## Investigation of <sup>18</sup>F and <sup>89</sup>Zr Isotopes Self-Absorption and Dose Rate Parameters for PET Imaging

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#### Abstract

This work concerns study of self-absorption factor (SAF) and dose rate constants of zirconium-89 ( $^{89}$ Zr) for the purpose of radiation protection in positron emission tomography (PET) and to compare them with those of <sup>18</sup>F-deoxyglucose ( $^{18}$ F-FDG). We analyzed the emitted energy spectra by <sup>18</sup>F and <sup>89</sup>Zr through anthropomorphic phantom and calculated the absorbed energy using Monte Carlo method. The dose rate constants for both radionuclides were estimated with 2 different fluence-to-effective dose conversion coefficients. Our estimated SAF value of 0.65 for <sup>18</sup>F agreed with the recommendation of the American Association of Physicists in Medicine (AAPM). The SAF for <sup>89</sup>Zr was in the range of 0.61-0.66 depending on the biodistribution. Using the fluence-to-effective dose conversion coefficients recommended jointly by the American National Standards Institute and the American Nuclear Society (ANSI/ANS), the dose rate at 1 m from the patient for <sup>18</sup>F was 0.143 µSv·MBq<sup>-1</sup>·hr<sup>-1</sup>, which is consistent with the AAPM recommendation, while that for <sup>89</sup>Zr was 0.154 µSv·MBq<sup>-1</sup>·hr<sup>-1</sup>. With the conversion coefficients currently recommended by the International Committee on Radiological Protection (ICRP), the dose rate estimates were lowered by 2.8% and 2.6% for <sup>89</sup>Zr and <sup>18</sup>F, respectively. Also, we observed that the AAPM derived dose is an overestimation near the patient, compared to our simulations, which can be explained by the biodistribution nature and the assumption of the point source. Thus, we proposed new radiation protection factors for <sup>89</sup>Zr radionuclide.

#### **Keywords**

fluorine-18, zirconium-89, dose rate constant, self-absorption factor, radiation protection

#### Introduction

Immuno-PET imaging uses monoclonal antibodies (mAbs) labeled with radionuclides for positron emission tomography. Unlike <sup>18</sup>F-FDG which is a relatively non-specific tracer, mAbs are designed to specific antigenic sites such as growth factor receptors. <sup>18</sup>F ( $t_{1/2} = 109$  min), <sup>64</sup>Cu (12.7 hr), <sup>68</sup> Ga (68 min), <sup>76</sup>Br (16 hr), <sup>86</sup>Y (14.7 hr), <sup>89</sup>Zr (78.4 hr) and <sup>124</sup>I (100.2 hr) have been used in immuno-PET.<sup>1</sup> Typical injected activity of <sup>18</sup>F-FDG ranges from 100 MBq to 500 MBq.<sup>289</sup>Zr gives good diagnostic quality at administered activities as low as 37 MBq<sup>2,3</sup> and is sufficiently long-lived ( $t_{1/2} = 78.4$  hr) to allow serial imaging up to approximately 1 week post-administration.<sup>4</sup> It can be produced in medical cyclotrons by proton irradiation of an <sup>89</sup>Y foil.<sup>5</sup> <sup>89</sup>Zr decays to <sup>89</sup>Y through  $\beta^+$  emission (22.7%) and electron capture (77.3%) followed by a prompt  $\gamma$ -ray at 909 keV (99.04%).<sup>6</sup>

In a Monte Carlo study, for radiation protection purposes, when using <sup>89</sup>Zr in an imaging facility that is optimized for the annihilation photons only, the results showed that the low

<sup>89</sup>Zr injected activity (75 MBq) would deliver a lower effective dose near the patient than <sup>18</sup>F (500 MBq) would; however, <sup>89</sup>Zr would give a higher effective dose than the <sup>18</sup>F outside the patient room due to the greater penetration power of the prompt γ-ray.<sup>6</sup>

In PET procedures utilizing <sup>18</sup>F-FDG, the 511 keV annihilation radiation is the main radiation protection concern, delivering a dose to the staff of 2.7-4.0 mSv during injection

