

# Assessment of the Agreement between Cerebral Hemodynamic Indices Quantified Using Dynamic Susceptibility Contrast and Dynamic Contrast-enhanced Perfusion Magnetic Resonance Imagings

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Received: 19-10-2017  
Accepted: 11-12-2017  
Published: 22-01-2018

ABSTRACT

**Background:** Brain tumor is one of the most common tumors. A successful treatment might be achieved with an early identification. Pathological investigation as the gold standard method for tumor identification has some limitations. Noninvasive assessment of tumor specifications may be possible using perfusion-weighted magnetic resonance imaging (MRI). Cerebral blood volume (CBV) and cerebral blood flow (CBF) could be calculated based on dynamic contrast-enhanced MRI (DCE-MRI) in addition to dynamic susceptibility contrast MRI (DSC-MRI) modality. Each category of the cerebral hemodynamic and permeability indices revealed the specific tumor characteristics and their collection could help for better identification of the tumor. Some mathematical methods were developed to determine both cerebral hemodynamic and permeability indices based on a single-dose DCE perfusion MRI. There are only a few studies available on the comparison of DSC- and DCE-derived cerebral hemodynamic indices such as CBF and CBV. **Aim:** The objective of the study was to validate first-pass perfusion parameters derived from T1-based DCE method in comparison to the routine T2\*-based DSC protocol. **Materials and Methods:** Twenty-nine patients with brain tumor underwent DCE- and DSC-MRIs to evaluate the agreement between DSC- and DCE-derived cerebral hemodynamic parameters. Agreement between DSC- and DCE-derived cerebral hemodynamic indices was determined using the statistical method described by Bland and Altman. The reliability between DSC- and DCE-derived cerebral hemodynamic indices was measured using the intraclass correlation analysis. **Results:** The achieved magnitudes for DCE-derived CBV (gray matter [GM]:  $5.01 \pm 1.40$  mL/100 g vs. white matter [WM]:  $1.84 \pm 0.74$  mL/100 g) and DCE-derived CBF (GM:  $60.53 \pm 12.70$  mL/100 g/min vs. WM:  $32.00 \pm 6.00$  mL/100 g/min) were in good agreement with other studies. The intraclass correlation coefficients showed that the cerebral hemodynamic indices could accurately be estimated based on the DCE-MRI using a single-compartment model ( $>0.87$ ), and DCE-derived cerebral hemodynamic indices are significantly similar to the magnitudes

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**How to cite this article:** Zakariaee SS, Oghabian MA, Firouznia K, Sharifi G, Arbabi F, Samiei F. Assessment of the Agreement between Cerebral Hemodynamic Indices Quantified Using Dynamic Susceptibility Contrast and Dynamic Contrast-enhanced Perfusion Magnetic Resonance Imagings. J Clin Imaging Sci 2018;8:2. Available FREE in open access from: <http://www.clinicalimagingscience.org/text.asp?2018/8/1/2/223734>.

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**DOI:** 10.4103/jcis.JCIS\_74\_17

achieved based on the DSC-MRI ( $P < 0.001$ ). Furthermore, an acceptable agreement was observed between DSC- and DCE-derived cerebral hemodynamic indices. **Conclusion:** Based on the measurement of the cerebral hemodynamic and blood–brain barrier permeability using DCE-MRI, a more comprehensive collection of the physiological parameters could be achieved for tumor evaluations.

**KEYWORDS:** *Bland and Altman analysis, brain tumor, cerebral hemodynamic indices, dynamic contrast-enhanced-magnetic resonance imaging, dynamic susceptibility contrast-magnetic resonance imaging, intraclass correlation coefficient*

## INTRODUCTION

Brain tumor is one of the most common tumors in the world. The primary incidence of the central nervous system (CNS) tumors is approximately 7 people per 100,000 in adults.<sup>[1]</sup> Studies show that the outbreaks of the disease are 6.04 and 3–5 per 100,000 population/year in the USA and Europe, respectively.<sup>[2,3]</sup> Most brain tumors are asymptomatic in the early stages of their progression. Symptoms and deadly characteristics appear in the final stages. Therefore, a successful treatment might be achieved with an early identification.

Histopathological investigation is the gold standard method for tumor identification. The main limitations of the histopathology-based tumor evaluation include inherent sampling errors associated with stereotactic biopsy, limited number of sampling, and the inability to evaluate residual tumor tissues after cytoreductive surgery. In cytoreductive surgery, the histological examination could only be performed on the resected specimens and remaining tumor tissues could not be evaluated.<sup>[4]</sup> Magnetic resonance imaging (MRI) method is a promising noninvasive method for tumor assessment. In the evaluation of brain tumors using MRI methods, metabolic and physiological changes of the peripheral tissues could be examined in addition to the abnormal sites. Conventional MRI method does not provide reliable information about the tumor physiology including microvasculature, angiogenesis, metabolism, micronecrosis, and cellularity (parameters that are important to determine tumor specifications).<sup>[5]</sup> Advanced MRI methods provide a comprehensive dataset from *in vivo* physiological and metabolic information that could complement the histopathology examinations.<sup>[6]</sup>

