Voxel-based comparison of Dynamic Contrast-Enhanced MRI and FDG-PET in head-and-neck cancer

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Purpose
Both FDG-PET and dynamic contrast-enhanced (DCE-) MRI can be useful to characterize radio-resistant parts of tumors in head-and-neck cancer. Previous studies carried out at patient-level showed mixed results about the correlations between standardized uptake values (SUV) and DCE pharmacokinetic parameters. For primary tumors a significant correlation was reported between SUV and one of the DCE parameters (ve)1, whereas no correlation was obtained for nodal metastases2. In this study we test the hypothesis that a voxel-based comparison will result in stronger correlations, because tumor heterogeneity is taken into account.

Methods
Twenty-one patients with head-and-neck cancer were retrospectively selected for whom a planning CT, FDG-PET/CT and DCE-MRI exam prior to radiotherapy were available. FDG-PET was performed one hour after injection of FDG (voxel size 2x2x2 mm3). SUV maps were used for subsequent analysis. The DCE-MRI exam was performed on a 1.5T MR scanner with flex coils using a transversal 3D spoiled-gradient echo sequence (voxel size 3x3x6 mm3, 29 slices, TR/TE 4/0/1.16 ms, flip angle 15 degrees; dynamic scan time 2.5 s, no. of volumes 60). The contrast agent (Dotarem, 15 mL) was injected at a flow rate of 3 mL/s using a power injector followed by a saline flush. The signal intensities were first converted to gadolinium concentration using a pre-contrast T1 map estimated from a variable flip angle series of 3, 6, 10, 20 and 30 degrees. The extended Tofts model was fitted to the concentration time curves for estimation of the pharmacokinetic parameters (Ktrans, kep, vp, and ve)3. A population-based arterial input function was used derived from the combined magnitude and phase data in the external carotid arteries in the neck of the patients. Patients were positioned in their radiotherapy mask during all examinations, except for six FDG-PET scans. All images were rigidly registered to the planning CT using a large region of interest containing the spinal cord, mandible and part of the skull. SUV and DCE parameter maps were compared at voxel-level and at patient-level. For the comparison at voxel-level Spearman’s correlation coefficient (ρ) was calculated between SUV and each of the DCE parameter values for all voxels within the gross tumor volume (GTV) for each patient separately. A two-sided Wilcoxon signed-rank test was used to test whether the average correlation coefficient across all patients was significantly different from zero. For the comparison at patient-level, ρ was calculated between median SUV values and median DCE values across all patients.

Results
The average median SUV value within the GTV was 4.6 ± 1.6, whereas the average median pharmacokinetic values of the Tofts model were 0.48 ± 0.16 min−1 for Ktrans, 1.30 ± 0.27 min−1 for kep, 0.34 ± 0.12 for vp, and 0.06 ± 0.01 for ve. The correlation between SUV and Ktrans was higher and significant at voxel-level compared to patient-level, ρ = 0.25 vs. ρ = 0.16. The same result was obtained for the correlation between SUV and kep (ρ = 0.42 at voxel-level, ρ = 0.19 at patient-level). These correlations were not dependent on the size of the tumor. vp and ve were not significantly correlated to SUV at both levels (Table 1). Fig. 1 illustrates the positive correlation between SUV and Ktrans and kep. However, it should be noted that the results are variable among patients (Fig. 2).

Discussion
The average SUV and DCE parameter values were in accordance with previously reported primary tumor values1. Also the results at patient-level were comparable to previous results, except that Bisdas et al. reported a significant correlation between SUV and vp1. At voxel-level our results show a positive correlation between SUV (indicative for metabolic activity) and Ktrans (representing vessel permeability) and kep (representing wash-out).

Conclusion
At voxel-level the correlations within the GTV for head-and-neck cancer were higher between SUV and Ktrans and kep compared to the results at patient-level. Therefore, tumor heterogeneity should be taken into account. However, the correlations are moderate indicating that FDG-PET and DCE-MRI provide partly complementary information.

![image](image1.png)

**Fig. 1** Patient example of comparison between SUV, Ktrans, and kep. White contour line indicates the GTV. For this patient ρ between SUV and Ktrans was 0.30, whereas ρ between SUV and kep was 0.66.

![image](image2.png)

**Fig. 2** Spearman’s correlation coefficients between SUV and DCE parameters at voxel-level for all patients.

**Table 1** Spearman’s correlation coefficients for comparison between SUV and DCE parameters at voxel-level and at patient-level (*p* < 0.01)

<table>
<thead>
<tr>
<th>Parameter Comparison</th>
<th>Patient-Level Correlation</th>
<th>Patient-Level Range</th>
<th>Patient-Level <em>p</em> Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>SUV vs. Ktrans</td>
<td>0.25* (0.25 – 0.52)</td>
<td>0.16</td>
<td>0.01</td>
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<tr>
<td>SUV vs. kep</td>
<td>0.42* (0.31 – 0.66)</td>
<td>0.19</td>
<td>0.03</td>
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<tr>
<td>SUV vs. vp</td>
<td>0.013 (-0.40 – 0.43)</td>
<td>0.26</td>
<td>0.05</td>
</tr>
<tr>
<td>SUV vs. ve</td>
<td>0.13 (-0.25 – 0.35)</td>
<td>0.22</td>
<td>0.01</td>
</tr>
</tbody>
</table>

References