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and 3.5-5.0 mSv during patient set-up.<sup>7</sup> The dose to the general public in the waiting area is about 0.8 mSv per procedure.<sup>2</sup> The dose to the staff and the general public depends on the positron emitter, the facility layout, the injected activity and the work load. The patient's physiology affects the accumulation of the tracer while his size affects the attenuation of the radiation. Thus, the dose rate D(x) in  $\mu$ Sv·hr<sup>-1</sup> at a distance *x*, in m, from the patient can be estimated by the following equation:

$$D(x) = A_0.DRC.SAF.x^{-2} \tag{1}$$

where  $A_0$  in MBq is the activity adjusted for biological and physical decay in the patient, DRC the dose rate constant in  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup> and SAF the dimensionless selfabsorption factor. The DRC is the constant of proportionality assuming that the dose rate is inversely proportional to the square of the distance from a point source. The SAF is defined as the ratio of the amount of energy absorbed by the patient's body to the amount of energy emitted by the radionuclide.<sup>6</sup> The American Association of Physicists in Medicine (AAPM) has recommended values of DRC and SAF for some commonly used radionuclides such as <sup>11</sup>C, <sup>18</sup>F, <sup>124</sup>I, for example, DRC =  $0.143 \ \mu \text{Sv} \cdot \text{m}^2 \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1}$  and SAF =  $0.64 \text{ for } ^{18}\text{F}$ .<sup>8</sup> The AAPM values for <sup>18</sup>F are further supported by studies in clinical setting<sup>2,9</sup> and with Monte Carlo methods<sup>10</sup> at distances beyond 1 m from the patient. However, studies near the patient (less than 1 m) concluded that the DRC provided by AAPM was a conservative value that over-estimated the dose.<sup>9</sup> The DRC for <sup>18</sup>F is not applicable to <sup>89</sup>Zr while the <sup>18</sup>F SAF may not be valid for <sup>89</sup>Zr. From previous work, we found that the estimated effective dose from  $^{89}$ Zr is 0.57  $\pm$  0.2 mSv/Mbq and the highest organ dose is  $1.87 \pm 0.2$  mSv/MBq in the liver.<sup>11</sup>

Furthermore, the AAPM recommendations are based on the ANSI/ANS fluence-to-dose conversion coefficients<sup>8</sup> that have been replaced by those in International Committee on Radiological Protection (ICRP) Publication 60<sup>12</sup> and subsequently in Publication 116.<sup>13</sup> The ANSI/ANS values are generally higher than the current ICRP recommendation. In particular, the difference between the 2 data sets is most prominent at energies below 0.1 MeV, which is relevant to the multiple scattered photons emerging from the patient.

The goal of this study was to investigate the SAF and the DRC for <sup>18</sup>F and <sup>89</sup>Zr using the Monte Carlo method and an anthropomorphic phantom.

#### Methods

We carried out the simulations to estimate the self-absorption factors and the dose rates in air using the Zubal voxel phantom<sup>14</sup> and MCNPX version 2.5.0.<sup>15</sup> For the benefit of computing efficiency, the simulations were done in photon-only mode. We did not model the positron source explicitly but implicitly with a combination of monoenergetic photon sources, each of which is either the annihilation photon (511 keV) or the prompt  $\gamma$ -ray (909 keV) emissions. Although secondary electrons were omitted, MCNPX accounted for their bremsstrahlung radiations with the thick-target bremsstrahlung model.<sup>6</sup> In each

Table 1. Typical Biodistributions of <sup>89</sup>Zr Used in This Study.

