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Biodistribution and Radiation Dosimetry for a Probe Targeting Prostate-Specific Membrane Antigen for Imaging and Therapy

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Abstract

Prostate-specific membrane antigen (PSMA) is a promising target for diagnosis and treatment of prostate cancer. EuK-Subkff-⁶⁸Ga-DOTAGA (⁶⁸Ga-PSMA Imaging & Therapy [PSMA I&T]) is a recently introduced PET tracer for imaging PSMA expression in vivo. Whole-body distribution and radiation dosimetry of this new probe were evaluated.

Methods—Five patients with a history of prostate cancer were injected intravenously with 91– 148 MBq of 68 Ga-PSMA I&T (mean \pm SD, 128 ± 23 MBq). After an initial series of rapid whole-body scans, 3 static whole-body scans were acquired at 1, 2, and 4 h after tracer injection. Time-dependent changes of the injected activity per organ were determined. Mean organ-absorbed doses and effective doses were calculated using OLINDA/EXM.

Results—Injection of 150 MBq of ⁶⁸Ga-PSMA I&T resulted in an effective dose of 3.0 mSv. The kidneys were the critical organ (33 mGy), followed by the urinary bladder wall and spleen (10 mGy each), salivary glands (9 mGy each), and liver (7 mGy).

Conclusion—⁶⁸Ga-PSMA I&T exhibits a favorable dosimetry, delivering organ doses that are comparable to (kidneys) or lower than those delivered by ¹⁸F-FDG.

Keywords

PET; PSMA; ⁶⁸ Ga; dosimetry; prostate cancer	

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DISCLOSURE

No other potential conflict of interest relevant to this article was reported.

More than 220,000 cases of prostate cancer will be diagnosed in 2015, accounting for 26% of all new cancer cases in the United States (1). Accurate staging and restaging remain a diagnostic challenge. Visualization of tumor metabolism with PET is gaining increasing interest particularly in the management of patients with biochemical disease relapse (2). Commonly used PET tracers include the glucose analog ¹⁸F-FDG for imaging the tumor glucose metabolism, especially in dedifferentiated tumors (3); amino acid transport probes such as ¹¹C-methionine (4) and the synthetic amino acid probe ¹⁸F-1-amino-3-fluorine 18-fluorocyclobutane-1-carboxylicacid (5); as well as probes of lipid metabolism such as choline (6) and acetate (7).

More recently, tracers targeting the prostate-specific membrane antigen (PSMA) have attracted the attention of clinicians and imaging specialists. The overexpression of PSMA in prostate cancer cells (8) is associated with the adverse outcome of prostate cancer patients (9). Glu-NH-CO-NH-Lys-(Ahx)-[⁶⁸Ga-(HBED-CC)] (⁶⁸Ga-DKFZ-PSMA-11), a ⁶⁸Ga-labeled PSMA-targeted ligand (10), has been studied by several groups (11–17). More recently, Wester et al. introduced a different PSMA ligand, EuK-Subkff-⁶⁸Ga-DOTAGA (⁶⁸Ga-PSMA I&T), that can also be labeled with ¹⁷⁷Lu (18) and can thus potentially serve as the therapeutic arm of a new theranostic approach in prostate cancer.

Because the whole-body distribution and dosimetry of ⁶⁸Ga-PSMA I&T must be determined before its clinical translation, we here report the dynamic biodistribution and dosimetry of this new PET imaging probe in a small cohort of patients with primary or recurrent prostate cancer.

MATERIALS AND METHODS

Research Design and Subjects

⁶⁸Ga-PSMA I&T was administered in compliance with The German Medicinal Products Act, AMG §13 2b, and in accordance with the responsible regulatory body (Regierung von [i.e., Government of] Unterfranken). The data analysis was presented to the ethics committee of the Universitätsklinikum Würzburg, and the need for a formal review was waived.

Five patients (age, 59.1–69.9 y; mean age \pm SD, 65.1 ± 4.1 y) with prostate cancer underwent 68 Ga-PSMA PET/CT. One patient had biopsy-proven primary prostate cancer, 3 had biochemical relapse, and 1 underwent restaging after the start of androgen-deprivation therapy. The mean prostate-specific antigen (PSA) level in the 3 patients with biochemical relapse was 2.9 ± 3.3 ng/mL, and the time interval between primary diagnosis and biochemical relapse was 2.2 ± 0.9 y. Detailed patient characteristics are presented in Table 1. Safety was assessed by monitoring adverse events up to 5 h after administration of 68 Ga-PSMA I&T.

