

# Automatic Brachytherapy Seed Placement Under MRI Guidance

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**Abstract**—The paper presents a robotic method of performing low dose rate prostate brachytherapy under magnetic resonance imaging (MRI) guidance. The design and operation of a fully automated MR compatible seed injector is presented. This is used with the MrBot robot for transperineal percutaneous prostate access. A new image-registration marker and algorithms are also presented. The system is integrated and tested with a 3T MRI scanner. Tests compare three different registration methods, assess the precision of performing automated seed deployment, and use the seeds to assess the accuracy of needle targeting under image guidance. Under the ideal conditions of the *in vitro* experiments, results show outstanding image-guided needle and seed placement accuracy.

**Index Terms**—Brachytherapy, IGI, image-guided robot, MR Compatible.

## I. INTRODUCTION

PROSTATE cancer is the most common cancer detected in males. It is estimated that approximately thirty three percent of the new cancer cases identified in 2006 will be prostate cancer [1]. Also, it is estimated that prostate cancer will be responsible for ten percent of cancer related deaths in the male population of the United States [1]. Radical surgery or radiotherapy are the most commonly used curative treatment options for prostate cancer [2]. Radiotherapy kills cancer cells by delivering ionizing radiation to the prostate. The radiation can be delivered using external beam irradiation, by high dose radiation sources temporarily delivered percutaneously, or by implanting radioactive pellets (brachytherapy).

Prostate brachytherapy is usually performed under transrectal ultrasound (TRUS) image guidance. The radioactive pellets (seeds,  $^{125}\text{I}$  or  $^{103}\text{Pd}$  isotopes) are deployed through transperineally inserted needles guided by a template according to a plan [3]–[5]. This pretreatment plan (dosimetry) should ensure that the prostate will receive a radiation dose high enough to destroy the cancer while sparing healthy surrounding tissues as much as possible. Long-term oncological results indicate that

brachytherapy is a good treatment option for properly selected patients [6]–[8].

An important factor that influences the outcome of the brachytherapy treatment is the accuracy of seed placement [9]. For example, if a cancer spot does not receive the adequate irradiation, this may lead to relapse of the disease. Accurate seed placement requires a good intra-operative image as well as a good delivery method. Arguably, the best prostate image is provided by MR imaging [10]. Moreover, studies showed promising results in prostate cancer imaging using MR spectroscopy [11]. Therefore, the use of MR imaging for treatment planning and guiding the delivery of therapy could potentially improve clinical outcomes compared to traditional methods

Currently, the needle placement is performed with the use of a template presenting a rectangular pattern of holes used to guide the needles. This allows for placing the needles only at the locations of the holes (usually 5 mm apart) and with parallel needle trajectories. In some patients, this impedes accessing part of the gland due to the interference of the needles with the pubic bone [12]. These problems may be overcome by replacing the needle template with a robotic device for manipulating the needle. Several research groups reported prototypes for robotic assisted brachytherapy of various levels of integration and complexity.

The systems reported for robotic assisted prostate interventions can be classified by the imaging modality used for guidance (i.e., US-guided and MR-guided). An example of an ultrasound-guided system is the one developed by Wei *et al.* [13]. Their system uses an industrial robot, and a TRUS probe with automated scanning motion. The coordinates of the ultrasound probe with respect to the robot are computed using a calibration procedure. This allows for orienting the robot to any target specified in the ultrasound image. In the first reported prototype, the robot was used only as a needle holder, with the physician being responsible for needle insertion and seed deployment.

MR-guided interventions require needle manipulators that are MR compatible. Building MR robotic systems is a challenging task due to the restrictions imposed by the high intensity magnetic fields of the MR as well as the EM/RF interference that can deteriorate the image signal-to-noise ratio [14], [15]. For example, the ubiquitous electromagnetic actuators may not be used in the MR environment. The use of electricity and some electronic devices in the magnet room may be possible, under certain specifications and shielding.