	<sup>89</sup> Zr-cmAb		<sup>89</sup> Zr-trastuzumab	
Organ	0 hr after injection [15]	3 days after injection [15]	4-5 days after injection [4]	
Bladder	-	-	-	
Brain	-	-	0.232	
Kidneys	0.008	0.003	0.399	
Ĺiver	0.105	0.291	0.184	
Lungs	0.036	0.085	-	
Skeleton	-	-	0.029	
Spleen	0.007	0.139	0.028	
, Total body	0.845	0.453	0.128	

simulation, a monoenergetic photon source was distributed uniformly in 1 of 7 organs or to the entire body—the "total body" distribution. The results for <sup>18</sup>F were derived from the 511 keV simulations assuming 1.93 annihilation photons per decay; those for <sup>89</sup>Zr were derived from the 511 keV and the 909 keV simulations assuming 0.46 annihilation photons and 0.99 prompt  $\gamma$ -ray per decay, respectively. The photon intensities per decay came from the Nuclide Chart by National Nuclear Data Center of Brookhaven National Laboratory (www.nndc.bnl.gov/chart). Thus, the results for a radiopharmaceutical were weighted sums of the tallies according to the emission intensities and the biological distribution of the radiopharmaceutical. Statistical error doesn't exceed 1% for all simulations carried out, with a total number of events of 10<sup>9</sup> for each run.

#### **Biological Distributions**

In this study, we assumed the following typical biodistribution for <sup>18</sup>F-FDG (where the percentages refer to the percent of the administered activity): 7.5% in the brain, 4.8% in the liver, 6.8% in the bladder content and 80.9% in the rest of the body.<sup>10</sup> For <sup>89</sup>Zr, we studied the distributions of 2 radiopharmaceuticals from literature <sup>89</sup>Zr labeled chimeric monoclonal antibody U36 (cmAb) for head and neck cancer patients<sup>4</sup> and trastuzumab for breast cancer patients.<sup>3</sup> The 2 studies gave uptakes in individual organs (Table 1). The activities in the rest of the body was derived by subtracting the sum of the organ activities at each time point from the total body activity decayed from the previous time point.<sup>11</sup> The uptake ratio for each organ was reassigned to account for of the total body distribution. For <sup>89</sup>Zr-cmAb, we can see that the radioactivity clearly concentrated in liver and spleen sites over time. However, kidneys, brain and liver organs presented the highest uptake after 4-5 days for <sup>89</sup>Zr-trastuzumab. Each organ has its own specific labeling response capability.

#### Self-Absorption Factors and Emission Spectra

For the self-absorption simulations, the phantoms were positioned at the center of a vacuum sphere of 100 cm radius. The total amount of photon energy crossing the spherical surface was calculated with the \*F1 tally, which is a track length estimator modified by the particle energy so that the energy is tallied instead of the number of photons. To obtain the total amount of energy emitted by a radionuclide in multiple organs, the \*F1 tallies were weighted by the biological distribution and summed together. Thus the self-absorption factor of a radionuclide was the weighted sum divided by the total amount of energy that it emitted.

#### Dose Rate Constants

Dose rates at 1, 4 and 8 m from the phantom center were estimated in concentric spherical shells at those positions. Outside of the phantom was modeled with air medium. The F4 track length estimators modified with fluence-to-effective dose conversion coefficients were applied in the spherical shells. The unit of the modified tallies was Sv/hr per source photon. Similar to the self-absorption and spectrum calculations, the estimation of the dose rate D at distance x took into account the photon emission intensity and biological distribution of the radiopharmaceutical.

The phantom scattered dose rate constant K that accounts for phantom scattering and self-absorption was obtained by least square fit of the estimated dose rates D(x) per MBq of activity in the phantom at x by the equation

$$D(x) = Kx^{-2} \tag{2}$$

Ideally, K should be close to the product of DRC and SAF.

#### **Results and Discussion**

# Dose Rate Constants From a Point Source, ANSI/ANS vs ICRP

The dose rate at 1 m from point source of <sup>18</sup>F was estimated to be 0.143  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> using the ANSI/ANS conversion coefficients. Our estimated dose rate for <sup>18</sup>F was the same as the AAPM recommended value. Since the AAPM model used a point source, the value of the dose rate at 1 m is the value for the DRC, that is, the DRC for <sup>18</sup>F is 0.143  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup>. When the current ICRP conversion coefficients were used instead of the ANSI/ANS coefficients, the DRC was reduced to 0.139  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup>, which was 2.8% below the AAPM value. For <sup>89</sup>Zr, the DRCs were 0.154  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup> and 0.150  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup> using ANSI/ANS and the ICRP conversion coefficients, respectively. The ICRP conversion coefficients reduces the dose by 2.6%.

