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Influence of the vials radioactive residue in Nuclear Medicine procedure applied to a new "in situ" activimeter calibration methodology



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ABSTRACT

In this study we present a new "in situ" calibration methodology establishment for activimeters – i.e., this equipment calibration can be made at the hospital and/or the nuclear medicine service (NMS) – and also the importance of considering vials radioactive residue when generating the calibration coefficient in terms of activity (N_A). For the calibration, two methods are presented here: in the first, 99m Tc sample activity (A_C) is measured in the NMS. Then it is sent to the Instruments Calibration Laboratory (LCI/IPEN) where the reference activity (A_R) is obtained under controlled conditions in a reference vial. After geometry and residue corrections, N_A is calculated. In the second method, A_R is obtained at the LCI, then 99m Tc sample is sent to the hospital/NMS, where A_C is measured and N_A calculated. Calibration methodologies were applied to three NMS and to four equipment belonging to the radiopharmaceutical production facility (CR) at the Nuclear and Energy Research Institute (IPEN). Both methodologies were tested with and without residual activity corrections under controlled conditions. Variations of up to 4% were obtained.

1. Introduction

In the last decades, numbers of radiological and nuclear medicine (NM) procedures in the United States have increased massively, from 54 in 1964 to about 500 in 2006 (Mettler et al., 2009). Therefore, radiation dose to population has also increased. In this same year nuclear medicine services (NMS) represented only 5% of total medical procedures, but were responsible for 26% of medical collective dose in the US (Mettler et al., 2009; Fahey et al., 2011).

Both workers and patients security assurance in NMS has been the focus of international organizations (International Atomic Energy Agency, IAEA, 2006; International Atomic Energy Agency, IAEA, 2010). A safe and efficient NMS performance depends, among other factors, on a complete quality assurance program. This includes, mainly, the activimeter quality control, which must meet quality standards provided by international authorities (American Association of Physicists in Medicine, AAPM, 2012; International Atomic Energy Agency, IAEA, 2015). Quality assurance for an activimeter includes tests such as high voltage verification, zero adjust, background radiation, accuracy, repeatability, etc., that must be performed periodically with a specific radioactive source according to international recommendations (National Physical Laboratory, 2006).

Some parameters, such as vials geometry and solution volume, can

$$F = \frac{A_R}{A} \tag{1}$$

where A_R is the reference activity and A is the indicated activity for a condition to be corrected (vial with different volume, geometry etc.) (National Physical Laboratory, 2006).

Other important procedure that must be performed in order to guarantee workers and patient security in NMS is the activimeter calibration, which establishes a relation between a reference quantity value, provided by measurement standards, and a measurement result from an indicator (International Bureau of Weights and Measures, BIPM, 2008), which is, in this case, the activimeter from an NMS and/or a hospital.

In this study we suggest another influence factor which is normally disregarded, that must be considered in these procedures: remaining residual radioactive solution from a radiopharmaceutical transference from one vial to another that results in an error in the final activity.

Furthermore, we propose a new "in situ" calibration methodology establishment for activimeters, which is performed without the need of

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change the activimeter response, as presented in previous studies (Martins and Potiens, 2012; Zimmerman et al., 2004; Calhoun et al., 1987). Correction factors, F, must be applied and can be determined by means of Eq. (1):

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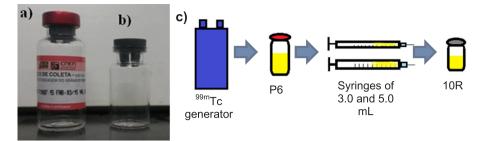


Fig. 1. (a) P6 and (b) 10R vials used in this study. (c) ^{99m}Tc handling scheme.

Table 1P6 and 10R vials dimensions (Baker, 2005). Numbers in parenthesis are the uncertainties.

	P6 vial	10R vial
Height (mm)	54.00(75)	45.0(5)
Diameter (mm)	21.75(25)	24.0(2)
Wall thickness (mm)	1.2(1)	1.00(4)
Maximum volume (mL)	13.8(1)	13.5(1)

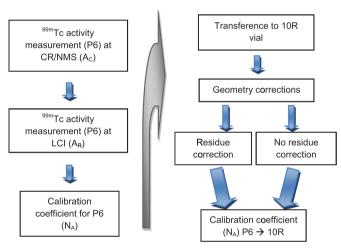


Fig. 2. Methodology 1 flowchart. First measurement is performed at radio-pharmaceutical production facility (CR) or Nuclear Medicine Service (NMS), and then P6 vial containing ^{99m}Tc is sent to calibration laboratory (LCI).

Table 2 Methodology 1 applied to radiopharmaceutical center (CR) at IPEN with and without residue corrections. Three central columns present N_A values with respective uncertainties in parenthesis.