Several electrically actuated systems for MR-guided interventions were reported. Chinzei *et al.* developed a MR compatible manipulator for prostate needle interventions [16]. This system automatically positions a needle guide allowing for prostate biopsies and radioactive seed placement under MR guidance in an open MRI scanner. The system is actuated using piezoelectric motors located outside the MR field.

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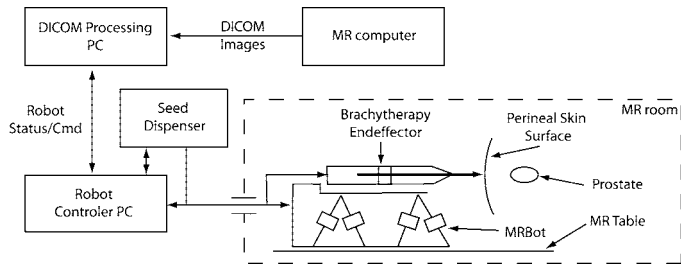


Fig. 1. System Diagram.

Other MR compatible robotic systems using piezoelectric actuators were reported by Larson *et al.* [17], Tsekos *et al.* [18], and by Hempel *et al.* [19]. The first system was developed for breast interventions and the later two were developed for general chest and abdominal interventions. All three manipulators use piezoelectric actuators to manipulate needle guides for manual needle insertion. The system developed by Hempel *et al.* was subsequently modified to use pneumatic pistons for actuation and a commercial version is presently being developed (Innomedic GmbH, Germany).

Krieger *et al.* presented a device for transrectal needle placement in the prostate under MR guidance [20]. Their device is manually operated but uses special localization coils to track the real-time position of the device in the MR image space. The system was successfully tested in animal and human studies at the NIH.

In contrast with the previously reported needle-guide systems, the one presented in this paper is more complex allowing for fully automatic needle insertion and seed deployment under MR guidance. In addition, this fully actuated system is entirely made of nonmagnetic and dielectric materials, and is electricity free exclusively using pneumatics and optics. The system consists of an end-effector mounted on a MR compatible robotic manipulator [21] (MrBot) and it incorporates a new type of pneumatic actuators (PneuStep) developed for fully MR compatible actuators [22]. This paper presents the MR compatible automated seed placement end-effector, image registration and guidance algorithms. The results of an accuracy study for MR-guided seed placement in agar and *ex vivo* models are also presented. The key novelty of the reported system is the automatic seed deployment under MR guidance.

## II. SYSTEM DESIGN

In conventional prostate brachytherapy, a number of needles preloaded with seeds are inserted into the prostate under US guidance. Alternatively, a single needle can be used to place all seeds according to a treatment plan [23]. In this case, the needle is sequentially placed at the desired seed locations and the radioactive seed is deployed through the cannula of the needle. The seeds located on the same needle trajectory can be deployed sequentially in the same needle insertion starting from the deepest location. Therefore, prostate seed placement requires for the robotic system to perform the following tasks: 1) needle orientation; 2) needle insertion; 3) seed deployment. The overall system diagram is presented in Fig. 1. Needle orientation is performed using the MR compatible manipulator, MrBot. Needle insertion and seed deployment are performed by a special brachytherapy end-effector.

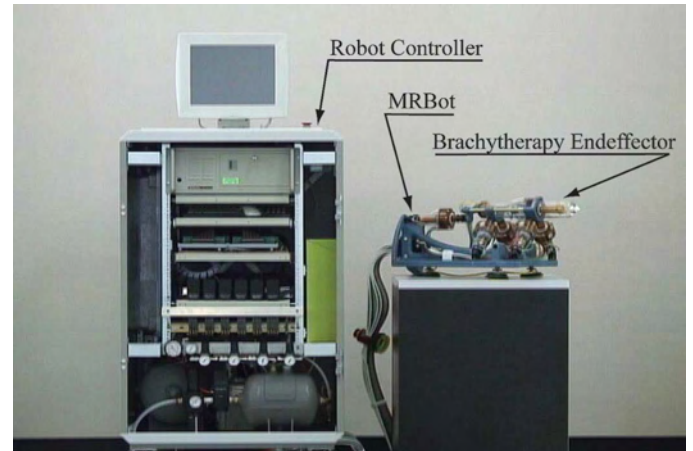


Fig. 2. MR compatible automated brachytherapy system components.

