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Research article

# Improved clinical workflow for simultaneous whole-body PET/MRI using high-resolution CAIPIRINHA-accelerated MR-based attenuation correction



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## ABSTRACT

*Purpose:* To explore the value and reproducibility of a novel magnetic resonance based attenuation correction (MRAC) using a CAIPIRINHA-accelerated T1-weighted Dixon 3D-VIBE sequence for whole-body PET/MRI compared to the clinical standard.

*Methods*: The PET raw data of 19 patients from clinical routine were reconstructed with standard MRAC (MRAC<sub>stal</sub>) and the novel MRAC (MRAC<sub>caipi</sub>), a prototype CAIPIRINHA accelerated Dixon 3D-VIBE sequence, both acquired in 19 s/bed position. Volume of interests (VOIs) for liver, lung and all voxels of the total image stack were created to calculate standardized uptake values (SUV<sub>mean</sub>) followed by inter-method agreement (Passing-Bablok regression, Bland-Altman analysis). A voxel-wise SUV comparison per patient was performed for intra-individual correlation between MRAC<sub>std</sub> and MRAC<sub>caipi</sub>. Difference images (MRAC<sub>std</sub>-MRAC<sub>caipi</sub>) of attenuation maps and SUV images were calculated. The image quality of in/opposed-phase water and fat images obtained from MRAC<sub>caipi</sub> was assessed by two readers on a 5-point Likert-scale including intra-class coefficients for inter-reader agreement.

Results: SUV<sub>mean</sub> correlations of VOIs demonstrated high linearity (0.95 < Spearman's rho < 1, p < 0.0001, respectively), substantiated by voxel-wise SUV scatter-plots ( $1.79 \times 10^8$  pixels). Outliers could be explained by different physiological conditions between the scans such as different segmentation of air-containing tissue, lungs, kidneys, metal implants, diaphragm edge or small air bubbles in the gastrointestinal tracts that moved between MRAC acquisitions. Nasal sinuses and the trachea were better segmented in MRAC<sub>caipi</sub>. High-resolution T1w Dixon 3D VIBE images were acquired in all cases and could be used for PET/MRI fusion. MRAC<sub>caipi</sub> images were of high diagnostic quality (4.2  $\pm$  0.8) with 0.92-0.96 intra-class correlation.

*Conclusions:* The novel prototype  $MRAC_{caipi}$  extends the value for attenuation correction by providing a high spatial resolution DIXON-based dataset suited for diagnostic assessment towards time-efficient whole-body PET/MRI.

### 1. Introduction

The fully integrated combination of positron emission tomography (PET) and magnetic resonance imaging (MRI), simultaneous PET/MRI, facilitates the fusion of molecular information provided by PET with high soft tissue contrast, numerous functional imaging methods and reduced radiation exposure by MRI compared to PET/CT [1]. For accurate quantitative PET-imaging, attenuation correction (AC) is the

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Abbreviations: AC, attenuation correction; MRI, magnetic resonance imaging; PET, positron emission tomography; VOI, volume of interest; VOI<sub>liver</sub>, VOI in liver; VOI<sub>totab</sub> VOI of total image stack; SUV, standardized uptake value; µmap, MR-based attenuation map

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#### Table 1

| Patient demographics included in the quantitative analysis. | The thorax was completely included in 9 | 9 patients, of which 4 received a whole-body examination. |
|---|---|---|
|---|---|---|

|    | age | gender | MBq | height | weight | diagnosis      | bed positions        | tracer                        |
|----|-----|--------|-----|--------|--------|----------------|----------------------|-------------------------------|
| 1  | 48  | m      | 114 | 177    | 81     | NET (pancreas) | Thorax-pelvis (3BED) | 68Ga-DOTATOC                  |
| 2  | 30  | f      | 113 | 160    | 46     | NET (appendix) | Abdomen (2BED)       | <sup>68</sup> Ga-DOTATOC      |
| 3  | 62  | m      | 121 | 187    | 108    | PCA            | Abdomen (2BED)       | <sup>68</sup> Ga-PSMA-HBED-CC |
| 4  | 59  | m      | 141 | 175    | 63     | PCA            | Abdomen (2BED)       | <sup>68</sup> Ga-PSMA-HBED-CC |
| 5  | 74  | m      | 144 | 182    | 86     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 6  | 66  | m      | 115 | 178    | 88     | PCA            | Head-pelvis (5BED)   | <sup>68</sup> Ga-PSMA-HBED-CC |
| 7  | 76  | m      | 105 | 173    | 87     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 8  | 68  | m      | 259 | 170    | 70     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 9  | 53  | m      | 214 | 183    | 80     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 10 | 64  | m      | 71  | 180    | 80     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 11 | 46  | m      | 232 | 189    | 83     | NHL            | Head-pelvis (5BED)   | <sup>18</sup> F-FDG           |
| 12 | 74  | m      | 282 | 177    | 76     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 13 | 56  | m      | 243 | 176    | 95     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 14 | 49  | m      | 227 | 180    | 80     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 15 | 66  | m      | 247 | 170    | 68     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 16 | 61  | m      | 158 | 178    | 82     | PCA            | Thorax-pelvis (3BED) | <sup>68</sup> Ga-PSMA-HBED-CC |
| 17 | 66  | m      | 250 | 180    | 80     | PCA            | Head-pelvis (5BED)   | 68Ga-PSMA-HBED-CC             |
| 18 | 71  | f      | 371 | 164    | 55     | BC             | Head-pelvis (5BED)   | <sup>18</sup> F-FDG           |
| 19 | 70  | m      | 238 | 170    | 75     | PCA            | Thorax-pelvis (3BED) | 68Ga-PSMA-HBED-CC             |