#### Self-Absorption Factor

The estimated SAFs in this study are listed in Table 2. The SAF calculated with a typical biodistribution of <sup>18</sup>F-FDG in the Zubal phantom was 0.65. Our value is between the AAPM value of 0.64<sup>8</sup> and the value of 0.66 when using the phantom developed by the Committee on Medical Internal Radiation

**Table 2.** Self-Absorption Factors of <sup>18</sup>F and <sup>89</sup>Zr in Various Biodistributions for Zubal Phantom.

Biodistribution	<sup>18</sup> F	<sup>89</sup> Zr
<sup>18</sup> F-FDG	0.65	-
Uniform total body	0.66	0.67
<sup>89</sup> Zr-cmAb, 0 hr	-	0.66
<sup>89</sup> Zr-cmAb, 72 hr	-	0.62
<sup>89</sup> Zr-trastuzumab, 4-5 days	-	0.61

**Table 3.** Phantom Scattered Dose Rate Constants (K,  $\mu$ Sv·m<sup>2</sup>·MBq<sup>-1</sup>·hr<sup>-1</sup>) of <sup>18</sup>F and <sup>89</sup>Zr for Various Biodistribution in the Zubal Phantom.<sup>a</sup>

Biodistribution	<sup>18</sup> F	<sup>89</sup> Zr
<sup>18</sup> F-FDG	0.085	-
Uniform total body	0.086	0.094
<sup>89</sup> Zr-cmAb, 0 hr	-	0.092
<sup>89</sup> Zr-cmAb, 72 hr	-	0.086
<sup>89</sup> Zr-trastuzumab, 4-5 days	-	0.089

<sup>a</sup>The R<sup>2</sup> values are at least 0.9998.

Dose (MIRD).<sup>10</sup> Since the fraction of administered activity of <sup>18</sup>F-FDG in total body is 0.849, the SAF is similar to the one from a uniform distribution in total body 0.66.

At the time of injection, the fraction of <sup>89</sup>Zr-cmAb in total body was 0.845; the SAF was estimated to be 0.66 which was similar to the one from total body, 0.67. In fact, the SAF was also close to the one of <sup>18</sup>F-FDG. After 72 hours, the SAF reduced to 0.62 because the fraction of the tracer in total body was reduced to 0.453. In the case of <sup>89</sup>Zr-trastuzumab, its ratio in total body was further reduced to 0.128 after 4-5 days. The SAF was estimated to be 0.61 which is similar to the SAF of <sup>89</sup>Zr-cmAb after 3 days. Hence our study suggested that the SAF for <sup>89</sup>Zr was about 0.66 immediately after injection and 0.61 a few days later.

## Phantom Scattered Dose Rate Constants of the Emissions From the Zubal Phantom

The phantom scattered dose rates were calculated at 1, 2, 4 and 8 m from the phantom center and normalized by the activity in the phantom. They had not been adjusted for the physical and biological decay. These data were fitted to equation (2) to obtain K. Furthermore, the ANSI/ANS conversion coefficients were applied instead of the current ICRP recommendation to maintain comparability with AAPM recommendations. Table 3 summarizes the values of K from various biodistribution. For comparison, the DRC and SAF product under the AAPM formalism gives  $0.094 \ \mu \text{Sv} \cdot \text{m}^2 \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1}$  for <sup>18</sup>F and it varies between  $0.094 \ \mu \text{Sv} \cdot \text{m}^2 \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1}$  and  $0.103 \ \mu \text{Sv} \cdot \text{m}^2 \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1}$  for <sup>89</sup>Zr. Thus, the AAPM formalism yields conservative estimates near the patient.