Preparation of PSMA-Targeting Probe ⁶⁸Ga-PSMA I&T

⁶⁸Ga-PSMA I&T was synthesized using a fully automated, good manufacturing practice—compliant procedure with a GRP module (SCINTOMICS GmbH) connected to a ⁶⁸Ge/⁶⁸Ga generator (Cyclotron Co. Ltd.) and equipped with a disposable single-use cassette kit

(ABX). A standardized labeling sequence with 40 µg of unlabeled PSMA I&T (SCINTOMICS GmbH) was used as previously described (19).

Before application, the radiopharmaceuticals were analyzed according to the monographs 2462 (Gallium Chloride) and 2482 (Gallium Edotreotide) of the European Pharmacopoeia by analytic high-performance liquid chromatography. Analytic high-performance liquid chromatography was performed on a SCINTOMICS system equipped with a RP-18 column (Nucleosil, 125×4.6 mm; CS-Chromatographie). The eluent had a linear gradient from 100% water (0.1% trifluoroacetic acid) to 100% MeCN (0.1% trifluoroacetic acid) over 20 min. The radiochemical purity of the tracer was determined with a miniGITA thin-layer chromatography scanner (Raytest) using Varian silica gel–impregnated glass fiber sheets and 0.1 M sodium citrate as eluent.

PET Imaging

All ⁶⁸Ga-PSMA I&T scans were obtained on a dedicated PET/CT scanner (Biograph mCT 64; Siemens Medical Solutions) after at least 4 h of fasting. Low-dose CT scans for attenuation correction were acquired (35 mAs, 120 keV, a 512 × 512 matrix, 5-mm slice thickness with a total of 201 slices, increment of 30 mm/s, rotation time of 0.5 s, and pitch of 0.8). The imaging field ranged from the head (patient 1; [P1]) or the base of the skull (patients 2–5 [P2–5]) to the proximal thighs. Immediately after injection, the PET imaging sequences started with a series of 3 (P1) or 4 (P2–5) 300-s whole-body scans.

Subsequently, 3 static whole-body scans encompassing 6–7 bed positions were acquired at 1, 2, and 4 h after tracer injection. All data were decay-corrected to the starting time of each individual scan. All PET images were corrected for photon attenuation, dead-time, random events, and scatter. The PET scanner is periodically checked for calibration accuracy as part of quality control according to published guidelines (20) and is accredited by European Association of Nuclear Medicine Research Ltd.

Imaging and Dosimetry

All images were analyzed qualitatively by experienced nuclear physicians for the presence of suggestive lesions. To determine the time point providing the best sensitivity and optimal lesion-to-background contrast, we first examined all available datasets to identify target lesions.

Semiquantitative analysis of visually detectable lesions was done by 3-dimensional volumes of interest (VOIs). Corresponding maximum standardized uptake values (SUV_{max}) were recorded as a function of time. The European Association of Nuclear Medicine recommendations for good dosimetry reporting were used (21).

Full organ segmentation was performed by a single observer on CT images and PET images for all segmentable organs (e.g., gallbladder, heart, kidneys, spleen, liver, salivary glands, and bladder) and lesions using the E.SOFT software VA60C (Siemens Medical Solutions). A whole-body VOI was also drawn. Bone marrow dosimetry was derived from CT-based VOIs placed over lumbar vertebrae 2–4. From the coregistered PET images, average organ activity per volume in kBq/mL was obtained for each frame. The total activity in the respective VOI

was calculated subsequently by multiplying the average organ activity per volume with the respective volume taken from the CT images. Because the scanning was done only from the head or base of the skull to the mid thigh, the total activity in the whole body (measured portions and the nonmeasured portions of the lower extremities and the head) was estimated by extrapolating the respective whole-body time—activity curves to the injection time and by calculating a corresponding scaling factor. This scaling factor was applied to all whole-body measurements of the same patient for calculating the time-integrated activity coefficients. All measured organ activities were normalized to the respective total injected activities.

The integration of time–activity curves was performed using the software NUKFIT (22). For this investigation, a systematic error in activity quantification of 10% was assumed. The time–activity curves of the urinary bladder contents were integrated using a trapezoidal integration and assuming physical decay after the last data point.