Noninvasive assessment of tumor hemodynamic characteristics (before surgery) is possible using perfusion MRI methods.<sup>[7]</sup> Angiogenesis as the main feature of tumor malignancy is a critical procedure in the solid tumor growth and cancer cell metastasis.<sup>[8]</sup> First-pass  $T_2^*$  dynamic susceptibility contrast (DSC) MRI method is considered as the most powerful tool to evaluate tumor

vasculature, which is known to be a clinical index for tumor grading and treatment response evaluation.<sup>[9]</sup>

Dynamic contrast-enhanced MRI (DCE-MRI) is the standard imaging modality to evaluate cerebral hemodynamic and permeability parameters, which provide additional pathological information for preoperative identification of the tumor. Each category of DSC and DCE indices revealed some specific tumor characteristics and their collection could help for better tumor identification and grading.<sup>[9-11]</sup>

Both imaging sequences could be implemented using the dual dynamic acquisition mode with a single-dose contrast administration, which is accessible only in the new MRI systems. However, this technique is not implemented in all available commercial scanners.

Although  $T_2^*$ -based MRI methods such as DSC is faster (about 2 min) and provides higher temporal resolution in comparison to T1-based methods (i.e., DCE), DCE-MRI has its own advantages such as lower susceptibility artifacts and an acceptable imaging time.<sup>[11-13]</sup> Recent studies showed that cerebral blood volume (CBV) and cerebral blood flow (CBF) parameters could be calculated using DCE-MRI method.<sup>[14,15]</sup> There is limited evidence to validate DCE-MRI method for deriving cerebral hemodynamic indices such as CBF and CBV for clinical tumor evaluation. In this study, an existing paradigm is re-evaluated to determine cerebral hemodynamic parameters using DCE-MRI, and the results were compared with perfusion parameters derived from DSC-MRI protocol. To validate these new DCE-derived parameters (CBF and CBV), our patient data were tested by this technique. We believe if a T1-based perfusion method can fulfill first-pass perfusion parameters as efficient as its inherent capability for permeability measurements, then more physiological characteristics of the tumors could be obtained using a single-dose DCE examination.

## Theory

To calculate the physiological parameters based on DCE-MRI method, signal intensities must be converted

to relaxivity changes. Relaxivity changes are directly proportional to the contrast concentration. Cerebral hemodynamic and permeability parameters quantified based on time-concentration curves.

#### *Quantification of cerebral blood volume and cerebral blood flow using dynamic contrast-enhanced-magnetic resonance imaging*

As mentioned above, cerebral hemodynamic parameters including CBV and CBF could also be derived based on the DCE-MRI using a single-compartment model. CBV (in mL/100 g) is the volume of blood in a given amount of brain tissue, which is calculated by equation 1.

$$CBV = \frac{H \int C(t) dt}{\rho \int C_a(t) dt} \quad (1)$$

Where  $C(t)$  and  $C_a(t)$  represent the concentration of the contrast in the tissue and artery at time  $t$ , respectively. Brain tissue density ( $\rho$ ) is set to 1.04 g/mL, and  $H = (1-Har)/(1-Hcap)$  is considered to differentiate capillary hematocrits ( $Hcap = 25\%$ ) from large vessel hematocrits ( $Har = 45\%$ ). Because of normalization to the arterial input function (AIF), the quantified CBV is independent of the injected contrast magnitude.

Determined CBV is the blood volume of the intravascular and leakage space. Corrected CBV reflects only blood volume in the intravascular space. Therefore, corrected CBV would be achieved by the removal of volume contribution of fractional leakage space (equation 2).

$$CBV_{corrected} = CBV_{uncorrected} - Ve \cdot CBV_{uncorrected} \quad (2)$$

Cerebral blood flow (CBF) is the volume of blood passing through a defined amount of brain tissue per unit of time (mL/100 g/min). Tissue concentration ( $C(t)$ ) determined using linear control theory as stated by equation 3:

$$C(t) = \rho \cdot H \cdot CBF \cdot (C_a(t) \otimes R(t)) \quad (3)$$

Where  $\otimes$  is the convolution operator, and  $R(t)$  is the residual function. In this study, Fourier transform deconvolution is used to quantifying CBF values.<sup>[15,16]</sup> Using the Fourier transform deconvolution, equation 4 thus becomes:

$$CBF \cdot R = \frac{1}{\rho \cdot H} F^{-1} \left\{ \frac{F\{C(t)\}}{F\{C_a(t)\}} \right\} \quad (4)$$

Where  $F^{-1}\{\}$  denotes the inverse Fourier transformation. CBF could be determined by taking the inverse Fourier transformation of  $\frac{F\{C(t)\}}{F\{C_a(t)\}}$  at every time point with known arterial input and time-concentration curve.

