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Keywords

high, dose, rate, prostate, online, brachytherapy, vivo, moskin, detectors, phantom, feasibility, study, dosimetry

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Online *in vivo* dosimetry in high dose rate prostate brachytherapy with MOSkin detectors: in phantom feasibility study

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Highlights

- A needles implant was set up in phantom to simulate prostate brachytherapy treatments
- In vivo dosimetry was performed in the urethral catheter with MOSkin dosimeters
- Dual-MOSkin detectors resulted to be accurate dosimeters to perform this task

Abstract

MOSkin detectors were studied to perform real-time *in vivo* dose measurements in high dose rate prostate brachytherapy. Measurements were performed inside an urethral catheter in a gel phantom simulating a real prostate implant. Measured and expected doses were compared and the discrepancy was found to be within 8.9% and 3.8% for single MOSkin and dual-MOSkin configurations, respectively. Results show that dual-MOSkin detectors can be profitably adopted in prostate brachytherapy treatments to perform real-time *in vivo* dosimetry inside the urethra.

Keywords

High dose rate, prostate brachytherapy , MOSFET, *in vivo* dosimetry

1. Introduction

1.1 *In vivo* dosimetry in radiotherapy

The recent developments of more sophisticated radiotherapy and brachytherapy (BT) techniques call for the improvement of instruments and methodologies employed for the quality control of the performed treatments. Due to the achievable high conformity of modern BT associated with steep dose gradients, a careful verification of the accuracy in the delivered dose distributions, as planned by the Treatment Planning System (TPS) through mathematical models, is gaining importance.

In vivo dosimetry is a reliable method to compare planned and delivered dose distributions, representing therefore a valid tool to systematically verify treatment accuracy and improve radiotherapy quality control (Lambert et al 2007, Mijnheer 2008). Particularly advantageous for *in vivo* dosimetry are detectors that allow on-line dose reading. These dosimeters provide in fact real-time measurements during treatment, avoiding therapy misadministration and allowing at the same time intraoperative dose re-planning for treatment error correction.

Current methods for *in vivo* dosimetry are mainly based on the application of thermoluminescence detectors (TLDs) (Toye et al 2009) or semiconductor diodes (Waldhäusl et al 2005). TLDs involve offline process providing the integral dose absorbed during patient treatment and require special procedures in order to achieve good precision of the results. On the other hand, diodes show rapid processing time, high sensitivity and immediate reuse, however, they show a high energy dependence and the delivered dose is therefore not promptly inferred from the diode reading. Moreover, the major disadvantages of diodes are their relative large sizes, which make them unable to be held in many catheters placed inside the patient to perform *in vivo* dosimetry.

New detectors such as fiber optic coupled scintillation dosimeters (Suchowerska et al 2007, Therriault-Proulx et al 2011) and metal oxide semiconductor field effect transistors (MOSFETs) (Zilio et al 2006, Fagerstrom et al 2008) have recently been introduced to perform *in vivo* dosimetry. In particular, MOSFETs show many advantages, such as good spatial resolution, high

sensitivity, real time read-out without deterioration of information, negligible radiation field perturbation owing to their small size and ease of use. In particular, great interest was dedicated to the application of MOSFETs to BT, because the typical large dose gradients achieved in BT necessitate a small detector with a reduced active volume for accurate dosimetry. In this work, a specific type of MOSFET dosimeter called “*MOSkin*” which has developed by the Centre for Medical Radiation Physics (CMRP) of the University of Wollongong (Australia) (Qi et al 2007, Kwan et al 2008, Kwan et al 2009) has been studied.

1.2 High dose rate prostate brachytherapy

High dose rate (HDR) prostate BT allows the delivery of local and high conformal dose directly into the tumour, minimizing exposure of the surrounding healthy tissues. Due to the large dose delivered to the target in a single fraction and the dose constraints to be simultaneously satisfied for organs at risk, it is very important to have as small as possible discrepancy between planned and delivered dose. The development and application of reliable and accurate methods for monitoring the dose delivered to critical organs is therefore crucial.