P6/P6	10 R/P6 w/ corrections	10 R/P6 w/o corrections	Variation (%)
0.999(9)	0.999(9)	0.972(15)	2.7
0.980(9)	0.981(9)	0.945(20)	3.7
0.988(9)	0.988(9)	0.961(13)	2.8
0.988(9)	0.988(9)	0.961(14)	2.8
	0.999(9) 0.980(9) 0.988(9)	0.999(9) 0.999(9) 0.980(9) 0.981(9) 0.988(9) 0.988(9)	corrections corrections 0.999(9) 0.999(9) 0.972(15) 0.980(9) 0.981(9) 0.945(20) 0.988(9) 0.988(9) 0.961(13)

Table 3 Methodology 1 applied to Nuclear Medicine Service A with and without residue corrections. Three central columns present N_A values with respective uncertainties in parenthesis.

Activimeter Model	P6/P6	10R/P6 w/ corrections	10R/P6 w/o corrections	Variation (%)
NMS-A	0.98(1)	0.98(1)	0.94(2)	4.0

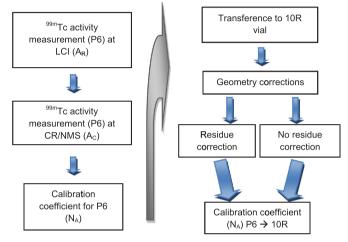


Fig. 3. Methodology 2 flowchart. First measurement is performed at calibration laboratory (LCI), then P6 vial containing ^{99m}Tc is sent to radiopharmaceutical production facility (CR) or Nuclear Medicine Service (NMS).

Table 4 Methodology 2 applied to Nuclear Medicine Service B and C with and without residue corrections. Three central columns present N_A values with respective uncertainties in parenthesis.

Activimeter Model	P6-P6	10R-P6 w/ corrections	10R-P6 w/o corrections	Variation (%)
NMS-B	0.990(9)	0.990(9)	0.964(15)	2.6
NMS-C	1.050(9)	1.051(9)	1.080(13)	2.7

taking the instrument to a calibration laboratory. Finally, a calibration coefficient comparison will be made when the calibration procedure is performed with and without residue corrections.

2. Methodology

2.1. Residual activity determination

For the residual activity determination ^{99m}Tc solution was obtained from a ^{99m}Tc generator (Osso et al., 2002) with a P6 vial (Fig. 1a). This solution was then transferred to the Instruments Calibration Laboratory (LCI/IPEN) reference vial, 10R (Fig. 1b), using 3.0 mL or 5.0 mL syringes. Each container was filled with saline solution to avoid volume errors and the residual activity was measured. A scheme for this procedure is presented in Fig. 1c. Vials dimensions are given in Table 1.

The total activity A_T is given by $A_T = A_F + A_{RC}$, where A_F is the solution in 10R vial and A_{RC} is the sum of the residual activity in the previous containers.

2.2. Calibration methodology establishment

Calibration methodology was established according to National

Physical Laboratory (NPL), United Kingdom (National Physical Laboratory, 2006) recommendation, using P6 and 10R vials. Reference activity was obtained with a Capintec NPL-CRC at LCI.

Activimeters calibration was performed in three NMS and in four different activimeters at IPEN radiopharmaceutical production facility (CR). The main advantage of an "in situ" calibration is the possibility to keep both radiopharmaceutical production and patient care working during the procedure. In both methodologies, a calibration factor in terms of the activity (N_A) was calculated from activities measured at NMS/CR, according to the equation:

$$N_A = \frac{A_R}{A_C} \tag{2}$$

where A_R and A_C are the activities measured with the reference and the NMS/CR equipment, respectively.

Letter N is normally used to indicate a calibration factor, and can be applied to different physical quantities, such as air-kerma (N_K), absorbed dose in water (N_D ,w) and activity (N_A) (International Atomic Energy Agency, 2007; International Atomic Energy Agency, 2000)

Having been determined for an activimeter, N_A must be applied for every measurement made in this equipment, along with any necessary correction factor, according to Eq. (3):

$$A_S = A_M N_A \prod_{1}^{n} F_n \tag{3}$$

where A_S is the corrected activity, A_M is the measured activity and $\prod_{i=1}^{n} F_n$ is the product of every necessary correction factors (volume, geometry, vial position inside the well-chamber etc.).

3. Results

Results obtained for radioactive residue measurement will be presented with calibration coefficients calculated in each calibration methodology. Two methodologies were performed with and without residue correction and N_A was obtained from both situations. Obviously, in a real calibration situation, this correction must always be made.

First methodology consists of measuring radiopharmaceutical activity at the laboratory where it was produced or in the NMS, using P6 vial. The sample is then sent to LCI for the reference activity measurement using the reference activimeter and subsequently it is transferred to the 10R vial. Geometry correction factor for P6 to 10R must be applied and it had already been determined in previous studies (Martins and Potiens, 2012; Kuahara et al., 2015).