## MATERIALS AND METHODS

### Imaging protocols

All images were acquired on a 1.5-Tesla clinical MRI scanner (TrioTim, Siemens Medical Solutions, Erlangen, Germany). The precontrast images were obtained using transverse T1-weighted spin echo (repetition time [TR]/echo time [TE], 370/8.7 msec; flip angle, 90; slice thickness, 5 mm; number of averages, 1; matrix, 512 × 464), transverse T2-weighted spin echo (TR/TE, 3300/99; echo train length, 11; flip angle, 120; slice thickness, 5 mm; number of averages, 2; matrix, 384 × 288), and transverse FLAIR (TR/TE, 5580/92 msec; TI, 1935; flip angle, 150; echo train length, 28; slice thickness, 4 mm; number of averages, 1; matrix, 512 × 512) sequences. For the baseline T1 calculation, four pre-contrast data sets were acquired using a three-dimensional T1-weighted gradient echo sequence with different flip angles ( $\alpha=2^\circ, 10^\circ, 20^\circ$ , and  $25^\circ$ ). The sequence parameters were as follows: TR=12 ms, TE=3.5 ms, matrix size=256×224, number of excitations (NEX)=1, and slice thickness=5 mm.

Dynamic contrast-enhanced T1-weighted imaging was performed using a gradient recalled echo sequence (GR) with TR = 4.13 ms, TE = 1.54 ms, field of view = 200 × 200 mm<sup>2</sup>, matrix size = 256 × 224, flip angle = 15°, NEX = 1, slice thickness = 5 mm, number of measurements = 70, and gap = 5 mm. At the sixth acquisition, Gd-DTPA (0.1 mL/kg) was injected intravenously with a rate of 2 mL/second, followed immediately by a bolus injection of 15 mL saline flush at the same rate.

DSC-MRI was performed using a gradient echo-planar pulse sequence with TR = 1860 ms, TE = 45 msec, field of view = 200 × 200 mm<sup>2</sup>, matrix size = 128 × 116, flip angle = 90°, NEX = 1, slice thickness = 5 mm, number of measurements = 64, and gap = 5 mm. At the sixth acquisition, Gd-DTPA (0.1 mL/kg) was injected intravenously with a rate of 2 mL/second, followed immediately by a bolus injection of 15 mL saline flush at the same rate.

The postcontrast images were obtained using transverse T1-weighted spin-echo sequence with an acquisition protocol similar to that of the precontrast T1-weighted images.

### Data analysis

#### *Dynamic contrast-enhanced-magnetic resonance imaging analysis*

In preprocessing step, motion correction was performed using FSL software (University of Oxford). MCFLIRT function was used to reduce motion artifacts. After motion correction, the slices of interest were extracted using MATLAB (ver. 2008a, The MathWorks TM, Natick, Massachusetts, United States). Data smoothing was

performed using a  $3 \times 3$  mean filter, and a semi-automatic MATLAB code was used for brain extraction.

DCE-MRI data were analyzed using in-house-developed software in MATLAB. The ROIs were determined by mouse pointer-aided method.

#### Dynamic susceptibility contrast-magnetic resonance imaging analysis

In DSC-MRI method, the relationship between the contrast concentration and the signal intensity is expressed by the following equation:

$$C(t) = \Delta R2^* = -k \frac{\ln \frac{S(t)}{S(0)}}{TE} \quad (5)$$

Where  $C(t)$  is the contrast concentration at time  $t$ ,  $\Delta R2^*$  is  $T_2^*$  relaxation rate change ( $=1/T_2^*$ ) due to contrast injection,  $S(t)$  is the signal intensity at time  $t$ ,  $TE$  is echo time, and  $k$  is a constant coefficient.

In this method, cerebral hemodynamic parameters included CBF, and CBV quantified using the standard singular value decomposition convolution algorithm. The constant  $k$  depends on the tissue, contrast agent, magnetic field strength, and imaging sequence.

#### Patients

Twenty-nine patients with brain tumors (male-female ratio, 20:9; mean age, 35.52 years; age range, 21–66 years) were scanned to evaluate the agreement between DSC- and DCE-derived cerebral hemodynamic parameters. The patients underwent DSC- and DCE-MRIs. The study was approved by the local committee for medical research ethics. Informed consent was obtained from all of the patients before the study.

Patients' data were analyzed using the mentioned algorithms. Agreement between DSC- and DCE-derived cerebral hemodynamic parameters was evaluated for

tumor and normal gray and white matters. The normal gray and white matters were segmented based on the structural MRIs of the contralateral normal side. The obtained masks were considered as the region of interests (ROIs) and automatically transferred into the parametric maps [Figure 1]. For evaluating of the tumor regions, the corresponding DSC- and DCE-derived cerebral hemodynamic maps were registered, and the ROI was selected on the enhanced area [Figure 1]. The means of the cerebral hemodynamic indices in the ROIs were determined.