Among these organs at risk, the urethra is most likely susceptible to acute and/or late toxicity resulting from the treatment (i.e. urethritis, stenosis), as it is inside the target volume (figure 1a). However, its localization for treatment planning purposes is particularly difficult due to images artefacts generated by the presence of source catheters, especially if transrectal ultrasound imaging is performed. Moreover, source catheters are themselves difficult to be accurately localized on the same images and therefore calculated dose distributions are susceptible to inaccuracies (figure 1b). The real time dosimetry in the urethra is therefore very important and will be supplementary to reinforce existing QA programs.

Studies aimed at characterizing the dosimetric properties of *MOSkin* dosimeters have already demonstrated that they are promising instruments for performing *in vivo* dosimetry during HDR BT treatments (Qi et al 2007, Kwan et al 2008, Qi et al 2012). Measurements finalized to detect the

accuracy of the dosimeters and the change in sensitivity as a function of depth and angle of incidence of the radiation, have already shown good agreement between *MOSkin* response and dose calculated by the TPS (Hardcastle et al 2010). Aim of this work was to study and develop the applicability of the *MOSkin* dosimeters for urethral dose measurement in prostate HDR BT.

2. Materials and Method

2.1 MOSkin dosimetry system

The design of this particular type of MOSFET is optimized to measure dose in steep dose gradients. Different from other commercial MOSFETs, *MOSkin* die is embedded in a thin kapton layer and hermetically sealed with water-equivalent flexible carrier of reproducible thickness and avoid traditional wire bonding with high $-Z$ wires. The sensitive volume, defined by the volume of the gate oxide, is $4.8 \times 10^{-6} \text{ mm}^3$. *MOSkin* detectors can be adopted alone or coupled in a face-to-face arrangement. This face-to-face dual-*MOSkin* arrangement is referred to in this text as the “dual-*MOSkin*”. The dual-*MOSkin*, proposed and developed at CMRP, allows for angular-independent measurements as it compensates the naturally asymmetrical structure of the MOSFET chip relative to the beam direction (Hardcastle et al 2010). The dosimetry system adopted with *MOSkin* detectors includes a microprocessor based reader which is connected to a laptop provided with a dedicated software (“MosPlot”). The computer data acquisition system measures periodically the instantaneous voltage signal with a user defined frequency. The “MosPlot” software allows for the online graphical representation of the change in the threshold voltage (proportional to accumulated dose) or increments in the threshold voltage for consecutive readouts (proportional to dose rate).

2.2 Nucletron Microselectron-HDR Brachytherapy facility

BT irradiations were performed using a Microselectron-HDR (Nucletron, Veenendaal, the Netherlands) high dose rate remote afterloading device. The facility is provided with a ^{192}Ir

radioactive source which has an active length of 3.6mm and a diameter of 0.65mm. The source is sealed inside a capsule that is welded to one end of a flexible steel cable and the treatment unit moves it to the required dwell positions.

Experimental determinations of absolute dose rates to water from high-energy sources should have direct traceability of the source Air Kerma Strength S_k to a primary or secondary standard dosimetry laboratory (PSDL OR SSDL) (Perez-Calatayud et al, 2012). In our case, the S_k was certificated by the vendor with 5% ($k=3$) uncertainty using a calibration instrument traceable to the standard of the PSDL Physikalisch-Technische Bundesanstalt (PTB, Braunschweig, Germany). Moreover, the certified S_k was independently verified at our hospital before its clinical application, by the qualified medical physicist using a calibrated well-type chamber traceable to a PSDL. The well-type chamber has a vented sensitive volume of 200cc (Source Dosimetry System. Nucletron, Veenendaal, the Netherlands) (Nath et al 1997) and has proven to be a fast and precise measuring instrument (Azhari et al, 2012). The delivered dose distribution is calculated by TPS Oncentra Brachytherapy (Nucletron, Veenendaal, the Netherlands) and is based on the recommendations of the American Association of Physics in Medicine (AAPM) TG-43 report (Rivard et al 2004)..