### A. MR Compatible Manipulator

The MrBot robot is a 5 degree of freedom (DOF) (3T+2R) MR compatible manipulator presented in Fig. 2 [21]. It allows for positioning the needle injector end-effector in all directions (3T) and its orientation about two directions normal to the needle axis (2R). Its workspace allows the placement and alignment of the needle towards any prostate target, assuming that initially the robot is roughly aimed towards the prostate.

The manipulator is pneumatically driven using custom designed MR compatible actuators, PneuStep [22]. The PneuStep is a new type of pneumatic motor, developed especially for MRI compatible robots. This achieves easily controllable precision of motion (0.055 mm in the design used here) by using a stepper motor principle. Position feedback is implemented using fiber-optic sensors. The manipulator is commanded from a control cabinet through a bundle of 6-m-long air hoses and optic fibers. Each PneuStep motor requires 3 air hoses, 4 fibers for incremental encoding, and 2 fibers for a limit switch sensor. During the intervention, the cabinet resides outside the MR room. The cable connecting the control cabinet to the manipulator is passed through the access port of the imager's room.

### B. MR Compatible Automatic Seed Placement System

The seed placement mechanism includes a MR compatible needle injector end-effector and the non MR compatible components that are attached to the control cabinet. Like for the robot, pneumatics and optics are used for the actuation and respectively sensing.

The needle injector was designed using the observation that high speed needle insertion reduces soft-tissue deflection. This was experimentally tested *ex vivo* and was also reported by other researchers [24]. The injector schematically represented in Fig. 3(b) is composed of two concentric cylinders ③ and ④ translated with respect to the case ① by the PneuStep motor ②. The case ① is attached to the robot. The motor ② allows the adjustment of the insertion depth by shifting needle cylinder and thus its stroke. The needle cylinder ③ allows for the rapid needle insertion, while the cylinder ④ allows for maneuvering the stylet of the needle for seed deployment. The insertion speed is determined by the pressure applied to the piston ③ through the port P2. Currently, during insertion the applied pressure with a value of 40PSI. This provides an