#### Statistical analysis

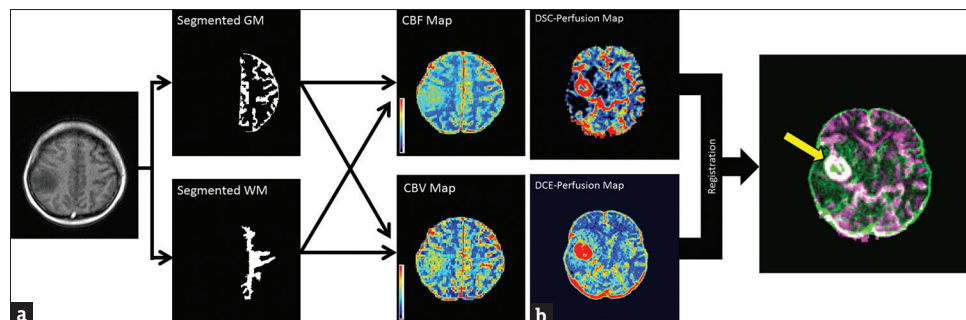
Agreement between DSC- and DCE-derived cerebral hemodynamic indices was determined using the statistical method described by Bland and Altman.<sup>[17]</sup> In this method, a plot of the difference between DSC- and DCE-derived cerebral hemodynamic indices against their mean is used to investigate any possible relationship between the measurement error and the true value. The mean of the two measurements is considered as the true value. The limits of agreement were determined using equation 6:

$$\text{Limits of agreement} = \text{Mean difference} \pm 1.96 \times \text{Standard deviation of the differences} \quad (6)$$

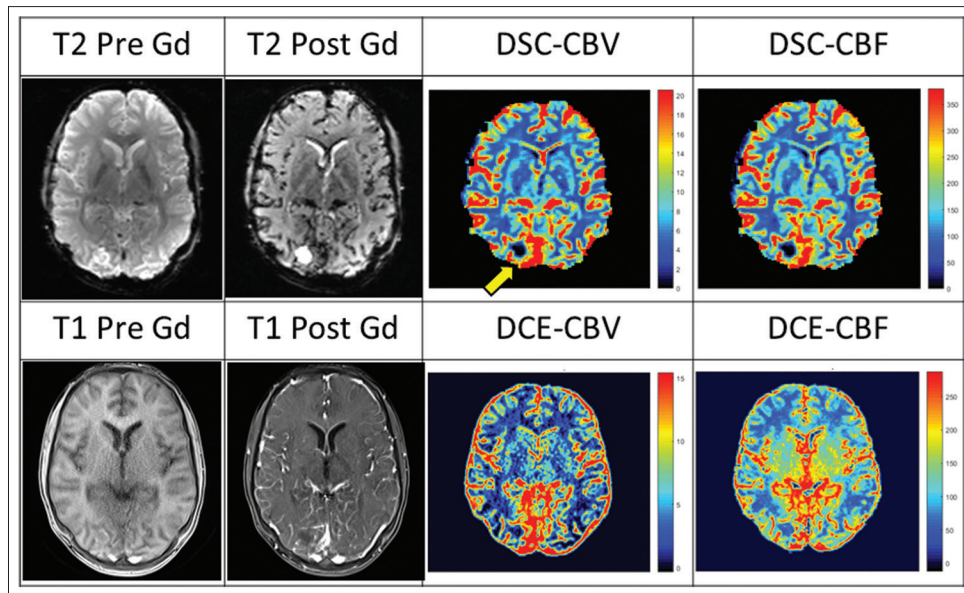
The reliability between DSC- and DCE-derived cerebral hemodynamic indices was determined using the intraclass correlation coefficient (ICC). The intraclass correlation analysis is a popular statistical method to evaluate the agreement between the continuous variables.<sup>[18]</sup> Data analysis was performed using SPSS 16.0 (SPSS Inc., Chicago, IL).

#### RESULTS

For evaluation of the accuracy of the proposed method for quantifying the cerebral hemodynamic indices, CBV and CBF were determined by both DSC- and DCE-MRI methods [Figure 2].



**Figure 1:** (a) Graphical presentation of the region of interest selection procedure for normal gray and white matters. The normal gray and white matters were segmented based on the structural magnetic resonance imagings of the contralateral normal side. The obtained masks were considered as the region of interests and automatically transferred into the parametric maps. The means of the cerebral hemodynamic indices in the region of interests were determined. (b) Graphical presentation of the region of interest selection procedure for tumor tissue. The corresponding dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral hemodynamic maps were registered, and the region of interest was selected on the enhanced area. The means of the cerebral hemodynamic indices in the region of interests were determined. The enhanced area is indicated using the yellow arrow in the registered image.



**Figure 2:** Dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral hemodynamic maps for a 21-year-old man with brain tumor. The region of interest is indicated using the yellow arrow in dynamic susceptibility contrast-derived cerebral blood volume map.

