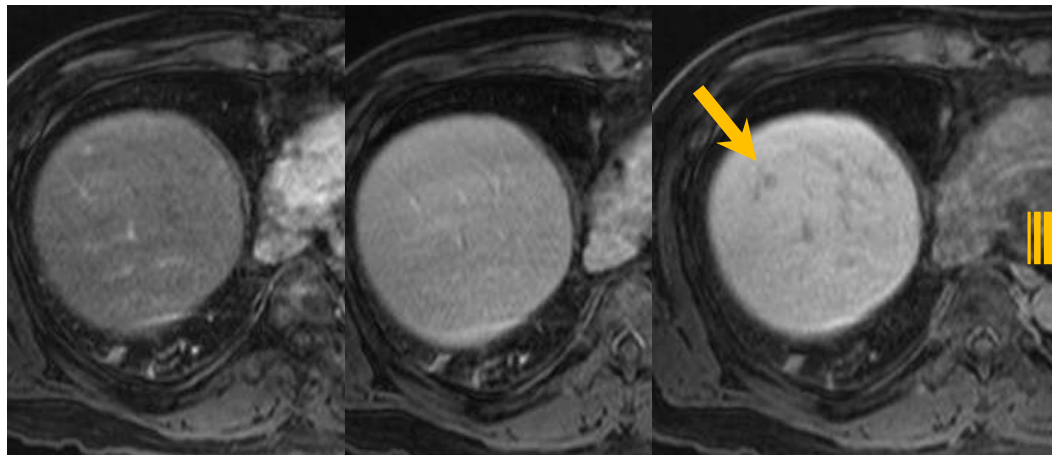
| MRI Finding | Sensitivity (%) | 95% CI | Specificity (%) | 95% CI |
|--|-----------------|-----------|-----------------|-----------|
| <u>Hyperintensity on T2-weighted images</u> | 90.7 | 80.3–95.9 | <u>78.8</u> | 55.1–91.8 |
| <u>Hyperintensity on diffusion-weighted images</u> | 73.2 | 60.3–83.0 | <u>84.9</u> | 61.7–95.1 |
| Hyperintensity on T2-weighted images and diffusion-weighted images | 67.6 | 54.5–78.4 | 87.9 | 65.2–96.6 |

Hypovascular HCC

-On EOB-MRI, hypovascular hypointense nodule on HBP



32 m later



8 m later

Hypovascular HCC

- Among hypovascular hypointense nodule on HBP of EOB-MRI,
→ Recognition of HCC or high risk nodule