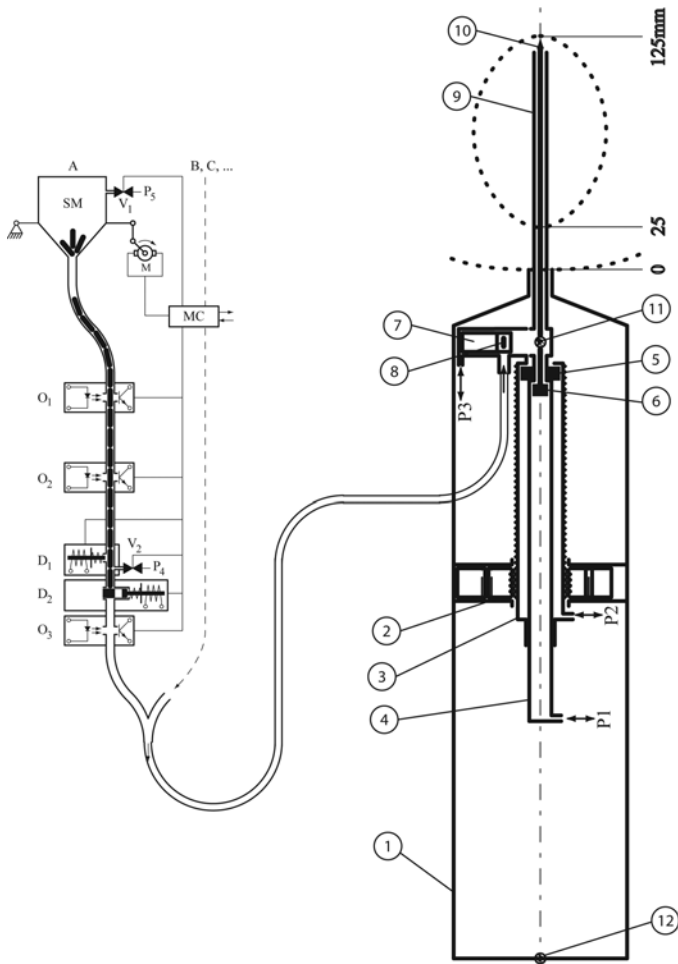


Fig. 3. Seed injector diagram; a) Seed dispenser b) Seed placement end-effector b.1) case; b.2) MR compatible actuator (PneuStep); b.3) needle cylinder; b.4) stylet cylinder; b.5) needle piston; b.6) stylet piston; b.7) seed drawer and cylinder; b.8) seed in the drawer; b.9) needle; b.10) stylet; b.11) seed fiber optic sensor; b.12) retract fiber optic sensor.

insertion speed of approximately 0.5 m/s. A third pneumatic piston is used for the seed drawer 7. Several holes were placed on the drawer mechanism to allow the possible piston leakage to be discharged in the air. These discharge holes will prevent blood and seeds from being aspirated into the mechanism.

The seed dispenser presented in Fig. 3(a) is composed of a jar with a funneled bottom that is shaken using a motor, the motor controller and the seed locking, sending and counting mechanisms. The seeds are preloaded into the jar and they drop into the funnel as the jar is shaken. The funnel proceeds to a tube leading to the sending system. The tube also serves as a sending buffer. Two photoelectric sensors  $O_1$  and  $O_2$  are used by the controller (MC) to detect the status of the seed column. MC actuates the motor  $M$  when  $O_2$  is open, and stops the jar shaking when  $O_1$  is obstructed. The sensor  $O_3$  is used to latch a bistable when a seed passes through. This provides a positive indication that a seed left the seed dispenser. It is possible to connect multiple seed dispenser channels to the same controller and sending tube, allowing for the programmatic use of different types of seeds. In the current implementation, the seed dispenser has two channels. The controller MC is connected to the main computer

allowing the user to command the operations required for seed sending.

The procedure for deploying a seed comprises the following steps.

- 1) MrBot is moved such that the tip of the needle is at the skin entry point and the injector is aligned towards the target.
- 2) The insertion depth is set by positioning the needle cylinder b.3. with the PneuStep motor b.2.
- 3) The stylet is inserted into the needle by applying pressure on port b.P1.
- 4) The needle is inserted to the desired depth by applying pressure on port b.P2.
- 5) The stylet is retracted by applying vacuum on port b.P1.
- 6) The seed drawer b.7 is already aligned with the seed transport tube, vacuum being applied on port b.P3.
- 7) Bring one seed from the dispenser:
  - a) The column of seeds in the seed dispenser is blocked by actuating the plunger a.D1. This holds the second seed in the column thus allowing for only one seed to be sent.
  - b) The first seed from the column is released by opening the gate a.D2.
  - c) The seed is sent through the feeding tube to injector by applying pressure on port a.P4. The seed reaches the drawer b.7.
- 8) Push the drawer to bring the seed at the axis of the needle by applying pressure on P3.
- 9) Restore the dispenser:
  - a) Release the pressure on P4.
  - b) Lock the seed column by releasing D2 and D1.
- 10) Retract the needle one seed length by actuating the PneuStep motor b.2.
- 11) Slowly push the seed through the needle with the stylet by applying flow limited pressure on P1.
- 12) Retract the stylet by applying vacuum on port P1.
- 13) Lift the drawer by applying vacuum on port P3.
- 14) If there are more seeds to be deployed on the same trajectory then adjust the needle position using the PneuStep 2 and go to Step 7).
- 15) Retract the needle by applying vacuum on port P2.
- 16) If there are more seeds to be deployed on a different needle path then go to Step 1).