**Table 1: The means and standard deviations of dynamic contrast-enhanced-derived cerebral blood volume (mL/100 g) and dynamic contrast-enhanced-derived cerebral blood flow (mL/100 g.min) for normal gray and white matters and the magnitudes reported in other studies**

Parameter	Study	GM	WM
CBF (mL/100 g/min)	This study	60.53±12.70	32.00±6.00
	Nael et al. <sup>[19]</sup>	68.7±23.1	19.4±6.4
	Reference - Singh et al. <sup>[15]</sup>	64.5±17.3	21.2±2.7
	Singh et al. <sup>[15]</sup>	65.9±23.8	36.5±14.8
	Sourbron et al. - simulation <sup>[14]</sup>	88	24
	Sourbron et al. - measured <sup>[14]</sup>	82	23
	Larsson et al. <sup>[20]</sup>	71.7±16.4	30.6±7.6
	CBV (mL/100 g)	This study	5.01±1.40
Nael et al. <sup>[19]</sup>		2.4±1	1.3±0.6
Reference - Singh et al. <sup>[15]</sup>		5.0±0.7	3.5±0.2
Singh et al. <sup>[15]</sup>		6.9±1.7	2.6±1.1
Bulte et al. <sup>[21]</sup>		3.93±0.90	2.52±0.78
Shin et al. - fast water exchange model <sup>[22]</sup>		5.81	2.73
Shin et al. - no water exchange model <sup>[22]</sup>		3.19	1.78
Ito et al. <sup>[23]</sup>		3.8±0.7	—
Lammertsma and Jones <sup>[24]</sup>		5.9±6.6	2.4±0.3
Sourbron et al. - simulation <sup>[14]</sup>		2.6	1.3
Sourbron et al. - measured <sup>[14]</sup>		2.6	1.3
Larsson et al. <sup>[20]</sup>	6.4±1.8	3.9±1.1	

CBV: Cerebral blood volume, CBF: Cerebral blood flow, GM: Gray matter, WM: White matter

The means and standard deviations of the DCE-derived cerebral hemodynamic indices for normal gray and white matters and the magnitudes reported in other studies are listed in Table 1.

Table 2 summarizes the descriptive statistics of DSC- and DCE-derived cerebral hemodynamic indices for tumor and normal gray and white matters.

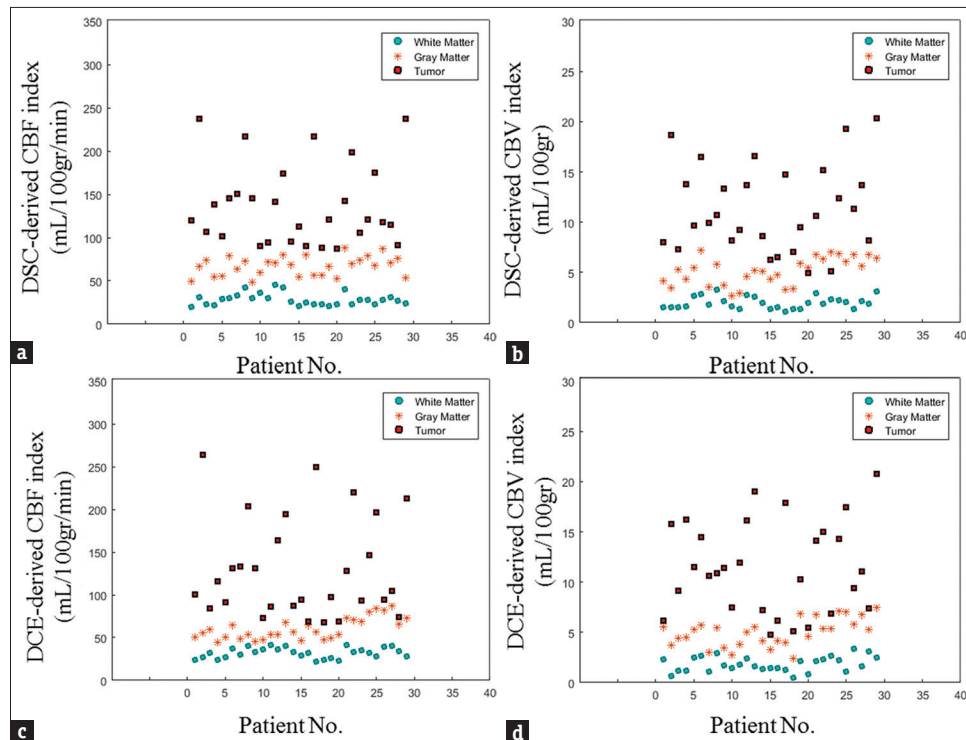
The scattergrams of DSC- and DCE-derived cerebral hemodynamic indices for tumor and normal gray and white matters are plotted to evaluate the distribution of these magnitudes [Figure 3].