- Size \geq 15mm

Kumada T et al. AJR 2011;197:58

- Hyperintensity on DWI

Kim YK et al. Radiology 2012;265(1):104

- Hyperintensity on T2WI, growth rate

Hyodo T et al. Radiology 2013;266(2):480

- Fat within nodule, Hyperintensity on T1WI, growth rate

Higaki A et al. JMRI 2013;37:1377

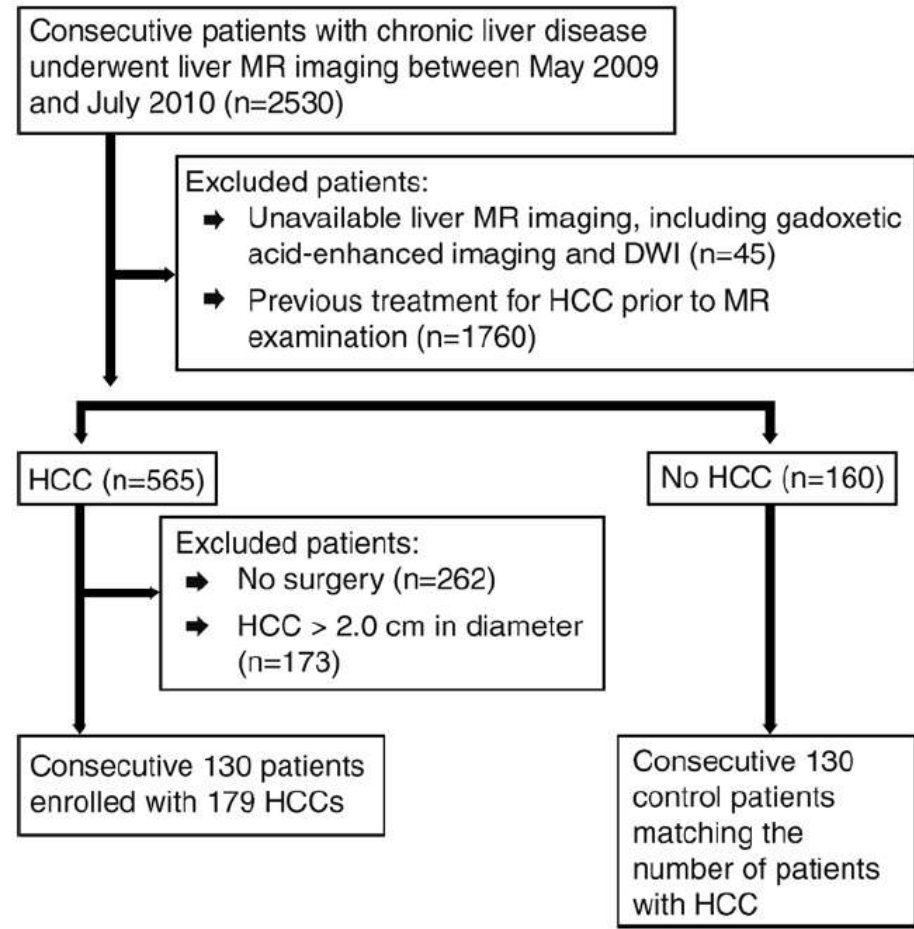
For early diagnosis of HCC

- Higher sensitivity using combination of contrast-enhanced MRI and DWI

Small Hepatocellular

Carcinomas: Improved Sensitivity by Combining Gadoxetic Acid-enhanced and Diffusion-weighted MR Imaging Patterns¹

Park MJ, Kim YK et al.
Radiology 2012;264(3):761



■ <Dx criteria for HCC>

① Gadoxetic acid set :

a. hypervascularity + WO + HBP low SI

② DWI set :

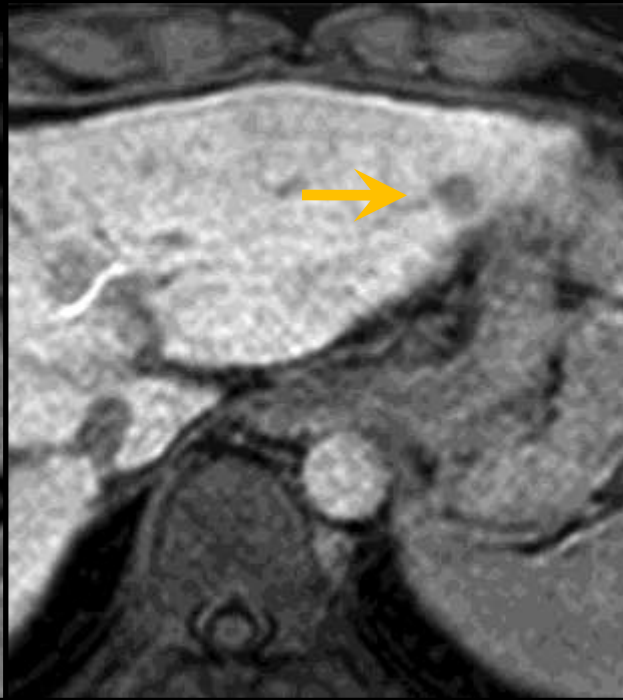
DWI high SI + ADC value \leq that of parenchyma