As pointed out by Zaidi and Xu,<sup>16</sup> the voxel phantoms allow the modeling of any geometrical shape of the body and any complex distribution of radioactive sources inside the body. Since the phantoms are developed from scans of real patients or volunteers, the modeled distribution and the scattering of the radiation will vary according to the phantom employed in the simulation. In general, larger phantom will attenuate the emitted radiation more than a smaller one. The Zubal phantom was developed to reflect an average adult male<sup>14</sup>; its size is similar to the MIRD phantom used by AAPM. Although our estimated SAF for <sup>18</sup>F was the same as the AAPM recommended value, the distributed source in our simulation yielded a smaller value in the phantom scattered dose rate constant than the AAPM suggested DRC-SAF product.

### Comparing <sup>18</sup>F Dose With AAPM Values

A point source is a crude approximation of the radioactivity distributed in a patient. Using the Zubal phantom and a typical biological distribution of <sup>18</sup>F, the dose rate at 1 m was 0.087  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> anterior to the phantom, which is 40% below the estimation with the AAPM model: 0.143  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> × 0.64 / 1 m<sup>2</sup> = 0.092  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>. The discrepancy between the 2 models decreases with increasing distance; it reduces to 4% at 2 m, below 1% at 4 m and beyond. Thus, the dose to the nurse or technician who administer the radiopharmaceutical injection may be overestimated based on the AAPM dose rate constant.

Dose survey suggested that the dose rates at 30 cm. 1 m and 2 m from the patient were 290  $\mu$ Sv·hr<sup>-1</sup>, 60  $\mu$ Sv·hr<sup>-1</sup> and 15 µSv·hr<sup>-1</sup>, respectively, during injection of 505 MBq of <sup>18</sup>F-FDG.<sup>2</sup> During the scanning after 1 hr of uptake, the dose rates were 190  $\mu$ Sv·hr<sup>-1</sup>, 47  $\mu$ Sv·hr<sup>-1</sup> and 20  $\mu$ Sv·hr<sup>-1</sup> at 30 cm, 1 m and 1.3 m, respectively. Estimation using our results gave 332  $\mu$ Sv·hr<sup>-1</sup> (30 cm), 44  $\mu$ Sv·hr<sup>-1</sup> (1 m) and  $12 \ \mu \text{Sv} \cdot \text{hr}^{-1}$  (2 m) during injection and 227  $\mu \text{Sv} \cdot \text{hr}^{-1}$  (30 cm),  $30 \ \mu \text{Sv} \cdot \text{hr}^{-1}$  (1 m) and  $19 \ \mu \text{Sv} \cdot \text{hr}^{-1}$  (1.3 m) during scanning. On the other hand, the doses derived from AAPM would be 514  $\mu Sv \cdot hr^{-1}$  (30 cm), 46  $\mu Sv \cdot hr^{-1}$  (1 m) and 12  $\mu Sv \cdot hr^{-1}$ (2 m) during injection and 352  $\mu$ Sv·hr<sup>-1</sup> (30 cm), 32  $\mu$ Sv·hr<sup>-1</sup> (1 m) and 19  $\mu$ Sv·hr<sup>-1</sup> (1.3 m) during scanning. Thus, the simulation results were broadly comparable to the measurements. Furthermore, the results were very close to the AAPM derived dose at 1 m and beyond.

The observation that the AAPM derived dose is an overestimation near the patient is also supported by Quinn et al.<sup>9</sup> The authors measured 0.37  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> at 30 cm from the patient during scanning. Our simulations gave a value of 0.42  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> while the AAPM values gave a value of 1.02  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>. Quinn et al suggested that the biodistribution might explain the discrepancy between their measured value and the one calculated using the AAPM parameters.