The actuation, seed sending tube, and sensors are included in the 6-m-long bundle of hoses connecting the robot to the control cabinet. The described deployment system was implemented and tested with several thousands of seeds.

The robot-seed injector ensemble was tested by automatically placing patterns of seeds in agar and *ex vivo* models at arbitrary locations [21]. Then, registration and image guidance algorithms were integrated to place the seeds at targets specified in the image.

### C. MRI – Robot Registration and Image Guidance Algorithms

In order to place an instrument to a location selected in a MR image, it is necessary to compute the 6 DOF transformation from the robot space to the image space. The registration is computed using a special marker embedded in the end-effector.

Image to robot registration methods have been previously reported by other authors and are frequently used in computed tomography (CT)-guided interventions. Susil *et al.* [25] reported

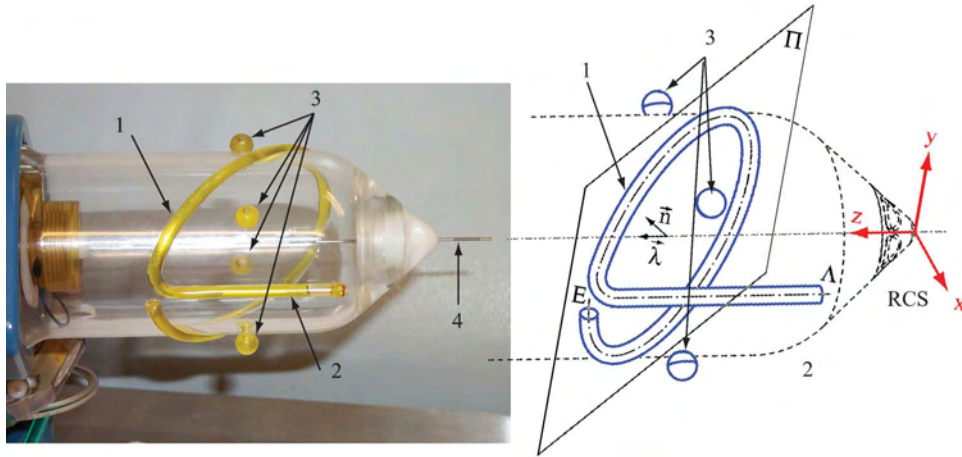


Fig. 4. MR Registration marker attached to the seed injector; 1) Ellipse, 2) Line, 3) Control Points, 4) Needle.

a registration method for CT-guided interventions that uses one single image for computing the parameters. The method was generalized by Lee *et al.* to an algorithm that allows the registration of a plane to a set of lines [26]. The method requires the placement of at least six line markers on the robot. Since the MRI does not raise radiation exposure problems, this one-slice CT method is not as appealing.

Registration methods have been developed purposely for the MR based on the image [27] or by using special coils [20]. While the coil-based registration is accurate and provides real-time data, it is not easily portable from one scanner to another. Image-based registrations are also accurate and imager independent, thus simplifying portability, but are not real-time. This, however, is not an issue because the MrBot is fully encoded.

A special marker, fitted to the seed injector end-effector, was developed, Fig. 4. The marker is composed of a line and an ellipse. The intersections of the marker with the planes of the MR slices are used to construct the registration transformation from the robot to the image.

Three different registration algorithms were implemented and tested. 1) line-ellipse (LE) algorithm. This algorithm automatically segments the image and computes the median point for each trace, then the set of points is matched to the line and the ellipse. 2) line-plane (LP) algorithm. This algorithm is similar with LE, the only difference is that instead of fitting an ellipse the intersection points are fitted to the plane of the ellipse. 3) image-model registration (IMR) algorithm. This registration works in two steps; first, an approximation of the registration is computed using the LP algorithm; second, the transformation computed in the first step is subsequently enhanced using an image to model registration.