2.3 MOSkin and dual-MOSkin calibration and response at increasing source-detector distance

Each single MOSkin and dual-MOSkin was calibrated in a water phantom at a distance of 38mm from the ^{192}Ir HDR source. A suitable support for accurate and reproducible detector and source positioning was developed to perform this task (figure 2) and the experimental set-up was accurately defined by means of CT imaging (slice thickness: 0.8mm) and catheter digital reconstruction. Five irradiations were performed for defined source dwell times and the mean values of MOSkin threshold voltage changes were obtained. Each detector was finally calibrated considering the correspondent dose value calculated by means of the TPS in the same position where the active component of the dosimeter was placed. To investigate MOSkin sensitivity variation vs source–detector distance due to changing of the photon energy spectrum from the Ir-

192 source, distance dependent correction factor (CF) was evaluated for measurements performed at locations other than the calibration position, ranging from 7mm to 46mm and from 11mm to 44mm along the source transverse axis in the water phantom for *MOSkin* and dual-*MOSkin* dosimeters, respectively. In each one of these locations, measurements were repeated five times.

2.4 In phantom simulations of prostate treatments

With the aim of performing dose measurements with implants and conditions simulating typical prostate treatments, a suitable gel phantom was realized. The phantom has a cylindrical shape with length of 14.5 cm, and a diameter of 17.6 cm. The urethral catheter and 14 interstitial needles were placed inside the gel phantom through a real prostate treatment template. Single or dual-*MOSkins* were inserted in the urethral catheter. CT images (0.8mm slice thickness) were acquired in order to precisely localize needles and the urethral catheter containing the dosimeter. In the treatment planning phase, the presence of a prostate was virtually created by drawing it on the CT images around the needles implant. The urethral catheter was as well segmented and different treatments were planned by prescribing the dose to the outlined prostate. The expected doses to the dosimeters were evaluated from the TPS. In fact, *MOSkin* detectors could be accurately recognized on CT images' reconstructions as small radiopaque spots inside the urethral catheter (figure 3). During irradiation, the plastic needles were connected to the BT afterloader system through transfer tubes and *MOSkins* were connected to the CMRP computerized reader for the on-line acquisition of dose measurements (figure 4).

A series of measurements of the same irradiation set-up was performed with different dose prescriptions according to the developed treatment plans. The total urethral dose measured with the single or dual-*MOSkin* detector was compared to the expected dose in the same point. Moreover, the contributions to the total dose given by the source moving in each single needle were calculated separately and compared to the measured ones.

3. Results and discussion

Single *MOSkin* and dual-*MOSkin* detectors calibration factors obtained in the water phantom on the Ir-192 source resulted 0.38 cGy/mV and 0.30 cGy/mV, respectively. Distance dependent CFs at different source-detector distances for single *MOSkin* data were in accordance with those reported in Qi et al (2012); the significant increase of CF at decreasing source-detector distance was observed. In contrary to single *MOSkins*, CFs for dual-*MOSkins* showed a relatively flat response with changing the Ir-192 source-detector distance (figure 5). It is important to note that reference dose data to perform dosimeter calibration were not obtained experimentally by a further dosimeter, but were those resulting from the TPS and cross-checked with a home made software based on the AAPM TG-43 algorithm (Rivard et al, 2004). The chosen Λ , $G(r,\theta)$, $F(r,\theta)$ and $g(r)$ factors were those tabulated in Daskalov et al (1998), which were obtained by means of a Monte Carlo (MC) photon transport code knowing the characteristics of the Nucletron model mHDR-v2 source. Consensus on these data has been widely achieved and the resulting AAPM TG-43 algorithm calculations, when performed in reference conditions of unbounded water medium, are highly accurate (Daskalov et al, 1998). In fact, the role of experimental dosimetry for high-energy BT may be more limited than MC-based dosimetry. With high energy BT sources, range and spatial resolution limitations are for instance not of concern for MC methods (Perez-Calatayud et al, 2012).

Comparison of total dose measured in the gel phantom for the BT prostate implant with the single-*MOSkin* detector and the expected doses in the same points are reported in Table 1. Measurements were repeated for three different treatment plans with different dose prescriptions. The maximum discrepancy between expected and measured dose was found to be 8.9%. Figure 6a shows measured and calculated contribution to the total dose resulting by the source dwelling in each of the 14 needles adopted to simulate the prostate implant, for one of the three different sets of measures.