To assess differences in bone marrow time-integrated activity coefficients while applying different methods, the time-integrated activity coefficient for the bone marrow was derived from 2 methods proposed by Ferrer et al. for radioimmunotherapy (23). Particularly for the blood-based method it is not conclusively proven which red marrow—to—blood ratio (RMBLR) is optimal. For [177Lu-DOTA0, Tyr3]octreotate, the activity in bone marrow aspirates correlates with the activity measured in blood (24), suggesting a RMBLR of 1. For ⁶⁸Ga-PSMA I&T, no such data are available. As, in a previous study with ⁶⁸Ga-pentixafor (25), the agreement between the image-based method and the blood-based method was better when the RMBLR for radioimmunotherapy was applied, we decided to apply the same method for this compound.

The individual time-integrated activity coefficients were used for calculating effective doses for the standard 70-kg adult male model using OLINDA/EXM (26) for each patient separately. Entering the mean values of the time-integrated activity coefficients into OLINDA/EXM provided organ-absorbed doses and effective doses (based on tissue-weighting factors from International Commission on Radiological Protection [ICRP] publication 60 (27)) for a standard patient. The lesion-absorbed doses for the salivary glands were calculated from the corresponding time-integrated activity coefficients with the OLINDA/EXM unit density sphere model for each of the glands separately, taking into account the individual gland volume. The resulting absorbed doses were averaged to obtain a mean absorbed dose. This model better represents more closely the real energy deposition pattern assuming that the energy is deposited in a uniform sphere of 85 g, which represents the salivary glands according to ICRP 89 (28). SDs were calculated using Excel (Microsoft).

In addition, the effective dose according to ICRP 103 (29) has been calculated despite the fact that the organ-absorbed dose rates per unit activity values for radiopharmaceuticals based on the latest ICRP voxel phantoms in ICRP 110 (30) have not been published yet (31). As there was no female patient, a sex-specific calculation has not been performed.

RESULTS

Radioligand and Patients

The administered amount of 68 Ga-PSMA I&T was less than 20 µg. The overall injected activity (radiochemical purity > 98%) ranged from 91 to 148 MBq (mean \pm SD, 128 \pm 23 MBq) with a specific activity greater than 5 MBq/µg. Activity remaining in the injection syringe was quantified and considered. Injection of 68 Ga-PSMA I&T was well tolerated by all subjects. No side effects or changes in vital signs were observed during the study.

Biodistribution

On qualitative image analysis, all but 1 patient (P2; PSA value, 0.5 ng/mL) presented suggestive focal lesions. The patient with biopsy-proven but untreated prostate cancer (P3; PSA value, 101.2 ng/mL) presented increased uptake in the prostate, mediastinal and iliac lymph nodes, and lumbar spine (Fig. 1). In the patient recently started on androgen ablation therapy, tracer uptake was observed in the prostate as well as in the lumbar spine lesions (P5). In 2 patients with biochemical recurrence (PSA values, 1.6 and 6.7 ng/mL), pathologic tracer uptake was identified in a rib lesion (P1) and an iliac lymph node (P4), respectively (Table 1).

The biodistribution of ⁶⁸Ga-PSMA I&T was determined for all major organs in all patients. Figures 1 and 2 depict whole-body maximum-intensity-projection images of 2 subjects (P1 and P3) at different time points. Figure 3 shows the time–activity curves for various organs and the blood (per liter of blood) expressed as percentage injected activity for the same patient as in Figure 2 (P1).

The highest uptake in any organ was observed in the liver of patient 5, with 13.3% of the injected activity after 5 min. The data at later time points for this patient showed a rapid washout phase, with an uptake of less than 0.8% after 4 h. For all patients the mean maximal liver uptake was $10.6\% \pm 1.7\%$. The mean uptake in the kidneys of all patients varied between $3.2\% \pm 0.6\%$ and $0.3\% \pm 0.2\%$ about 20 min and 4 h after administration, respectively. Significant tracer uptake was also observed in the heart, with a mean uptake of $2.5\% \pm 0.6\%$ after 5 min, declining to less than 0.1% after 4 h.