The Bland–Altman plots of DSC- and DCE-derived cerebral hemodynamic indices are illustrated [Figure 4].

**Table 2: Descriptive statistics of the dynamic susceptibility contrast and dynamic contrast-enhanced -derived perfusion indices for twenty-nine patients with central nervous system tumor. The normal gray and white matters were segmented and the obtained masks were considered as the region of interests (region of interests). For tumor regions, the region of interest was selected on the enhanced area of the registered image**

Parameter	DCE-derived perfusion indices			DSC-derived perfusion indices		
	Minimum	Maximum	Mean±SD	Minimum	Maximum	Mean±SD
GM						
CBV	2.42	7.52	5.01±1.40	2.69	7.17	5.15±1.36
CBF	44.10	87.64	60.53±12.70	48.97	88.09	67.35±11.22
WM						
CBV	0.52	3.36	1.84±0.74	1.12	3.26	2.00±0.60
CBF	22.07	41.68	32.00±6.00	20.29	45.65	28.64±6.91
Tumor						
CBV	4.79	20.80	11.53±4.57	4.92	20.34	11.35±4.29
CBF	68.32	263.89	130.28±57.95	87.54	236.95	137.08±46.27

SD: Standard deviation, DCE: Dynamic contrast-enhanced, DSC: Dynamic susceptibility contrast, CBV: Cerebral blood volume, CBF: Cerebral blood flow, GM: Gray matter, WM: White matter



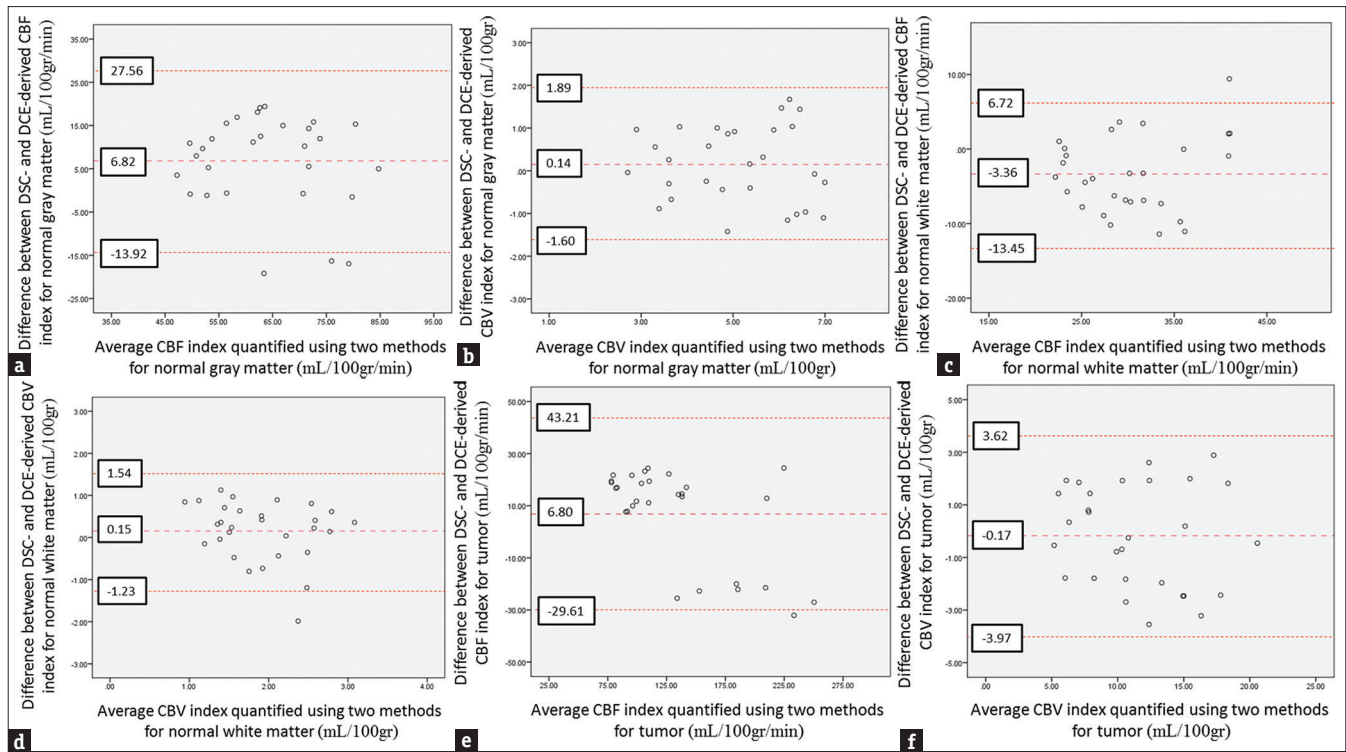
**Figure 3:** Scatterplots of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral hemodynamic indices for tumor and normal gray and white matters. (a) Scatterplot of dynamic susceptibility contrast-derived cerebral blood flow quantities for tumor and normal gray and white matters. (b) Scatterplot of dynamic susceptibility contrast-derived cerebral blood volume quantities for tumor and normal gray and white matters. (c) Scatterplot of dynamic contrast-enhanced-derived cerebral blood flow quantities for tumor and normal gray and white matters. (d) Scatterplot of dynamic contrast-enhanced-derived cerebral blood volume quantities for tumor and normal gray and white matters.