③ Combined set:

a. hypervascularity + WO + HBP low SI

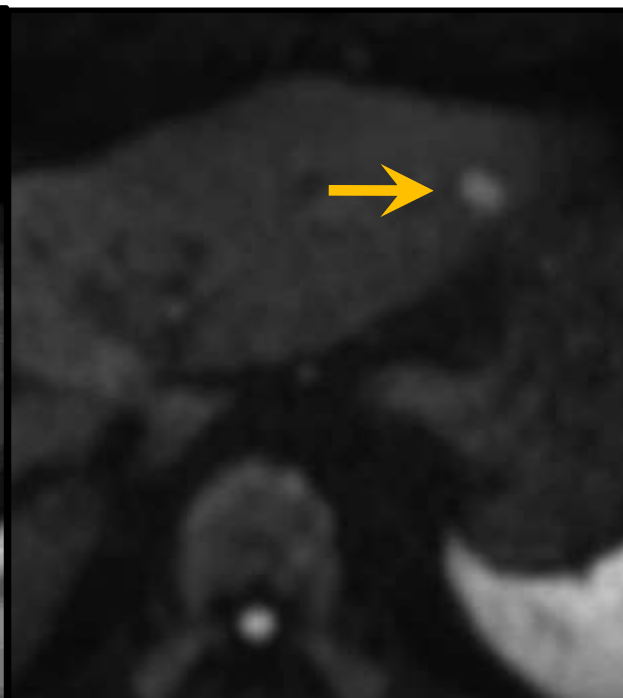
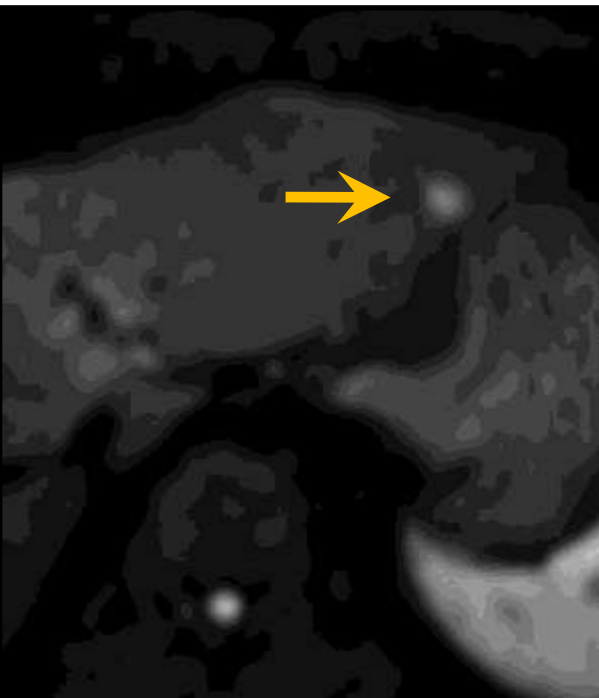
a. hypervascularity + HBP iso- or low SI + DWI high SI

