

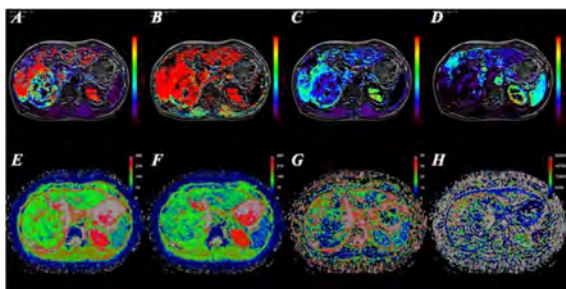
# Characterize Hepatocellular Carcinoma with IVIM-DWI and DCE-MRI in Combination: Preliminary Experience

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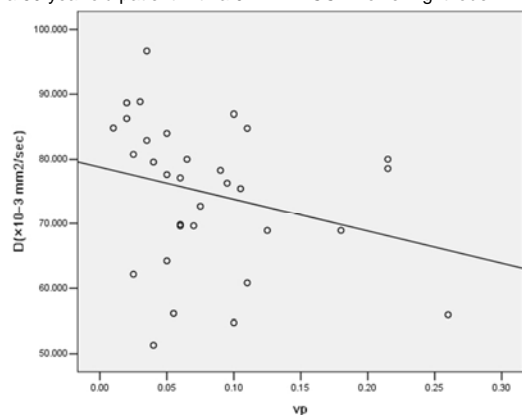
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**Purpose:** To quantify diffusion and perfusion characteristics of hepatocellular carcinoma (HCC) with both intravoxel incoherent motion (IVIM) diffusion-weighted imaging (DWI) and dynamic contrast enhanced (DCE) MRI, and to evaluate the correlation between the resulting parameters.

**Materials and Methods:** This prospective study was approved by the institutional review board and written informed consent was obtained. 35 patients (male-female ratio, 31:4; mean age, 47.6 years old; range, 23~66) with cirrhosis and HCCs underwent IVIM-DWI (respiratory triggered SS-EPI using 12 b values from 0 to 800 sec/mm<sup>2</sup>) and DCE-MRI scans at 1.5 T MR scanner (Achieva, Philips Healthcare, Best, Netherlands). DCE-MRI protocol consisted of one pre-contrast scan for T<sub>1</sub> mapping and a dynamic axial 3D-FFE sequence centering on the lesions (TR/TE 11/3, FA 5 and 15°, matrix 204×177, slice thickness 4 mm, 22 slices, temporal resolution 6~8 sec, 36 dynamics for average 10 min) before and after IV bolus of 0.1 mmol/kg of Gd-DTPA (Magnevist). Time-activity curves were converted to gadolinium concentration, then analyzed using a single-input two compartment model and the following DCE-MRI quantitative parameters of HCCs were obtained: volume transfer coefficient ( $K^{trans}$ ), reverse reflux rate constant ( $k_{ep}$ ), the fractional volume of extravascular extracellular space ( $v_e$ ) and the fractional volume of plasma ( $v_p$ ). ROIs were placed on the IVIM-DWI parametric maps in order to extract the apparent diffusion coefficient (ADC), true diffusion coefficient (D), pseudo-diffusion coefficient (D\*) and perfusion fraction (f) of tumors and the liver parenchyma using a bi-exponential model<sup>[1]</sup>. Pearson or Spearman correlation was computed to assess correlation between IVIM-DWI and DCE-MRI parameters of HCCs. IVIM-DWI parameters were compared between liver parenchyma and tumors using Paired-Samples T Test or Wilcoxon Test.



**Figure 1.** (A-D)  $K^{trans}$ ,  $k_{ep}$ ,  $v_e$ ,  $v_p$  maps and (E-H) ADC, D, f, D\* maps in a 59 year old patient with a 94 mm HCC in lower right lobe.



**Figure 2.** Graph demonstrating a significantly weak negative correlation between  $v_p$  and D ( $r=-0.391$ ,  $P<0.05$ ).

**Results:** 35 HCCs were evaluated (mean size 56 mm, range 33~168 mm).  $v_p$  was inversely correlated with D in HCCs ( $r=-0.391$ ,  $P=0.027$ ) (Fig.2); While no significant correlation between the other IVIM-DWI and DCE-MRI parameters was detected (correlation coefficients range  $-0.278\sim0.17$ ,  $P>0.05$ ) (Fig.1). Furthermore, when compared with liver parenchyma, HCCs showed significantly lower ADC, D, f and D\* values ( $P<0.001$ ) (Table, Fig.1).

Parameter	Liver	HCC	t/Z value	P value
ADC ( $\times 10^{-3}$ mm <sup>2</sup> /sec)	1.14 $\pm$ 0.07	0.87 $\pm$ 0.12	-6.113	<.001
D ( $\times 10^{-3}$ mm <sup>2</sup> /sec)	0.99 $\pm$ 0.06	0.76 $\pm$ 0.13	-5.822	<.001
f (%)	19.66 $\pm$ 3.46	11.12 $\pm$ 3.16	9.692	<.001
D* ( $\times 10^{-3}$ mm <sup>2</sup> /sec)	118.25 $\pm$ 20.24	73.62 $\pm$ 21.35	8.219	<.001

**Table** Comparison of IVIM and DCE-MRI metrics of liver and HCCs.

**Discussion:** As previously reported, the combination of IVIM-DWI and DCE-MRI can provide an accurate diagnosis of cirrhosis, but without significant correlation between those parameters, since IVIM-DWI and DCE-MRI may reflect different aspects of tissue perfusion<sup>[2]</sup>. However, our study demonstrated that  $v_p$  shows weak negative correlation with D. Because  $v_p$  represents the fractional volume of vascular in tissue, higher  $v_p$  suggests poor differentiation of tumor, which results in diffusion restriction from decreased extracellular space. ADC value and IVIM-DWI parameters were all decreased in HCCs. Increased cellular density in tumor is believed to impair molecules diffusion, thus leading to reduction in ADC and D values. A lower f value in HCCs compared to liver may be explained by blood flow derivation into small, leaky and poorly efficient tumor capillaries, resulting in a decrease of the fast moving blood pool<sup>[3]</sup>. The distortion, small diameter and thus slow blood flow of generated vessels in HCCs can explain the lower D\* value.

**Conclusion:** This study indicates that perfusion parameter  $v_p$  derived from DCE-MRI shows a weak negative correlation with diffusion coefficient D from IVIM-DWI in HCC. IVIM-DWI provides parameters that are significantly different in HCC compared to liver parenchyma, and could potentially be used for characterization of HCC.

**References:** [1] Le Bihan, D., et al., Radiology, 1988. [2] Patel J, et al. J MRI, 2010. [3] Lewin M, et al. Eur Radiol, 2011.