The first two algorithms start by binarizing the DICOM acquired MR volume and finding the marker intersections within each image slice. Each image blob is analyzed and the median point is selected. The  $x$  and  $y$  in slice coordinates and the  $z$  position of the slice provide a point in the 3D space. The collection of points is divided in two sets  $\mathcal{P}_L$  and  $\mathcal{P}_E$ ;  $\mathcal{P}_L$  containing all the points belonging to the line and  $\mathcal{P}_E$  containing all the points belonging to the ellipse. These two sets are the primary data for the LE and LP algorithms.

1) *Line-Ellipse Algorithm*: comprises two main steps. In the first step, the coordinates of the line are reconstructed from the

set  $\mathcal{P}_L$ . This provides 4 DOF of the transformation. The remaining 2 DOF, namely a rotation about the line and the translation along the line, are resolved by fitting the ellipse over the set  $\mathcal{P}_E$ . The algorithm for computing the transformation from the robot space to the image space is outlined in Appendix A.

2) *Line-Plane Algorithm*: *Line-plane algorithm* uses the points  $\mathcal{P}_L$  to identify a line and the set of points  $\mathcal{P}_E$  to identify a plane. The line is described by a position vector  $\mathbf{v}_l$  and a unitary direction vector  $\mathbf{d}_l$ . The plane is identified by a position vector  $\mathbf{v}_p$  and a unitary plane normal  $\mathbf{n}_p$ . The line and plane parameters are identified using a least-squares fitting. Then, taking into account the particular configuration of the marker, the robot coordinate system vectors are expressed in image coordinates as

$$\mathbf{z}_r^i = \mathbf{d}_l; \mathbf{y}_r^i = \mathbf{z}_r^i \times \mathbf{n}_p; \mathbf{x}_r^i = \mathbf{y}_r^i \times \mathbf{z}_r^i.$$

The rotation matrix from the robot space to the image space becomes

$$\mathbf{R}_r^i = (\mathbf{x}_r^i \quad \mathbf{y}_r^i \quad \mathbf{z}_r^i).$$

The translation from RCS to ICS is computed by matching the intersection point  $vi$  between the ellipse plane and the line in ICS and RCS. The intersection point in ICS,  $\mathbf{v}^i$ , is computed from the two sets of points  $\mathcal{P}_E$  and  $\mathcal{P}_L$ . The position of the intersection in RCS,  $\mathbf{v}^r$  is known from the marker construction. The translation between the RCS and ICS is computed as

$$\mathbf{T}^i = \mathbf{v}^i - \mathbf{R}_r^i \mathbf{v}^r.$$

3) *Image-Model Registration*: tries to match the 3D marker model over the image. This method uses the registration algorithm implemented by the class *ImageToModelRegistration* class from the Insight Toolkit [28]. The initial approximation of the registration transform was computed using the LP-algorithm. The class required the implementation of a model to image error metric used in the optimization process. The metric was defined as

$$M(\mathbf{R}_r^i, \mathbf{T}^i) = \sum_{\mathbf{p} \in \text{Model}} \text{Pixel}(\mathbf{R}_r^i \mathbf{p} + \mathbf{T}^i)$$

where  $\text{Pixel}(\mathbf{p})$  is the value of the pixel at the position  $\mathbf{p}$ .

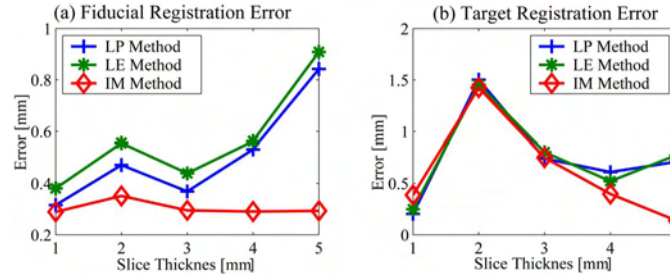


Fig. 5. Registration Error.