faint or no a. hypervascularity + HBP low SI + DWI high SI

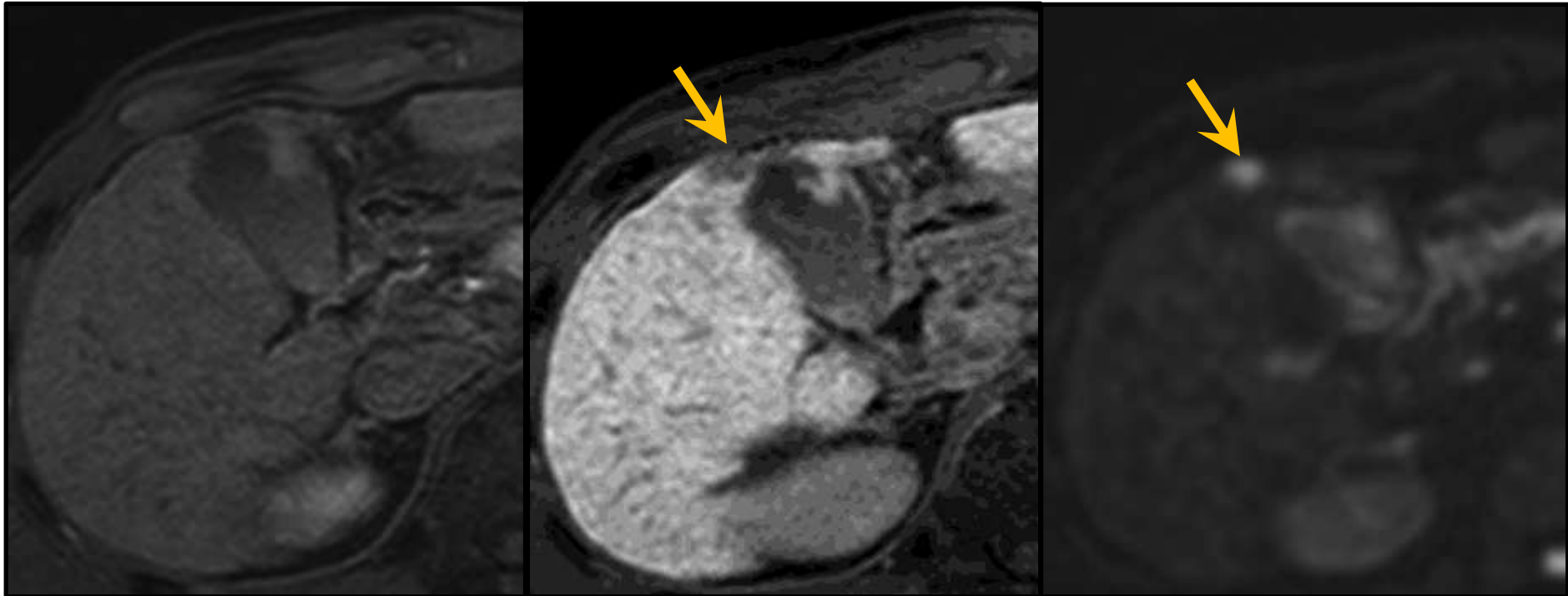


62/M,
HCC grade II

- Gadoxetic acid set:4
- DWI set: 4
- Combined set: 4

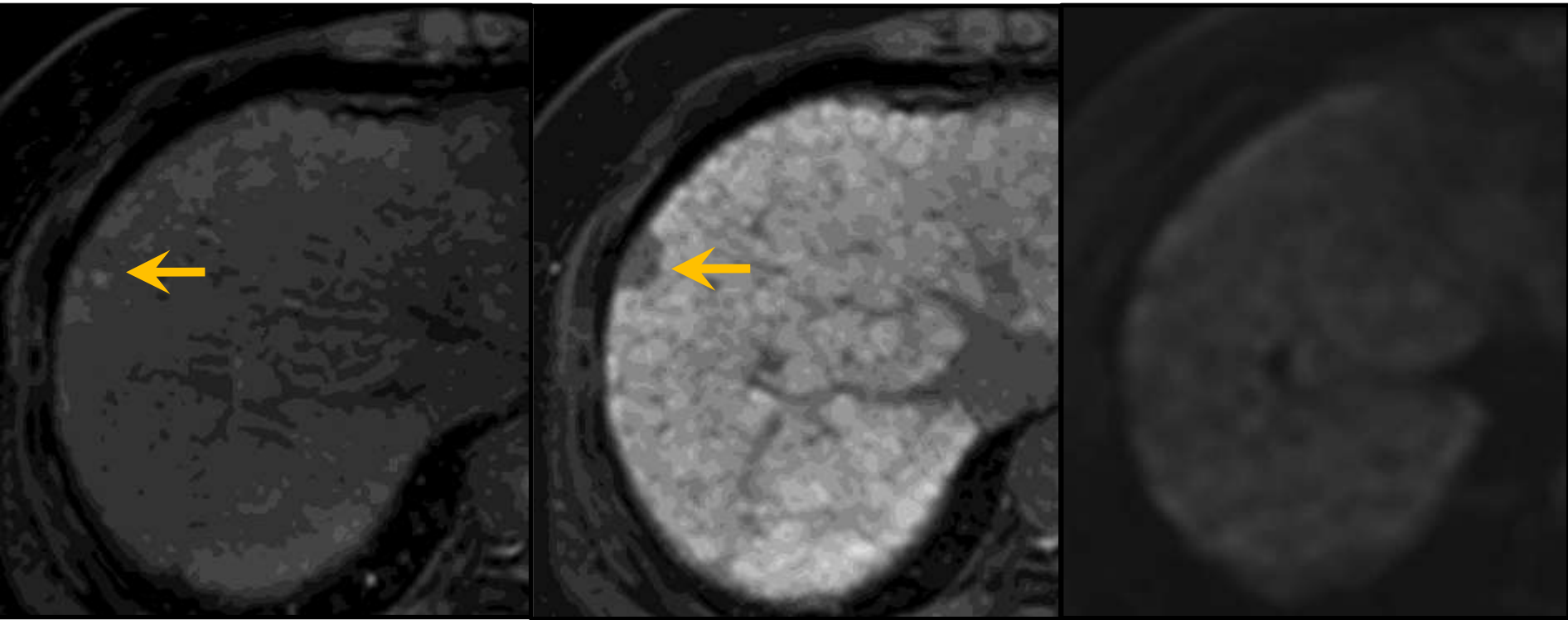


66/M, HCC grade I



- **Gadoxetic acid set: 1**
- **DWI set: 3 or 4**
- **Combined set: 3**

55/M, HCC grade I > II



- **Gadoxetic acid set: 3**
- **DWI set: 1**
- **Combined set: 3**

Sensitivity and Positive Predictive Values for the Detection of 179 HCCs

| Lesion Group and Imaging Modality | Observer 1 | | Observer 2 | | Observer 3 | | Pooled Data | |
|-------------------------------------|--------------------------|------------------|--------------------------|------------------|--------------------------|------------------|-------------------------|------------------|
| | Sensitivity* | PPV [†] | Sensitivity* | PPV [†] | Sensitivity* | PPV [†] | Sensitivity* | PPV [†] |
| All lesions (n = 179) | | | | | | | | |
| Gadoxetic acid set | 81.0 (145) | 98.6 (2) | 82.1 (147) | 98.7 (2) | 80.5 (144) | 98.0 [3] | 81.4 (437) [‡] | 98.4 [7] |
| DW imaging set | 79.9 (143) | 96.6 (5) | 77.7 (139) | 97.2 (4) | 78.8 (141) | 96.6 [5] | 78.8 (423) [‡] | 96.8 [14] |