In Fig. 2a procedure overview is presented, and the respective N_A values are presented in Tables 2 and 3.

The second methodology is very similar to the first. However, in this case, LCI receives ^{99m}Tc for the reference activity determination with P6 vial at reference activimeter. The sample is sent to NMS where a measurement and the calibration are made. The transference to 10R vial, geometry corrections and residue measurement are also performed, as previously presented. An overview of this methodology is presented is Fig. 3, and respective results in Table 4.

From Tables 1–3, a variation of up to 4% between procedures performed with and without residue correction can be observed, which is a high percentage considering that the test was made under controlled conditions and less residue was left inside vials. A much higher variation may be expected in NMS measurements. It is important to highlight

that these procedures were performed for P6 vial, but can be applied to any other vial used in the NMS.

4. Conclusions

Results show the importance of applying radioactive residue corrections when performing activimeters calibration. Variation of up to 4% in the measurement with and without residue correction was found, even under controlled laboratorial conditions. Greater errors are expected in a NMS. Both calibration procedures applying proper corrections presented high reliability and low uncertainties (lower than 1%).

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References

American Association of Physicists in Medicine, AAPM, 2012. The selection, use, calibration, and quality assurance of radionuclide calibrators used in nuclear medicine (AAPM-TG-181).

Baker, M., 2005. Calibration of the NPL secondary standard radionuclide calibrator for the new 10R Schott, Type 1+ vials. Appl. Radiat. Isotop. 63 (71–77), 2005.

Calhoun, J.M., Golas, D.B., Harris, S.G., 1987. Effects of varying geometry on dose calibrator response: cobalt-57 and technetium-99m. J. Nucl. Med. 28, 1478–1483.
Fahey, F.H., Treves, T.S., Adelstein, S.J., 2011. Minimizing and communicating radiation

risk in pediatric nuclear medicine. J. Nucl. Med. 52 (8), 1240–1251.

International Atomic Energy Agency, 2000. IAEA. Absorbed Dose Determination in

External Beam Radiotherapy. Vienna, Austria. (Technical Reports Series No. 398).

International Atomic Energy Agency, 2007. IAEA. Dosimetry in Diagnostic Radiology: An International Code of Practice. Vienna, Austria (Technical Report Series No. 457).

International Atomic Energy Agency, IAEA, 2006. Quality Assurance for Radioactivity Measurement in Nuclear Medicine. Vienna. TECHNICAL REPORTS SERIES No. 454. International Atomic Energy Agency, IAEA, 2010. Competency Based Hospital Radiopharmacy Training. Vienna. TRAINING COURSE SERIES No. 39.

International Atomic Energy Agency, IAEA, 2015. Activimetros – protocolo para el aseguramiento de la calidad de las mediciones. Vienna, Austria. (Proyecto ARCAL RLA 6074)

International Bureau of Weights and Measures, BIPM, 2008. International vocabulary of metrology – Basic and general concepts and associated terms (VIM).

Kuahara, L.T., Correa, E.L., Potiens, M.P.A., 2015. Determination of geometry correction factors to different vials used to radiopharmaceutical activity measurements. In: International Nuclear Atlantic Conference - INAC 2015, S\u00e3o Paulo.

Martins, E.W., Potiens, M.P.A., 2012. Determination of the influence factors of the radiopharmaceutical vials dimensions used for activimeter calibration at IPEN. Appl. Radiat. Isot. 70, 1281–1283.

Mettler Jr., F.A., Bhargavan, M., Faulkner, K., Gilley, D.B., Gray, J.E., Ibbott, G.S., Lipoti, J.A., Mahesh, M., McCrohan, J.L., Stabin, M.G., Thomadsen, B.R., Yoshizumi, T.T., 2009. Radiologic and nuclear medicine studies in the United States and worldwide: frequency, radiation dose, and comparison with other radiation sources—1950–2007. Radiology 253 (2), 520–531.

National Physical Laboratory, 2006. NPL. Protocol for establishing and maintaining the calibration of medical radionuclide calibrators and their quality control. A National Measurement Good Practice. Middlesex, United Kingdom.

Osso, J.A.Jr, Silva, N.C., Lima, A.L.V.P., Landini, L., Nieto, R.C., Moraes, V., 2002.

Development of ⁹⁹Mo-^{99m}Tc gel type generators at IPEN-CNEN/SP. World J. Nucl.

Med. 1, 189

Zimmerman, B.E., Cessna, J.T., Millican, M.A., 2004. Experimental determination of calibration settings for plastic syringes containing solutions of ⁹⁰Y using commercial radionuclide calibrators. Appl. Radiat. Isot. 60, 511–517.