The gallbladder exhibited low uptake, compared with other organs. The highest uptake in the urinary bladder before voiding (5.6%) was observed in P4. All other patients showed bladder uptake of less than 5% of the injected activity. All patients voided for the first time 40–60 min after administration of the radioligand. Comparable time–activity curves were recorded in all other patients.

Image Analysis

Immediately after injection there was only minimal urinary bladder activity. Thus, uptake in the prostate or prostate bed is least affected by activity spillover (Figs. 1 and 2). However, lymph node metastases adjacent to blood vessels can be detected better on later images because of the high blood-pool activity at early time points. For the remainder of the body, image acquisition after 1 h provided excellent image contrast.

 SUV_{max} increased in all but 1 lesion from early to late imaging (Fig. 4), but the visual impression was better after 1 h. The SUV_{max} for P4, derived from subcentimeter lymph nodes prone to partial-volume effects, was found at 1 h. In none of the lesions was there a washout observed until the last scanning time point.

For assessing quantitatively the optimal time point for scanning the relative uptake ratios of the lesion activity to the activity in the whole body was assessed for the time points 30 min, 1 h, and 2 h after administration and was highest after 2 h (mean uptake ratios for all lesions compared with t = 30 min, 1:1.4:1.9).

Because the lesion volumes were small (<3 mL), absorbed doses were not calculated for these lesions as there is no reliable method for determining the uptake in these volumes due to the partial-volume effect. Data on the partial-volume effect for ⁶⁸Ga and small volumes are shown by, for example, Preylowski et al. (32).

Dosimetry

Time-integrated activity coefficients of segmented organs were calculated for each patient individually. In addition, the mean values for all patients are given (Table 2). All corresponding errors for calculating the individual time-activity curves were less than 5%. The corresponding values are not shown in Table 2 as they were much smaller than the SD when comparing all patients (Table 2, column 7). Therefore, all errors were neglected for the calculation of the mean absorbed organ doses.

The highest number of disintegrations per organ occurred in the liver and kidneys, with an average time-integrated activity coefficient of 0.15 h (liver) and 0.14 h (kidneys). The average absorbed dose/dose coefficients across all subjects are shown in Table 3 (± SD). The highest absorbed dose per unit activity was observed in the kidneys (2.20E–01 mGy/MBq), followed by the urinary bladder wall (6.7E–02 mGy/MBq), spleen (6.3E–02 mGy/MBq), salivary glands (6.1E–02 mGy/MBq), and liver (4.3E–02 mGy/MBq).

The average effective doses reported individually for each patient with the tissue-weighting factors from ICRP publication 60~(27) was $1.99E-02\pm0.09E-02$ mSv/MBq. The SD of the effective dose was calculated by taking the mean of the individual patients' effective doses. The effective dose when the mean time-integrated activity coefficients were used resulted in a value of 1.99E-02 mSv/MBq (Table 3). Both values agreed well. The effective dose for an injection of 150 MBq of 68Ga-PSMA I&T was 3.0 mSv. The effective dose coefficient using ICRP 103~(29) tissue-weighting factors for the standard patient was 1.57E-02 mSv/MBq.

DISCUSSION

⁶⁸Ga-PSMA I&T shows a favorable dosimetry and biodistribution in prostate cancer patients. Its dosimetry compares favorably with that of other novel prostate cancer–specific imaging probes labeled with ⁶⁸Ga, ¹²³I, or ¹²⁴I (33–36) (Table 4). In addition, values for ⁶⁸Ga-labeled pentixafor (25), DOTATOC, and DOTATATE (34) and for ¹⁸F-FDG (35) are provided for comparison. Although the highest lesion-to-background ratio was observed 2 h after administration of the radiopharmaceutical, we recommend, for practical reasons

and because the lesion-to-background ratio is also acceptable, scanning the patients at 1 h after administration of 68 Ga-PSMA I&T.

Several theranostic PET probes of PSMA expression have been developed recently. The dosimetry of a ¹²⁴I- and ¹²³I-labeled small-molecule inhibitors of PSMA has been recently reported (MIP-1095, MIP-1072) (33,36). The kidney-absorbed dose coefficients for ⁶⁸Ga-PSMA I&T are higher than for other ⁶⁸Ga- and ¹²³I-labeled compounds but much lower than that of ¹²⁴I-MIP-1095. As for ¹²⁴I-MIP-1095, high physiologic uptake was observed in the salivary glands for ⁶⁸Ga-PSMA I&T. Both organs could be at risk and might need to be considered when developing ¹⁷⁷Lu-labeled PSMA I&T as a clinical theranostic. The effective dose is highest for the ¹²⁴I- and ¹²³I-labeled small-molecule inhibitors of PSMA for their respective administered activities (39, 12, and 9 mSv, Table 4).