| Combined set | 92.7 (166) [§] | 98.2 (3) | 91.1 (163) [§] | 98.2 (3) | 93.3 (167) [§] | 97.1 [5] | 92.4 (496) [‡] | 97.8 [11] |
| Lesions ≤1.0 cm (n = 55) | | | | | | | | |
| Gadoxetic acid set | 58.2 (32) | 94.1 (2) | 61.8 (34) | 94.4 (2) | 56.4 (31) | 93.9 [2] | 58.8 (97) | 94.2 [6] |
| DW imaging set | 63.6 (35) | 87.5 (5) | 56.4 (31) | 88.6 [4] | 60.0 (33) | 89.2 [4] | 60.0 (99) | 88.4 [13] |
| Combined set | 85.5 (47) [§] | 94.0 (3) | 81.8 (45) [§] | 93.8 [3] | 87.3 (48) [§] | 92.3 [4] | 84.8 (140) [§] | 93.3 [10] |
| Lesions >1.0 cm (n = 124) | | | | | | | | |
| Gadoxetic acid set | 91.13 (113) | 100 (0) | 91.13 (113) | 100 [0] | 91.13 (113) | 99.1 [1] | 91.1 (339) [‡] | 99.7 [1] |
| DW imaging set | 87.1 (108) | 100 (0) | 87.1 (108) | 100 [0] | 87.1 (108) | 99.1 [1] | 87.1 (324) [‡] | 99.7 [1] |
| Combined set | 96.0 (119) | 100 (0) | 95.2 (118) | 100 [0] | 96.0 (119) | 99.2 [1] | 95.7 (356) [‡] | 99.7 [1] |

- The combination of gadoxetic acid–enhanced MRI and DWI yielded better sensitivity in the detection of small HCCs than each MR imaging technique alone.

Conclusions

- Increased detection of small HCC due to advanced imaging
- More studies necessary for indeterminate liver nodule
- Guidelines for HCC diagnosis; HBP (and DWI) MRI