The scatterplots of DSC- and DCE-derived cerebral hemodynamic indices and the equality line are depicted to demonstrate the agreement of the indices [Figure 5]. The intraclass correlation and significant coefficients between DSC- and DCE-derived cerebral hemodynamic indices are listed in Tables 3.

## DISCUSSION

In our study, DCE-derived CBF magnitudes for normal

gray and white matters were  $60.53 \pm 12.70$  mL/100 g/min and  $32.00 \pm 6.00$  mL/100 g/min, respectively. DCE-derived CBV quantities for normal gray and white matters were  $5.01 \pm 1.40$  mL/100 gr and  $1.84 \pm 0.74$  mL/100 gr, respectively. There were statistically significant differences for the DCE-derived cerebral hemodynamic indices between the normal gray and white matters ( $P < 0.001$ ), which were correlating with the known increased vascularity of gray matter. DCE-derived cerebral hemodynamic indices of the



**Figure 4:** Bland–Altman plots of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral hemodynamic indices for tumor and normal gray and white matters. (a) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood flow for normal gray matter. (b) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood volume for normal gray matter. (c) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood flow for normal white matter. (d) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood volume for normal white matter. (e) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood flow for tumor tissues. (f) Bland–Altman plots of dynamic susceptibility contrast- versus dynamic contrast-enhanced-derived cerebral blood volume for tumor tissues. The middle line is the mean difference and the two extreme lines are the limits of agreement.

**Table 3: The intraclass correlation and significant coefficients between dynamic susceptibility contrast and dynamic contrast-enhanced-derived perfusion indices for tumor and normal brain tissues**

	Intraclass Correlation	ICC		<i>F</i> test with true value 0			
		Lower bound	Upper bound	Value	df 1	df 2	Significant
Normal tissue							
CBV	0.914 <sup>a</sup>	0.859	0.948	22.285	57	57	<0.001
CBF	0.877 <sup>a</sup>	0.801	0.926	15.325	57	57	<0.001
Tumor							
CBV	0.905 <sup>a</sup>	0.807	0.954	19.958	28	28	<0.001
CBF	0.937 <sup>a</sup>	0.871	0.970	30.870	28	28	<0.001

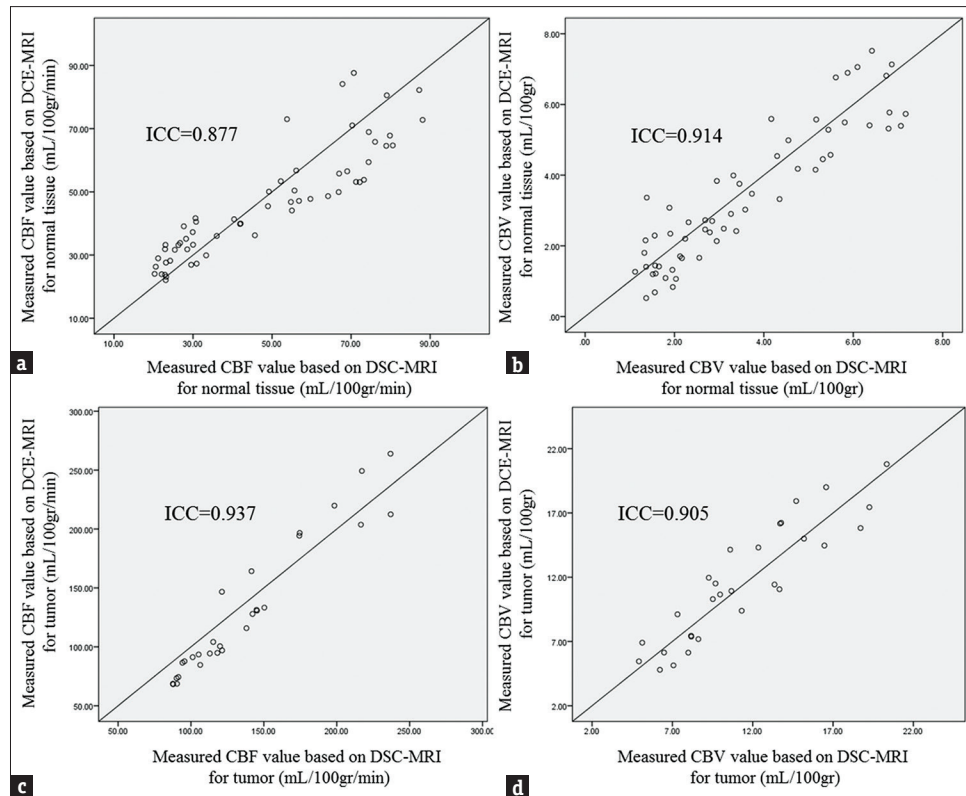
<sup>a</sup>The estimator is the same, whether the interaction effect is present or not. Two-way mixed effects model where people effects are random and measures effects are fixed. ICC: Intraclass correlation coefficient, CI: Confidence interval, CBV: Cerebral blood volume, CBF: Cerebral blood flow

