

# The relationship between the quantitative pharmacokinetic parameters of DCE-MRI and the types of uterine fibroids based on T2WIs

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## Introduction

Uterine fibroids are the most common benign tumors in women of reproductive age. Menorrhagia, dysmenorrhea, pelvic pain, and bulk-related symptoms are common symptoms [1]. Hysterectomy or myomectomy is standard treating approach of fibroids. Magnetic resonance-guided high-intensity focused ultrasound (MRg HIFU) ablation, along with uterine fibroid embolization(UAE), is currently major nonsurgical options in the treatment of uterine fibroids [2]. The safe and effective use of MRg HIFU is affected by fibroid type in T2-weighted images (T2WIs) which were the current clinical criterion to select patients for MRg HIFU ablation[3]. The  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  based on Tofts model [4] of dynamic contrast-enhanced MR images (DCE-MRI) represent the process of Gd-DTPA transfer across the capillary wall, which is a surrogate of capillary permeability [5]. The aim of this study is to investigate the relationship between the quantitative pharmacokinetic parameters of DCE-MRI and the fibroid type based on T2WIs ,with expectation to select suitable fibroids for MRg HIFU ablation.

## Methods

This study was approved by the local institutional review board. Twenty-three fibroids in 23 females (37~53 years, mean age:43.96±4.38 years) underwent T1 weighted images(T1WIs), T2WIs and DCE-MRI. The fibroids were divided into three types based on signal intensity in T2WIs[6]: type 1, low signal compared to skeletal muscle, 11 cases; type 2, lower signal than the myometrium and higher than skeletal muscle,7cases; type 3, signal equal to or higher than that of myometrium, 5 cases. In this study, the turbo field echo (TSE) T1WIs, two-dimensional TSE T2WIs and three-dimensional T1-weighted DCE-MRI were performed in a 1.5T scanner (Achieva, Philips Medical System, the Netherland ) with 16-channel XL torso coil. DCE-MRI was performed including a precontrast scan (with a flip angle of 5) and a dynamic scan (with a flip angle of 15) after a single dose injection of Gadopentetate dimeglumine (Magnevist) at a dose of 0.1 mmol/kg through a peripheral vein at a rate of 2 mL/s. Dynamic scan was performed by 60 dynamics, TR/TE=4.84/2.35ms, field of view = 200mm×200mm×96mm, matrix = 176×176, voxel size= 1.7×1.7×4mm<sup>3</sup>,NSA=2, scan duration is 330second.The analysis of DCE-MRI data was performed using IDL version 6.4 Software. The process mainly included: Load the data, define the AIF, map calculation, review results, export. The quantitative parameters of  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  were obtained in workstation. The differences of  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  among the three types were determined using One-Way ANOVA. All statistical analyses were considered to be significant with  $P<0.05$ .

## Results

The quantitative pharmacokinetic parameters ( $K^{trans}$ ,  $K_{ep}$  and  $V_e$ ) of each type were shown in Table 1 as mean ± standard deviation.  $P$ -value of ANOVA analysis among the three fibroid types were shown in Table 2. The  $K^{trans}$  of the three fibroid types is very close and there are no statistic differences among them. The  $K_{ep}$  of type 3 is higher than type 1 and type 2.The significant differences of  $K_{ep}$  were shown between type 1 and type 3, type 2 and type 3. Moreover, there were no significant differences of  $V_e$  among the three fibroid types , although average values of  $V_e$  seem to be large difference among them from Table 1. The three fibroid types were shown typically with T2WIs and color maps of  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  in Fig 1.

**Table 1**  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  of three fibroid types

Group	$K^{trans}$ (min <sup>-1</sup> )	$K_{ep}$ (min <sup>-1</sup> )	$V_e$
Type1(n=11)	0.38±0.25	0.16±0.05	1.94±2.53
Type2(n=7)	0.47±0.30	0.14±0.05	2.27±2.55
Type3(n=5)	0.35±0.23	0.28±0.13	0.62±0.40

**Table 2** Differences of  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  among 3 fibroid types

	$K^{trans}$	$K_{ep}$	$V_e$
Type1&2	0.673	0.294	0.762
Type1&3	0.892	0.023*	0.293
Type2&3	0.642	0.005*	0.233

\*:  $P<0.05$

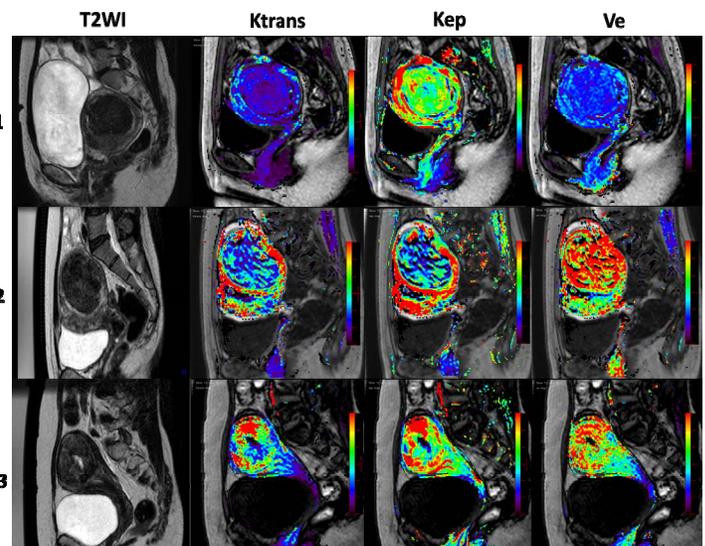
## Discussion & Conclusions

To our knowledge, this is the first study to compare the DCE-MRI derived contrast agent kinetic characters among the three fibroid types by using  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  based on Tofts model. There were no statistic differences of  $K^{trans}$  among the three fibroid types, suggesting the similar volume transfer constant of Gd-DTPA between the capillary and extravascular extracellular space (EES). In current study, the  $V_e$  representing the volume of EES per unit volume of tissue were no significant differences among the 3 fibroids types , but the  $V_e$  of type 1 and type 2 were greater than 1. It indicated that the length of the dynamic scanning for this study may not be long enough to accurately estimate  $V_e$  in voxels. On the other hand, it demonstrated that type 3 fibroids have the haemodynamic characters of a fast wash-out. In addition,  $K_{ep}$  of type 3 was higher than type 1 and type 2. All of these demonstrated that the wash-out of type 3 is faster than other two types. It implied that the more heat energy may be taken away if carry out the HIFU ablation. Therefore, the patients with fibroids of type 3 may be not very suitable for MRg HIFU.

In sum, the  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  based on Tofts model, as an objective and sensitive method for DCE-MRI data analyzing, can reveal differences of capillary permeability and blood flow among the three fibroid types, and be proposed to guide patients selection for MRg HIFU ablation.

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**Fig.1** T2WI and color maps of  $K^{trans}$ ,  $K_{ep}$  and  $V_e$  in the 3 fibroid types