Regarding methodologic considerations, the mean time-integrated activity coefficient is higher when using the blood-based method for bone marrow dosimetry as compared with the image-based method. Although this notion conflicts with the observations of Ferrer's report (23), we decided to assess the absorbed bone marrow dose conservatively using the higher values of the blood-based method. Overall, the absorbed doses to the bone marrow in our group of patients are of the order of several mGy for an administered activity of 150 MBq, thus showing that the bone marrow is not an organ at risk.

The urinary excretion does not rely on model assumptions. In fact, the current data were obtained after an observation period of at least 4 h after injection. At this time, 4%–7% of the injected activity was still retained in the whole-body. A comparison of the time-integrated activity coefficients of the bladder contents to the activity in the remainder of the body shows that, as a conservative estimate, less than 10% of the injected activity was excreted through the urinary tract. Thus, the absorbed dose to the bladder wall from 150 MBq of ⁶⁸Ga-PSMA I&T is significantly lower than that from an injection of 370 MBq of ¹⁸F-FDG (31).

As expected, ⁶⁸Ga-PSMA I&T was well tolerated by all patients. No acute or subacute adverse events were observed, and no significant changes in total blood count, kidney, or hepatic function occurred.

⁶⁸Ga-labeled PSMA-targeted radioligands (10) allow the in vivo visualization of PSMA expression. Because PSMA expression is associated with an adverse prognosis, this probe may provide important phenotypic in addition to the diagnostic information.

The high tumor-to-background ratios and the favorable biodistribution have resulted in rapid clinical adoption in several centers (11–17). In addition, initial therapeutic attempts with ¹³¹I-labeled PSMA ligands have been reported (33). However, ⁶⁸Ga-PSMA I&T allows for the labeling with ¹⁷⁷Lu (18), which is a potentially more desirable therapeutic compound (18).

In 4 of the 5 scans, pathologic tracer uptake consistent with malignant disease was observed. The 1 negative scan was found in a patient with biochemical relapse and a low PSA level of 0.5 ng/mL. Metabolic PET probes such as 11 C- or 18 F-choline and 11 C-acetate also failed to

detect sites of recurrence in patients with low serum PSA levels. The potential strength of ⁶⁸Ga-PSMA I&T is its ability to serve as a predictive biomarker for response to its ¹⁷⁷Lulabeled therapeutic analogs. This theranostic concept has been highly successful in the context of somatostatin receptor imaging and therapy (37).

There is an urgent clinical need to develop effective systemic treatments in advanced prostate cancer. ⁶⁸Ga-PSMA I&T will be explored as a theranostic that may also apply to other PSMA-expressing malignancies such as pancreatic cancer (38), breast cancer, and primary gliomas (39).

CONCLUSION

⁶⁸Ga-PSMA I&T exhibits promising dosimetry and is not associated with any toxicity. It shows favorable imaging characteristics with high lesion-to-background uptake ratios already 30 min after tracer injection. The best image contrast is achieved 1 h after intravenous injection. The kidney is the critical organ. Low tracer uptake in normal bone marrow may be of particular interest for future therapeutic applications. Organ-absorbed doses associated with ⁶⁸Ga-PSMA I&T are lower than those of many other PET radiopharmaceuticals with the exception of the kidneys.

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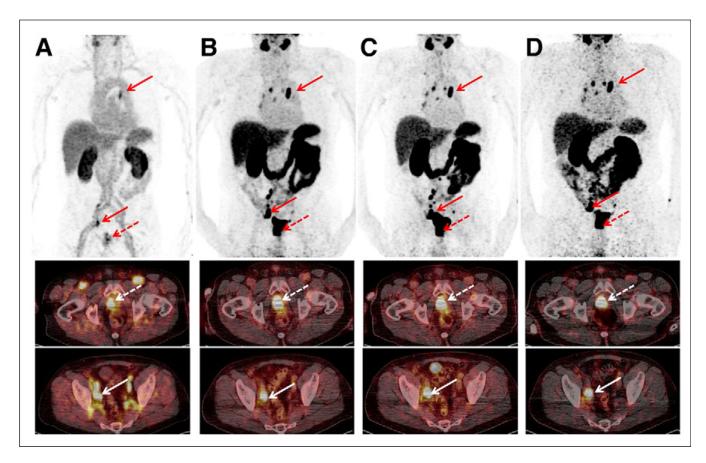


FIGURE 1.