Once the registration information is available the user needs to specify the entry point  $\mathbf{p}_e^r$  for the robot and the target point  $\mathbf{p}_t^i$ . These two points completely specify the trajectory of the needle and the last seed placement position. The user will also specify the number of seeds and the spacing between seeds. The points are translated in RCS using the registration transformation. Then,  $\mathbf{p}_e^r$  and  $\mathbf{p}_t^r$  are used to compute the desired robot coordinates

$$\mathbf{T}_R = \mathbf{p}_e^r; d = \|\mathbf{p}_t^r - \mathbf{p}_e^r\|_2; \alpha_x = \arcsin\left(\frac{d_n[1]}{\cos \alpha_y}\right)$$

$$\alpha_y = -\arcsin(d_n[0]); d_n = \frac{\mathbf{p}_t^r - \mathbf{p}_e^r}{\|\mathbf{p}_t^r - \mathbf{p}_e^r\|_2}.$$

The joint positions are computed from  $\mathbf{T}_R$ ,  $\alpha_x$ ,  $\alpha_y$ , and  $d$  using the inverse kinematics of the robotic manipulator.

### III. RESULTS

#### A. Image Registration Tests

The three registration methods were implemented and tested. The following procedure was carried through. The robot was placed in the MRI scanner. Six volumes were acquired with slice thicknesses of 0.5, 1, 2, 3, 4, and 5 mm with the end-effector in the same position. The registration was performed using the datasets with 1-, 2-, 3-, 4-, and 5-mm slice thickness and the registration transformation was verified using the traces of the control points in the 0.5-mm dataset. The registration error was computed as follows; the registration transformation was used to compute the center of each control point. The result was compared against the control marker center retrieved from the .5-mm dataset. Usually, the registration accuracy is evaluated in terms of fiducial localization error (FLE), fiducial registration error, and the target registration error [29]. The FLE degrades with the slice thickness. Specifically, the localization along the Z-axis of the scanner can not be less than the slice thickness. However, thin slices require a long acquisition time in order to obtain a good signal to noise ration. Therefore, it is desirable to be able to compute an accurate registration with thick slices. The FLE for the verification scan was in the order of half millimeter. Therefore, the spatial localization of the control points is very accurate. The FRE was estimated as the mean localization error for the four control points. Fig. 5(a) shows the FRE computed for the different datasets and for each registration algorithm presented above.

The TRE was estimated assuming that the target is usually placed along the needle at a distance of 125 mm from the RCS. The position of the target was computed using the control points retrieved from the .5-mm scan and by using the registration

transformation computed from the other five datasets. Fig. 5(b) presents the estimated TRE corresponding to different registration algorithms used. The graphs show that all three methods performed reasonably well. We noticed that the accuracy deteriorates as the slice thickness increases for LP and LE methods while the IMR method is more robust. However, the computational cost of the IMR method is significantly higher than the others. The decrease in performance with the increase in slice thickness can be explained by the fact that the FLE increases with the slice thickness. Therefore, the user will have the option of choosing the registration method to be used but the default will be set to the LP registration. Our results also show that 3- and 4-mm-thick slices are suitable for registration.

#### B. Image-Guided Seed Placement Tests

The following tests assessed robots ability to place seeds at random locations in a gel model. The gel used for these tests was purposely prepared such that it is firmer than the regular tissue. This prevented the gel to deflect during needle insertion. The goal of these tests was to test the robot in an ideal setup which is relevant for this development stage. For this the following procedure was used.