normal gray and white matters were in good agreement with other studies.<sup>[14,15,19-24]</sup> CBV and CBF of the tumor tissues were considerably higher than that of the normal gray and white matters [Figure 3]. There were statistically significant differences for CBV and CBF indices between the tumor and normal tissues ( $P < 0.001$ ).

Concurrent estimation of cerebral perfusion, blood volume, and blood–brain barrier permeability using DCE-MRI was evaluated by Larsson et al.<sup>[13]</sup> and Sourbron et al.<sup>[14]</sup>

Cerebral hemodynamic indices were quantified based on the DCE-MRI data, a method similar to our study. However, the correlation and agreement of these data with the quantified magnitudes using the DSC-MRI were not evaluated.

In this study, Bland–Altman plots showed that the cerebral hemodynamic indices quantified using DCE-MRI data have good agreement with the standard cerebral hemodynamic indices calculated using DSC-MRI



**Figure 5:** (a) The scatterplot of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral blood flow for normal brain tissues. (b) The scatterplot of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral blood volume for normal brain tissues. (c) The scatterplot of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral blood flow for tumor tissues. (d) The scatterplot of dynamic susceptibility contrast- and dynamic contrast-enhanced-derived cerebral blood volume for tumor tissues. For each scatterplot, the equality line was plotted to demonstrate the agreement of the indices, and the intraclass correlation coefficient was expressed.

data. DSC- and DCE-derived CBV magnitudes were compared by Haroon et al.<sup>[25]</sup> For normal brain tissues of nine patients with intra-axial tumors, an excellent correlation ( $r = 0.96$ ,  $P < 0.001$ ) was achieved using pixel-by-pixel scattergrams of values. Leakage-free T1 CBV maps provide comparable biological information with that of the T2\* CBV maps. In Haroon et al. study, the correlation of DSC- and DCE-derived CBV was evaluated for normal brain tissues. The correlation and scattergram of the quantified indices were also studied in tumor area for one patient. The correlation is not a strong statistical analysis to evaluate the agreement between two methods. It seems that a more sophisticated statistical method (such as Bland–Altman analysis) with a larger sample size is needed for determining the agreement between DSC- and DCE-derived cerebral hemodynamic indices.<sup>[18]</sup>

In our study, the ICCs between DSC- and DCE-derived CBF and between DSC- and DCE-derived CBV for normal brain tissues were 0.877 and 0.914, respectively. The ICCs between DSC- and DCE-derived CBF and between DSC- and DCE-derived CBV for tumor tissues were 0.937 and 0.905, respectively. The ICCs were significant at less than 0.001% level

for DSC- and DCE-derived cerebral hemodynamic indices. The ICCs showed that CBV and CBF could accurately be estimated based on the DCE-MRI using a single-compartment model ( $>0.87$ ), and DCE-derived magnitudes are significantly similar to that of the DSC-MRI technique ( $P < 0.001$ ). Furthermore, the acceptable agreements were achieved between DSC- and DCE-derived cerebral hemodynamic indices. These results validated the accuracy of the method to quantify CBV and CBF indices.

The cerebral hemodynamic and permeability indices are conventionally quantified based on different perfusion MRI techniques, separately. Both parameter categories could be extracted using single-dose DCE perfusion MRI. DCE-derived cerebral hemodynamic indices could be considered as a promising method to evaluate brain tumors. DCE-MRI permits multiparametric characterization of tumor vasculature which is not affected by susceptibility artifacts.<sup>[11,13]</sup>

## CONCLUSION

In this study, the agreement between DSC- and DCE-derived cerebral hemodynamic indices was evaluated for tumoral and normal brain tissues. Cerebral



hemodynamic parameters including CBV and CBF were calculated from DCE-MRI data accurately. Based on the proposed method, CBV and CBF could be determined based on the DCE-MRI using a single-compartment model with fair agreement with that of DSC-MRI and a more comprehensive collection of the physiological parameters achieved for tumor evaluations.

### Acknowledgment

Authors would like to thank the research affair of Medicine faculty of Tehran University of Medical Sciences, Medical Imaging Center of Imam Khomeini Hospital (Tehran), and participants in the study.

### Financial support and sponsorship

This research has been supported by Tehran University of Medical Sciences, grant number 94-01-30-27363, 2015/04/19.

### Conflicts of interest

There are no conflicts of interest.

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