# Morphometry of Intratumoral Enhancement Patterns on 4D Spectral Images for Differential Diagnosis of Breast Tumors in Dynamic Contrast-enhanced MRI

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## Introduction

The characterization of intratumoral enhancement patterns in dynamic contrast enhanced (DCE) MRI is important for the non-invasive diagnosis of breast tumors, and yet remains challenging due to the spatio-temporal complexities in the tumor uptake and clearance kinetics of contrast agents. A recent study has demonstrated the superiority of capturing the spatio-temporal enhancement pattern in accurate tumor characterization for breast MR-based diagnosis [1]. However, there were limitations that its analysis was based on a selected 2D image and tumors were segmented from a manually delineated contour. Here we present an integrated framework for 3D semi-automatic tumor segmentation and characterization of 3D spatio-temporal properties of intratumoral enhancements by using a quantitatively combined approach of tumor kinetics and morphometry. The influence of the combined spatio-temporal enhancement features on the performance is tested for differentiation of benign and malignant tumors in breast DCE-MRI.

### **Materials and Methods**

Twenty female patients were recruited to this study and one primary lesion from each patient was used for analysis. DCE MRI was carried out on a 1.5T scanner (Magnetom Sonata; Siemens, Erlangen, Germany) using T1-weighted 3D FLASH sequence: 448×448, TE=1.83 msec, TR=4.9 msec, flip angle=12°, slice thickness=1-1.5 mm without a gap, temporal resolution=84 sec per 96-111slices. One pre-enhanced and four post-enhanced series following a bolus injection of Gd-DTPA (Magnevist, Schering, Berlin, Germany; 0.1 mmol/kg at 2 ml/sec for 5 sec) were acquired as unilateral sagittal images. After 3D rigid registration of MR-time-series, our proposed perfusion index (PI) map was generated for enhanced tumor contrast, which widens the difference of variations in enhancement kinetic features between a lesion and normal parenchyma and allows effective identification of tumors [2]. On the PI map, tumor segmentation was performed by using Otsu thresholding, 3D region growing algorithm, hole-filling and iterative morphological erosion and dilation. Temporal enhancement (TE) features were extracted by singular value decomposition (SVD) of a lower-triangular Toeplitz matrix representing convolution operation, and their SVD-based eigenvalue (EV) maps were generated. The spatial variations of EVs within each tumor were captured by 3D geometric moment invariants (GMIs) [3]. The binary classification for tumor differentiation was performed by Least Square Support Vector Machines (LS-SVM) with a Radial Basis Function (RBF) kernel. Leave-one-out cross-validation was used in the classification process, which was repeated fifty times by iteratively retraining the LS-SVM in order to obtain an average test result. The average test performance was measured by Receiver Operating Characteristic (ROC) curve analysis.

#### Results

All of the malignant lesions were 10 invasive ductal carcinomas and the histological distributions of the benign lesions were 2 fibroadenomas, 3 papilomas, 3 phylloides tumors, 1 hamartoma and 1 atypical hyperplasia. The PI map successfully identified the margins of tumors in most instances and the segmentation results showed good agreement with tumor boundaries. A total of twenty-four SVD-based GMI features were extracted from each tumor to represent the spatio-temporal properties within the tumor. In an evaluation experiment with 10 malignant and 10 benign cases, the average test performance showed the area under ROC curve (AUC) of 0.795 (95% confidence interval, 0.768-0.819) with sensitivity and specificity of 96.4% (95% confidence interval, 94.4-97.9%) and 75.6% (95% confidence interval, 71.5-79.3%), using LS-SVM RBF applied to the SVD-based GMI features.

## Conclusion

With this pilot study for MR-based computer-aided diagnosis (CAD) of breast tumors, we have shown that: (1) 3D tumor segmentation problem of breast tumors in DCE-MRI can be solved effectively in the way that made the best use of tumor perfusion characteristics; (2) Combination of SVD and 3D GMI yields a promising descriptor to characterize and differentiate the spatio-temporal enhancement patterns within tumors in breast DCE-MRI.

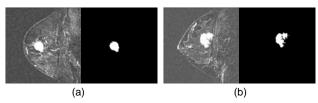


Figure 1. Examples of the PI map (left) and tumor segmentation outcome (right) for (a) malignant (invasive ductal carcinoma) and benign (fibroadenoma) tumors

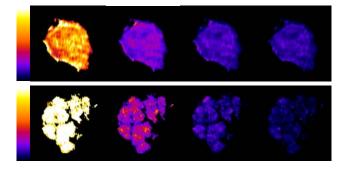


Figure 2. Examples of SVD-based EV maps for malignant (top) and benign (bottom) tumors: the 1st, 2nd, 3rd and 4th EV maps from left to right

## References

- [1] Y Zheng, S Englander, et al. IEEE ISBI 2007: 520-523.
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- [3] D Xu, H Li, Pattern Recognition 2008: 41, 240-249.

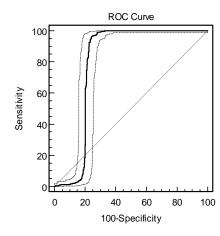


Figure 3. ROC curve of SVD-based GMI features for tumor differentiation (dashed line: 95% confidence limits)