

Live mouse imaging with ^{44}mSc by a multiple-isotope PET

T. Fukuchi,^{*1} M. Shigeta,^{*1} D. Mori,^{*2} T. Yokokita,^{*2} Y. Komori,^{*2} H. Haba,^{*2} and Y. Watanabe^{*1}

We have been developing a multiple-isotope positron emission tomography (MI-PET) system that can analyze the dynamics of multiple tracers. Using a positron- γ emitter, which emits de-excitation γ -ray as a tracer after the positron emission in β -decay, the MI-PET system identifies the tracer by detecting the prompt γ -ray emitted after the positron emission. Figure 1 shows a schematic illustration of the developed MI-PET prototype system. This system is composed of a PET scanner and additional γ -ray detectors.¹⁾ In this system, in addition to conventional PET imaging, coincidence among the additional detectors and the PET scanner can be performed. We expect that MI-PET will be used for drug discovery research by direct comparison between old and new drugs.

For multiple-isotope imaging using MI-PET, at least one positron- γ emitter is necessary as a tracer. Scandium-44 is one of the promising radioactive-tracer candidates for MI-PET because of its large positron and γ -ray emission ratio and moderate half-life (^{44}Sc : 3.97 h, $^{44\text{m}}\text{Sc}$: 58.61 h). In our previous work, we performed dual-isotope phantom imaging using $^{44\text{m}}\text{Sc}$ and ^{18}F (pure positron emitter) and evaluated the basic imaging performance of MI-PET for $^{44\text{m}}\text{Sc}$.²⁾

Therefore, before the future development of a $^{44\text{m}}\text{Sc}$ -labeled drug for MI-PET applications, in order to test the practical imaging ability of the MI-PET system for $^{44\text{m}}\text{Sc}$, we conducted dual-isotope live mouse imaging using $^{44\text{m}}\text{Sc}$, which is a simple substance.

Scandium-44m was produced at the RIKEN AVF cyclotron via the reaction $^{44}\text{Ca}(d, 2n)^{44\text{m}}\text{Sc}$ with a 24-MeV deuterium beam. As the irradiation target, ^{44}CaO

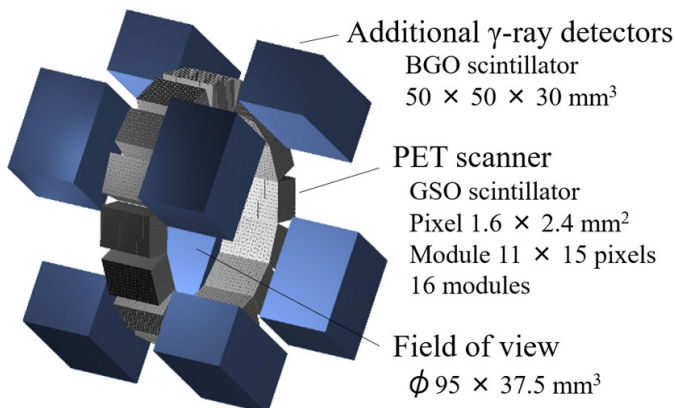


Fig. 1. Schematic illustration of the developed MI-PET prototype system.

^{*1} RIKEN Center for Biosystems Dynamics Research

^{*2} RIKEN Nishina Center

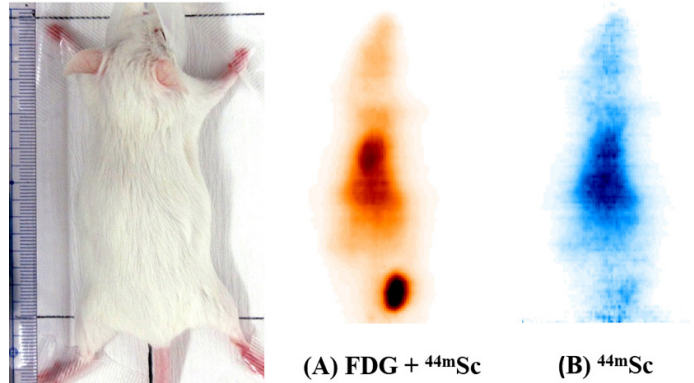


Fig. 2. Images of a live mouse with $^{44\text{m}}\text{Sc}$ and ^{18}F -FDG injections (maximum intensity projection images). The reconstructed images were acquired with the absence (A) or presence (B) of γ -ray coincidence.

(97.0% enriched ^{44}Ca isotope) powder was pressed into a disk of 10-mm diameter. The irradiated target was dissolved in 6 M HCl and purified by the chemical processes.²⁾ Finally, approximately 2 MBq of pure $^{44\text{m}}\text{Sc}$ was produced and transported to RIKEN Kobe campus for the imaging experiment. Most of the short-lived by-product, ^{44}Sc , decayed out during transportation.

In the mouse imaging experiment, 197 kBq $^{44\text{m}}\text{Sc}$ (simple substance) and 198 kBq ^{18}F -FDG were administered to an 8-week-old normal male mouse by tail-vein injection. Five minutes after administration, a 30-min whole-body scan was performed under anesthesia.

The result of dual-isotope mouse imaging is shown in Fig. 2. From the reconstructed images with the absence (A) or presence (B) of the prompt γ -ray detection, image (A) reflects the distribution of both ^{18}F -FDG and the $^{44\text{m}}\text{Sc}$ tracer, whereas image (B) reflects the isolated image of the $^{44\text{m}}\text{Sc}$ tracer. In these images, we can clearly observe the difference between ^{18}F -FDG and $^{44\text{m}}\text{Sc}$ distributions, *i.e.*, ^{18}F -FDG is distributed in the heart and urinary bladder, whereas $^{44\text{m}}\text{Sc}$ distributed in the liver.

From this experiment, we successfully demonstrated the practical feasibility of dual-isotope imaging using MI-PET with $^{44\text{m}}\text{Sc}$ as the second tracer. In the future, we will synthesize a useful MI-PET drug labeled by $^{44\text{m}}\text{Sc}$ and perform multiple-drug imaging on disease model animals.

References

- 1) T. Fukuchi *et al.*, *Med. Phys.* **44**, 2257 (2017).
- 2) T. Fukuchi *et al.*, *RIKEN Accel. Prog. Rep.* **52**, 207 (2019).