MBq: injected activity in megabecquerel; NET: neuroendocrine tumor; PCA: prostate carcinoma; BC: bronchial carcinoma; NHL: Non-Hodgkin-Lymphoma

#### Table 2

| Comparison of r | most relevant sequence | parameters of MRAC <sub>std</sub> | versus MRAC <sub>caipi</sub> . | CAIPIRINHA-acceleration | settings are given in methods. |
|-----------------|------------------------|-----------------------------------|--------------------------------|-------------------------|--------------------------------|
|-----------------|------------------------|-----------------------------------|--------------------------------|-------------------------|--------------------------------|

| Parameter                                    | TE1/TE2/TR (ms)                 | Resolution (mm <sup>3</sup> )                              | Acquisition Time (s) | FOV (mm <sup>2</sup> )  | Matrix  |
|--|---------------------------------|--|----------------------|---|---|
| MRAC <sub>std</sub><br>MRAC <sub>caipi</sub> | 1.23/2.46/3.6<br>1.24/2.47/4.26 | $4.1 \times 2.6 \times 3.1$<br>$1.9 \times 1.4 \times 3.2$ | 19<br>19             | $\begin{array}{l} 500 \times 328 \\ 500 \times 500 \end{array}$ | $\begin{array}{c} 192 \times 126 \\ 264 \times 352 \end{array}$ |

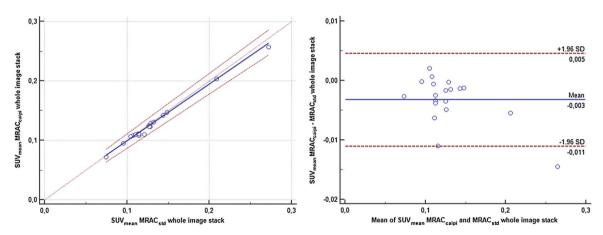


Fig. 1. Passing-Bablok regression (left column) and Bland-Altman plot (right column) for inter-method correlation between VOIs of the whole image stack for SUV<sub>mean</sub>. Refer to Fig. 1 (Supplementary) for voxel-wise SUV comparison. A high linearity for each plot is demonstrated indicating strong agreement between both methods. MRAC<sub>std</sub> is plotted on the x-axis, MRAC<sub>caipi</sub> on the y-axis.

clinical standard to correct for tissue-dependent differences in photon attenuation. In the case of PET/CT, an axial CT scan provides robust attenuation coefficients reflecting tissue density per voxel after conversion to the 511 keV energy of the PET photons [2]. To reduce radiation exposure, a low-dose scan between 30 and 80 mAs is sufficient. However, the downside of this method is the poorer contrast of the lowdose CT scan compared to a normal-dose contrast-enhanced CT scan. Hence, low-dose CT has intrinsic limitations for depicting the anatomy clearly due to low soft tissue contrast, i.e. to correlate a PET-positive finding with the underlying tissue and to differentiate it from adjacent structures such as vessels. In contrast, MRI potentially allows for better correlation images of the underlying anatomy ('landmarking') in areas where high soft tissue contrast is demanded such as brain, head/neck, abdomen and extremities. In contrast to PET/CT, the attenuation map for PET data using MRI is based on automatic tissue classification and assignment of attenuation values for each tissue type [3]. MRAC using

T1-weighted Dixon 3D-VIBE-images (MRACstd) is a valid approximation and produces reliable results compared to CT-based AC in clinical routine for <sup>18</sup>F-FDG, based on previous work including data on more than 2300 patients [4–6].  $MRAC_{std}$  features a comparable poor spatial resolution if re-formatted to axial orientation. However, it can be acquired in only 19 s per bed position. The generalized autocalibrating partially parallel acquisition (GRAPPA) algorithm yields an accelerated acquisition of MRI images [7] by acquiring fewer k-space lines. The method then compensates for the undersampled k-space data by exploiting the redundant information of different receiver coil channels. GRAPPA can be extended using controlled aliasing in parallel imaging results in higher acceleration (CAIPIRINHA) to improve the parallel imaging results by exploiting the coil sensitivity patterns more efficiently and, thus, allow for larger acceleration factors [8,9]. This technique facilitates high-resolution images for breath-hold examinations [10] in short acquisition times. However, in the case of PET/MRI,

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