Our simulations showed that the highest dose rates occurred if the radionuclide concentrated in the brain  $(0.90 \,\mu \text{Sv} \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1} \text{ at } 0.3 \,\text{m})$ , the skin  $(0.78 \,\mu \text{Sv} \cdot \text{MBq}^{-1} \cdot \text{hr}^{-1} \text{ at } 0.3 \,\text{m})$ .

at 0.3 m) and in the muscle (0.64  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> at 0.3 m). If the radionuclide was distributed uniformly over the entire body, the dose rate at 0.3 m would be 0.66  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>. When the radionuclide were distributed in deeper tissues, the dose rate predictable would be less than the foregoing values. All values were well below the AAPM dose rate, 1.02  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>. Our estimated dose rate of 0.42  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup> from a typical biodistribution was a weighted sum of the dose rates from individual organs. It is compatible with measurements interference.<sup>8</sup> While the biodistribution is an important factor in the dose rate close to the patient, the AAPM assumption of a point source is the main cause of the discrepancy.

### Comparing <sup>89</sup>Zr With <sup>18</sup>F Doses

The dose rate estimated at 1 m from a point source of <sup>89</sup>Zr in our simulation was 0.154  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>, which was 8% above the <sup>18</sup>F dose rate. With the radionuclide distributed in the Zubal phantom, we estimated that the patient body absorbed 33%-38% of the emitted energy depending on the distribution. Therefore, the self-absorption factor was estimated to be 0.67 for a uniform total body distribution, 0.66 for a typical U36 distribution and 0.61 for the trastuzumab distribution. In contrast, we estimated the self-absorption factor for <sup>18</sup>F was 0.65 while AAPM estimated it to be 0.66. The differences between <sup>18</sup>F and <sup>89</sup>Zr values were generally small because the curve of mass energy absorption coefficients for soft tissue is relatively flat in the range of 100 keV to 1 MeV. Using log-log interpolation of the NIST mass energy absorption coefficient data (https://www.nist.gov/pml/x-ray-mass-a ttenuation-coefficients), the values are  $3.267 \times 10^{-2}$  cm<sup>2</sup>/g and  $3.117 \times 10^{-2}$  cm<sup>2</sup>/g for 511 keV and 909 keV photons, respectively. The absorption of the prompt  $\gamma$ -ray in the body is only about 5% less than that of the annihilation photon.

We calculated the <sup>89</sup>Zr dose rates for 3 biodistributions: (1) a hypothetical situation same as the typical <sup>18</sup>F-FDG distribution by Elschot et al,<sup>10</sup> (2) in breast cancer patients by Dijkers et al,<sup>4</sup> and (3) in head and neck cancer patients by Alfuraih et al.<sup>11</sup> We repeated the <sup>18</sup>F calculations assuming that it followed the <sup>89</sup>Zr biodistributions (2) and (3) for comparison. The values are summarized in Table 4.

Due to the presence of the 909 keV prompt  $\gamma$ -ray, the dose rates from <sup>89</sup>Zr labeled antibodies are higher than the <sup>18</sup>F-FDG in organ-by-organ comparisons. It is interesting to note that the dose rate from a typical <sup>18</sup>F-FDG distribution in the body was very similar to a uniformly distributed <sup>18</sup>F source in the entire body (0.087  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>). Although the <sup>89</sup>Zr radiopharmaceutical accumulated more in the tumor deeper in the body, the estimated dose rates were about 3% (cmAb, 0.093  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>) to 5% (trastuzumab, 0.091  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>) below a that of a uniform total body distribution (0.096  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>).

Despite the higher dose rate of <sup>89</sup>Zr due to the greater penetration power of the prompt  $\gamma$ -ray, a typical immuno-PET scan using <sup>89</sup>Zr may not give a higher dose to the clinical staff than

	<sup>18</sup> F Dose Rates at 1 m Anterior to the
Phantom [µSv·MBq <sup>-1</sup> ·hr <sup>-1</sup> ].ª	

Distribution	<sup>89</sup> Zr	<sup>18</sup> F	
Uniform distribution in selected organ	s		
Bladder	0.073	0.065	
Brain	0.131	0.112	
Kidneys	0.074	0.066	
Liver	0.079	0.070	
Skin	0.112	0.103	
Total Body	0.096	0.087	
Typical radiopharmaceutical distributions in patients			
<sup>18</sup> F-FDG	0.096	0.087	
U36 in head and neck cancer	0.093	0.085	
trastuzumab in breast cancer	0.091	0.082	

<sup>a</sup>Notice that the <sup>18</sup>F would not follow the U36 or the trastuzumab distributions in reality and neither would <sup>89</sup>Zr follow the FDG one. They are included for comparisons only.

