

## DIGITAL AUTORADIOGRAPHY OF BONE-SEEKING RADIONUCLIDES IN HUMAN

George Tabatadze\*, Brian W. Miller†, Sergei Y. Tolmachev\*

\*U.S. Transuranium and Uranium Registries, 1845 Terminal Drive, Richland, WA 99354, USA,

[george.tabatadze@wsu.edu](mailto:george.tabatadze@wsu.edu), [stolmachev@wsu.edu](mailto:stolmachev@wsu.edu)

†College of Optical Sciences, University of Arizona, Tucson, AZ 85719, USA, [bwmiller@optics.arizona.edu](mailto:bwmiller@optics.arizona.edu)

*This paper describes the ionizing-radiation quantum imaging detector (iQID) system and its applicability for imaging of bone-seeking alpha-emitters. The United States Transuranium and Uranium Registries (USTUR) studies actinide (plutonium, americium, and uranium) biokinetics and tissue dosimetry by following up occupationally exposed individuals. Estimation of the micro-distribution of radionuclides in tissues is an important task to support biokinetic modeling and dose assessment. A newly developed iQID system was used to study radionuclide distribution in human bones. Results showed that iQID imaging approach is proven to be an effective method for micro-scale heterogeneous distribution studies, where traditional counting methods do not apply.*

### IONIZING-RADIATION QUANTUM IMAGING DETECTOR

The ionizing-radiation Quantum Imaging Detector (iQID) is a newly developed digital autoradiography (radiation imaging) system<sup>1</sup>. The iQID allows for real-time quantitative autoradiography and study of radionuclide micro-distribution at a low radionuclide activity level ( $<10^{-3}$  events per second). The iQID system is a portable, laptop-operated unit. Single-particle imaging with sub-pixel position estimation enables imaging studies to be performed at spatial resolutions as high as 20  $\mu\text{m}$ . Large-area iQID configurations (up to 200 mm diameter) accommodate studies requiring simultaneous imaging of an array of samples. The high detection efficiency (50-100%), low background rate, and event-by-event spatiotemporal information allow activity distributions to be quantified, even with short-lived radionuclides.

The iQID is comprised of a scintillator in direct contact with a micro-channel plate (MCP) image intensifier and a lens for imaging the intensifier screen to a CCD or CMOS camera sensor, all within a compact light-tight enclosure. iQID is sensitive to a broad range of radiation including gamma-/X-rays, neutrons, spontaneous fission, conversion electrons, alpha and beta particles<sup>1</sup>.

In order to localize the origin of a radioactive particle precisely, a iQID image is superimposed over the structural image of a sample. The iQID image carries information on the spatial distribution of radioactive particles, while a structural image represents a sample geometry. The structural digital image is acquired using a scanner, digital camera, or microscope.

### FEASIBILITY STUDIES

The iQID system was used for microdosimetry of targeted radionuclide therapy using  $\alpha$ - and  $\beta$ -emitters:  $^{211}\text{At}$ ,  $^{90}\text{Y}$ , and  $^{177}\text{Lu}$  in soft tissues<sup>2</sup>. At the USTUR, the application of iQID is successfully extended for imaging of bone-seeking  $\alpha$ -emitters:  $^{241}\text{Am}$ ,  $^{239}\text{Pu}$ , and  $^{226}\text{Ra}$  in humans<sup>3</sup>. For the internally deposited radionuclides, activity distribution was visualized and quantified in various bone sections. Radionuclide activity distribution ranged between 0.002 and 0.003  $\text{mBq mm}^{-2}$  for  $^{239}\text{Pu}$  0.1 and 0.7  $\text{mBq mm}^{-2}$  for  $^{226}\text{Ra}$ , and 1.0 and 10.0  $\text{mBq mm}^{-2}$  for  $^{241}\text{Am}$ . Mapping of radionuclide distribution was successfully achieved on a macro-scale. However, it was challenging to distinguish whether  $\alpha$ -events originated from the surface or volume of a sample. The  $\alpha$ -interference can be eliminated by preparing micron-thick slides. (Fig. 1).

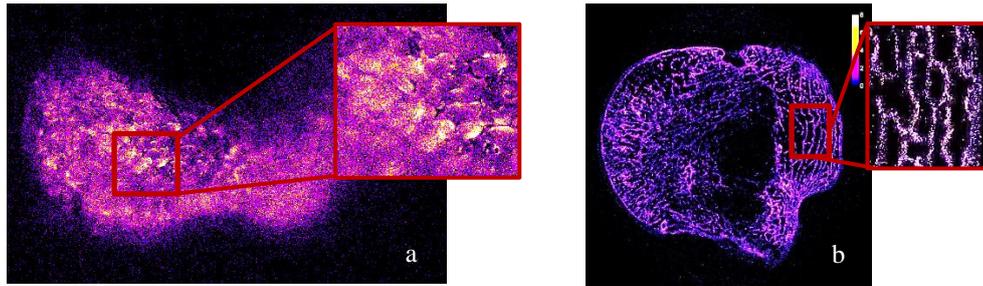


Fig.1. Distribution of  $^{241}\text{Am}$  in clavicle acromial end (a) – unpolished bone surface, and in humerus proximal end (b) – polished 100- $\mu\text{m}$ -thick slide.

## BONE MICRO-DOSIMETRY

To study bone micro-dosimetry, bone specimens were sampled from humerus proximal end, humerus proximal shaft, and clavicle acromial end. These specimens were embedded in methyl methacrylate plastic and processed to produce multiple 100  $\mu\text{m}$ -thick sections. Bone sections were polished to a fine surface.<sup>4</sup> This allowed to investigate distribution of metabolized  $^{241}\text{Am}$  within trabecular bone regions on a micro-scale. The  $^{241}\text{Am}$  activity distributions were visualized and quantified in cortical bone and trabecular spongiosa (Fig. 1b). The  $^{241}\text{Am}$  activity concentration ratios within different bone regions were used to represent the radionuclide distribution. The trabecular-to-cortical bone and trabecular spongiosa-to-cortical bone ratios are reported in Table I for the humerus and clavicle. The iQID values are in agreement with those obtained from radiochemical analysis but not consistent with the ICRP biokinetic model predictions.

TABLE I.  $^{241}\text{Am}$  activity concentration ratios within different bone regions

Bone Region	Humerus Bone		Clavicle Bone		iQID/ICRP	
	iQID	$\alpha$ -spec	iQID	$\alpha$ -spec	Humerus	Clavicle
Trabecular-to-cortical	$2.76 \pm 0.04$	$2.15 \pm 0.13$	$1.29 \pm 0.15$	$1.48 \pm 0.11$	5.5	2.6
Spongiosa-to-cortical	$1.09 \pm 0.01$	$1.28 \pm 0.08$	$0.65 \pm 0.06$	$0.68 \pm 0.05$	5.5	3.3

## CONCLUSIONS

The iQID digital imager allows for real-time visualization and quantitative digital autoradiography of bone-seeking alpha-emitters. To reduce a signal-to-noise ratio and improve an image resolution, appropriate sample preparation is required. The  $^{241}\text{Am}$  micro-distribution measurements showed that ICRP defaults underestimate  $^{241}\text{Am}$  concentration ratios within cortical bone regions at least by a factor of 3.

## ACKNOWLEDGMENTS

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