Sequential scan of 70-y-old patient (P3; PSA level, 101.2 ng/mL) with initial diagnosis of prostate cancer (dotted arrow) showing high tumor-to-background ratio. Maximum-intensity projections (upper row) and axial slices (middle and lower rows) at different time points are displayed ([A] early rapid scan, [B] 1-h scan, [C] 2-h scan, [D] 4-h scan). Primary prostate cancer (middle row; 1-h SUV_{max}, 55.0) as well as numerous iliacal (lower row; 1-h SUV_{max}, 57.0) and mediastinal lymph node metastases (upper row; 1-h SUV_{max}, 31.4) can be depicted immediately after injection and up to 4 h.

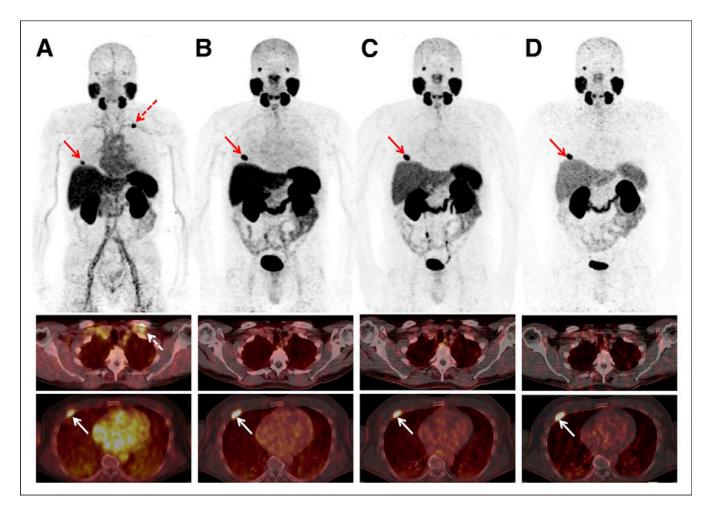


FIGURE 2.

Sequential patient scan (P1) of 67-y-old patient with biochemical relapse (PSA level, 6.7 ng/mL) 1.8 y after curative radiotherapy. Maximum-intensity projections (upper row) and axial slices (middle and lower row) of early rapid scan (A) and scans after 1 h (B), 2 h (C), and 4 h (D) show increased uptake in rib metastasis (A–D, arrows) and physiologic uptake in lacrimal glands, salivary glands, liver, spleen, kidneys, and slightly in bowel (A–D). Red dotted arrow in A depicts unspecific uptake in left subclavian vein.

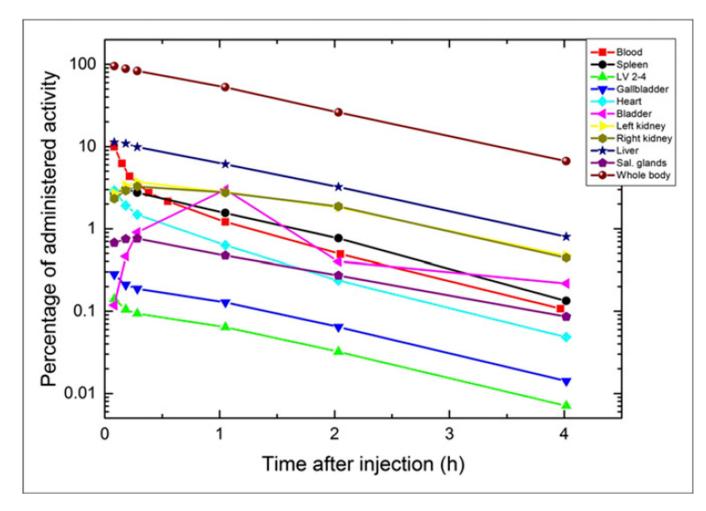


FIGURE 3.Time–activity curves for P1 for all organs showing uptake, for whole body, and for blood. For blood, percentage of activity is given per liter of blood.