**Table 5.** Comparing <sup>89</sup>Zr and <sup>18</sup>F Dose Rates Per Scan at 1 m Anterior to the Phantom  $[\mu Sv \cdot hr^{-1}]^{a}$ .

Distribution	<sup>89</sup> Zr	<sup>18</sup> F
At the time of injection		
Total body	7.2	44
After I hr		
<sup>18</sup> F-FDG	-	30
cmAb in head and neck cancer	6.9	-
trastuzumab in breast cancer	6.8	-

 $^{a}$ lt was assumed that the injected activity of  $^{89}\text{Zr}$  was 75 MBq while that of the  $^{18}\text{F}$  was 500 MBq.

<sup>18</sup>F-FDG because immuno-PET generally utilizes a lower administered activity (75 MBq) than that typically used for <sup>18</sup>F-FDG (~500 MBq). Table 5 compares the dose rates of the 2 radionuclides during injection and during scanning. We assumed that the radiopharmaceuticals followed a uniform total body distribution at the time of injection and subsequently accumulated in a typical distribution after 1 hour of uptake time. Because of the long half-life of <sup>89</sup>Zr, the dose rates changed very little from 7.2 μSv·hr<sup>-1</sup> during injection to 6.8-6.9 μSv·hr<sup>-1</sup> during scanning. Furthermore, the low injected activity made the <sup>89</sup>Zr dose rates *per scan* significantly lower than those of a typical <sup>18</sup>F-FDG study that gave 44 μSv·hr<sup>-1</sup> during injection and 28 μSv·hr<sup>-1</sup> during scanning.

#### ICRP Conversion Coefficients Applied to the Emission From the Zubal Phantom

The dose rate at 1 m from a point source of <sup>18</sup>F and <sup>89</sup>Zr decreased to 0.139  $\mu$ Sv/hr and 0.150  $\mu$ Sv/hr, respectively, when the current ICRP conversion coefficients were employed. They were 2.8% and 2.6% below the values obtained with the ANSI/ANS coefficients. When the estimation was estimated again with the Zubal phantom, the estimations using the ICRP coefficients were also smaller than those using the ANSI/ANS

ones. However, it is interesting to note that the reduction was larger in the case of <sup>18</sup>F (4.5%) but smaller than in the case of <sup>89</sup>Zr (1.4%) because of the greater penetration power of the <sup>89</sup>Zr emission. The <sup>89</sup>Zr emissions from the distal parts of the body were less attenuated than those of <sup>18</sup>F when they arrived at the point of calculation.

#### Conclusion

We have presented our study on the <sup>18</sup>F and <sup>89</sup>Zr dose rates and the dose rate constants from point source in air and also from an anthropomorphic phantom. The dose rate at 1 m for <sup>18</sup>F was found to be 0.143  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>, which was consistent with the AAPM recommendation, while that for <sup>89</sup>Zr was found to be 0.154  $\mu$ Sv·MBq<sup>-1</sup>·hr<sup>-1</sup>. We estimated SAF for <sup>18</sup>F to be 0.65, which also agreed with the AAPM recommendation. The SAF for <sup>89</sup>Zr was in the range of 0.61–0.66 depending on the biodistribution. We further examined the effect of fluence-toeffective dose conversion coefficients on the effective dose rate estimations. Also, we found that the AAPM derived dose is an overestimation near the patient, compared to our simulations, which can be explained by the biodistribution nature and the assumption of the point source. Compared to using the current ICRP coefficients, the AAPM method using the ANSI/ANS coefficients overestimated the dose rate by about 3% for both radionuclides.

#### **Declaration of Conflicting Interests**

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