Excluding needles that give a low contribution to the cumulative dose ($<20\text{cGy}$), single point dose differences resulted less than 15%.

Comparison of total doses measured with the dual-MOSkin detector and the expected doses in the same point is reported in Table 2 for three different measurement sessions. It was found that discrepancy was within 3.8%. Figure 6b shows measured and calculated contribution to the total dose resulting by the source moving in each of the 14 needles adopted to simulate the prostate implant, for one of the measured treatment sessions. Excluding needles that give a low contribution to the cumulative dose ($<20\text{cGy}$), single dose differences resulted in general smaller than those resulting with a single MOSkin readings (i.e. max 8%).

Results obtained for both single MOSkin and dual-MOSkin dosimeters are summarized in the plot in figure 7. Dual-MOSkin configuration provides greater accuracy than a single MOSkin detector as angular isotropy and depth dose are greatly improved. Single MOSkin accuracy could be partially increased by applying measured CFs at changing source-detector distances and further CFs that take into account its orientation with respect to the source position. However, CFs are almost impossible to be correctly adopted whenever MOSkins are used as real-time *in vivo* dosimeters because source-detector distances and MOSkin orientation cannot be easily quantified and taken into account.

4. Conclusions

In prostate HDR BT treatments, a very high dose over a few fractions is delivered. In such conditions, real time dosimetry is most valuable in detecting a dose error at the onset of treatment, providing the capability of immediate corrections. Measurements performed in a phantom simulating a typical prostate implant with MOSkin detectors placed within the urethral catheter have shown that they allow evaluation of the actual dose to the urethra during a treatment fraction. Through an online comparison between the measured and expected dose, the dosimetry system

might offer a further security to detect and prevent significant errors in the clinical routine of HDR BT prostate treatments.

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Figure captions

Figure 1: a) 3D graphical representation of an HDR brachytherapy plan of the prostate (dark red). Urethra, rectum and bladder are represented in yellow, green and blue, respectively. Source catheters are drawn as green lines, source dwell positions are represented by red spheres and resulting dose distribution (95% isodose) is given in light blue; b) transversal prostate image with the resulting dose distribution. Characteristic hot spots are present around some of the implanted needles.

Figure 2: Support to be placed in a water phantom which was developed for dosimeters calibration. Source catheter can be placed at different distances from the dosimeter.

Figure 3: a) sagittal CT reconstruction and b) transversal CT image including the urethral catheter. *MOSkin* dosimeter is clearly recognizable as a radiopaque spot inside the catheter.

Figure 4: Experimental set-up used for in phantom measurements.

Figure 5: Correction factors evaluated at different source-detector distances for single- (white circles) and dual-*MOSkin* (gray diamonds) detectors. Plotted error bars are given by the standard deviation of the measurements and result in the values of 0.025, 0.018, 0.018, 0.019, 0.020, 0.019 and of 0.046, 0.042, 0.043, 0.048, 0.034, 0.032 for single- and dual-*MOSkin*, respectively.

Figure 6: Measured and expected doses given by the source moving in each of the 14 needles for a) *MOSkin* and b) dual-*MOSkin* detectors.

Figure 7: Comparison between single *MOSkin* and dual-*MOSkin* in terms of measured mean dose differences between expected and measured values, standard errors and standard deviations.

Tables

Table 1: Total urethral dose measured with a single-MOSkin dosimeter for three different treatments.

Total urethral dose			
	Measured Dose (cGy)	Calculated Dose (cGy)	Dose difference (%)
1	727.2	798.1	-8.9
2	809.2	828.1	-2.0
3	732.0	799.0	-8.0

Table 2: Total urethral dose measured with a dual-MOSkin dosimeter for three sessions of the same treatment.

Total urethral dose			
	Measured Dose (cGy)	Calculated Dose (cGy)	Dose difference (%)
1	1139.5	1148.5	-0.8
2	1126.7	1148.5	-1.9
3	1104.3	1148.5	-3.8

Figures

Figure 1

Figure 2

Figure 3

Figure 4

Figure 5

Figure 6

Figure 7