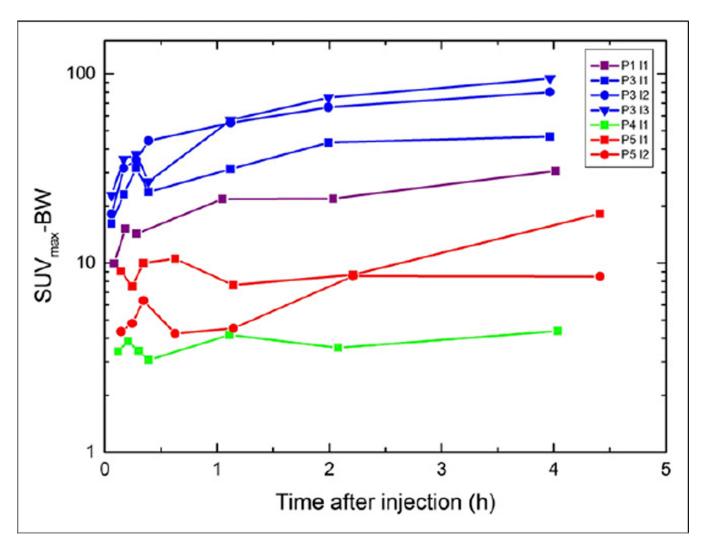


FIGURE 4.

Temporal variation of SUV_{max}-body weight in visible lesions in P1 (red), P3 (light blue), P4 (green), and P5 (blue). P1-l1 = bone lesion; P3-l1 = lymph node mediastinal; P3-l2 = tumor tissue prostate; P3-l3 = lymph node iliacal; P4-l4 = lymph node; P5-l1 = tumor tissue prostate; P5-l2 = bone lesion.

Herrmann et al.

TABLE 1

Patients' Demographic Data

PSA level Gleason Previous PET (ng/mL) Score treatment findings	6.7 7 RTx B	0.5 8 RPT None	101.2 7 — Loc, B, LNs	1.6 8 RPT and LNs RTx	1.4 8 ADT B, Loc
PET indication	Relapse	Relapse	Staging	Relapse	Follow- up
PD	148 09/12	124 01/13	146 07/14	133 07/11	91 07/14
Activity (MBq)	148	124	146	133	91
Weight (kg)	74	95	81	110	70
Height (cm)	170	178	172	185	177
Age (y)	8.99	66.3	6.69	59.1	63.5
Patient no.	1	2	3	4	5

PD = primary diagnosis; RTx = radiotherapy; B = bone metastasis; RPT = radical prostatectomy; Loc = tumor tissue in the prostate; LN = lymph node metastasis; ADT = androgen-deprivation therapy.

Page 15

Author Manuscript

TABLE 2

Time-Integrated Activity Coefficients

		Time-ii	ntegrate	Time-integrated activity coefficients (h)	7 coeffici	ents (h)	
Source organ	P1	P2	P3	P4	P5	Mean	SD
Remainder	1.023	1.029	0.952	1.098	0.964	1.013	0.059
Liver	0.181	0.141	0.100	0.142	0.173	0.147	0.032
Right kidney	0.080	0.068	0.075	0.078	0.038	0.068	0.018
Left kidney	0.084	0.076	0.087	0.073	0.042	0.073	0.018
Kidneys, sum	0.164	0.144	0.163	0.151	0.080	0.140	0.035
Heart	0.021	0.018	0.015	0.024	0.017	0.019	0.003
Bladder contents	0.042	0.029	0.055	0.039	0.088	0.051	0.023
Gallbladder	0.002	0.001	0.001	0.002	0.002	0.002	0.001
Spleen	0.042	0.010	0.011	0.019	0.032	0.023	0.014
Parotid glands	0.010		0.004		0.005	0.006	0.003
Submandibular glands	0.005	0.003	0.004	0.002	0.003	0.004	0.001
LV2-4	0.002	0.001	0.001	0.001	0.001	0.001	0.000
Red marrow lumbar vertebrae	0.027	0.013	0.009	0.018	0.016	0.017	0.007
Red marrow blood	0.028	0.026	0.019	0.024	0.022	0.024	0.003

Calculation methods for red marrow lumbar vertebrae and red marrow blood are presented in "Material and Methods" section.

P = patient; LV2-4 = lumbar vertebrae 2-4.

Herrmann et al.

 $\label{table 3}$ Absorbed Organ Dose Coefficients and Absorbed Organ Doses (150 MBq of $^{68}\text{Ga-PSMA})$

Page 17

Target organ	β (mGy/MBq)	Photon (mGy/MBq)	Total (mGy/MBq)	Absorbed dose (mGy) (150 MBq)
Adrenals	5.87E-03	7.62E-03	1.35E-02	2.0
Brain	5.87E-03	2.43E-03	8.29E-03	1.2
Breasts	5.87E-03	2.58E-03	8.44E-03	1.3
Gallbladder wall	1.16E-02	7.99E-03	1.96E-02	2.9
Lower large intestine wall	5.87E-03	4.68E-03	1.06E-02	1.6
Small intestine	5.87E-03	5.24E-03	1.11E-02	1.7
Stomach wall	5.87E-03	5.24E-03	1.11E-02	1.7
Upper large intestine wall	5.87E-03	5.25E-03	1.11E-02	1.7
Heart wall	1.48E-02	5.36E-03	2.02E-02	3.0
Kidneys	1.96E-01	2.31E-02	2.20E-01	33.0
Liver	3.30E-02	1.01E-02	4.31E-02	6.5
Lungs	5.87E-03	3.91E-03	9.78E-03	1.5
Muscle	5.87E-03	3.65E-03	9.52E-03	1.4
Ovaries	5.87E-03	4.94E-03	1.08E-02	1.6
Pancreas	5.87E-03	7.38E-03	1.32E-02	2.0
Red marrow	8.07E-03	4.35E-03	1.24E-02	1.9
Osteogenic cells	1.16E-02	4.04E-03	1.57E-02	2.4
Skin	5.87E-03	2.28E-03	8.15E-03	1.2
Spleen	5.26E-02	1.09E-02	6.34E-02	9.5
Testes	5.87E-03	3.28E-03	9.14E-03	1.4
Thymus	5.87E-03	3.62E-03	9.49E-03	1.4
Thyroid	5.87E-03	3.15E-03	9.01E-03	1.4
Urinary bladder wall	5.70E-02	1.03E-02	6.74E-02	10.1
Uterus	5.87E-03	5.72E-03	1.16E-02	1.7
Salivary glands	6.07E-02	_	6.07E-02	9.1
Total body	7.88E-03	3.85E-03	1.17E-02	1.8
Effective dose coefficient (mSv/MBq)			1.99E-02	
Effective dose coefficient (mSv/MBq)*			$1.93E-02 \pm 0.09E-02$	
Effective dose (mSv)				3.0

^{*} Mean effective dose coefficient of P1–P5.

Herrmann et al.

TABLE 4

Comparison of Absorbed Dose Coefficients and Absorbed Doses for Several Prostate-Specific Compounds

		124I-PSMA	¹²³ I-MIP- 1072	¹²³ I-MIP- 1095	Pentixafor	DOTATOC	DOTATOC DOTATATE	¹⁸ F-FDG	PSMA-IT
Target organ	Unit	Zechmann (33)	Barrett (36)	Barrett (36)	Herrmann (25)	Sandstrom (34)	Sandstrom (34)	ICRP 106 (35)	This work
Kidneys	mSv/ MBq	1.39E100	5.4E-02	1.10E-2	3.50E-02	8.20E-02	9.30E-02	1.70E-02	2.20E-01
Liver	mSv/ MBq	1.66E-00	2.4E-02	5.8E-2	1.75E-02	4.10E-02	5.00E-02	2.10E-02	4.31E-02
Spleen	mSv/ MBq	7.7E-01	2.3E-2	4.7E-2	5.38E-02	1.08E-01	1.09E-01	1.10E-02	6.34E-02
Urinary bladder wall	mSv/ MBq	5.7E-01	9.2E-2	2.1E-2	8.14E-02	1.19E-01	9.80E-02	1.30E-01	6.74E-02
Effective dose coefficient	mSv/ MBq	5.8E-01	2.5E-2	3.2E-2	1.56E-02	2.10E-02	2.10E-02	1.90E-02	1.99E-02
Typical injected activity	MBq	29	370	370	150	185	150	370	150
Effective dose	mSv	38.9	9.3	11.8	2.3	3.9	3.2	7.0	3.0

Page 18