- 1) The robot was placed on the MR table together with an agar model as shown in Fig. 6(a).
- 2) After an initial scout image with a coarse resolution, the position of the markers was registered with a 3D gradient echo acquisition with the following parameters: repetition time TR/echo time TE = 7.7/2.3 ms; field of view FOV = 100 mm; matrix = 192 × 256; 60 slices with a slice thickness ST = 3 mm; acquired/reconstructed voxel size 0.52 × 0.52 × 3 mm<sup>3</sup>/0.39 × 0.39 × 3 mm<sup>3</sup>; acquisition bandwidth BW = 687 Hz/pixel; scan duration dur = 90 s. The robot and the gel were scanned and the registration transformation was computed using the LP algorithm described in Section II-C. Fig. 6(b) shows the image of the marker in an axial MR slice.
- 3) Random target positions were chosen in the image. These were uniformly distributed and limited to the workspace of the robot.
- 4) Seeds were deployed at the selected targets. Fig. 6(c) presents the agar model with ceramic seeds deployed.
- 5) A verification scan was acquired. The localization of the ceramic seeds in the gel was determined with a three dimensional gradient echo acquisition with the following acquisition parameters: TR/TE = 8.0/2.3 ms; FOV = 180 mm; matrix = 512 × 512; 80 slices with a ST = 0.5 mm; acquired/reconstructed voxel size 0.35 × 0.44 × 0.5 mm<sup>3</sup>/0.18 × 0.18 × 0.5 mm<sup>3</sup>;

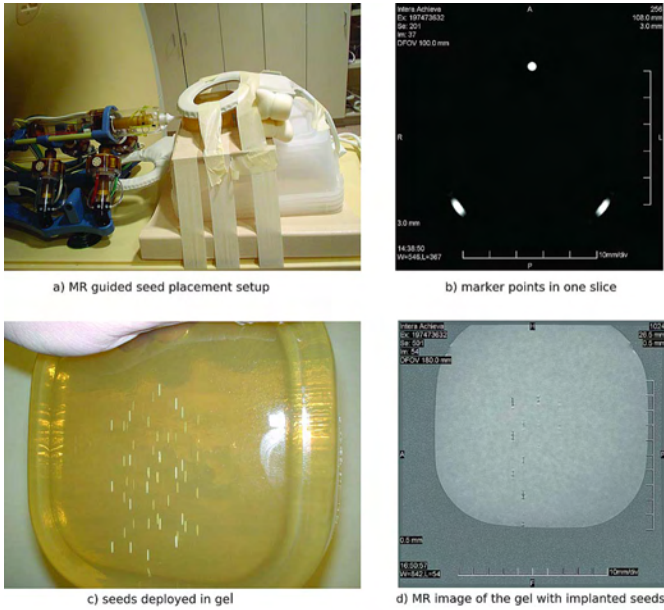


Fig. 6. Seed placement setup.

BW = 145 Hz/pixel; dur = 336 s. Fig. 6(d) illustrates one seed localization slice. In this slice, one can notice that the image artifacts are minimal allowing for accurate seed localization. Custom made ceramic seeds were used for this purpose.

6) The seeds positions from the control scan were compared against the desired seeds locations.

The system was able to deploy all seeds without incidents. This shows that the automatic seed delivery system was properly designed and built.

Fig. 7 presents the placement error computed over sixty seeds. The tests showed that the mean seed placement error is less than 1.2 mm. This value cumulates all system errors including imaging, registration, robot positioning, and seed deployment errors. Table I presents the mean values and standard deviations for each component of seed placement error. Table I indicates also that the main error component is in the direction of the needle. This is likely to be caused by seed migration along the needle tract, phenomenon common in brachytherapy procedures. Without this component, the XY positioning error is 0.6 mm mean with 0.2-mm standard deviation.

The accuracy computed in these tests can be considered the *intrinsic system accuracy*, meaning that the system, under ideal conditions, can place seeds within 1.2 mm from the specified target. In living tissues, this accuracy will likely worsen due to soft tissue deflection and seed migration. We conjecture that fast needle insertion will overcome some of the tissue deflection problems. However, the system presented in this paper has an excellent *intrinsic accuracy*. This justifies the continuation of testing with animal models.

#### IV. CONCLUSION

The paper presents a robotic system for prostate brachytherapy under MRI guidance. Even though the system is entirely constructed of nonmetallic components and does not use classic actuation methods, it is fully actuated and

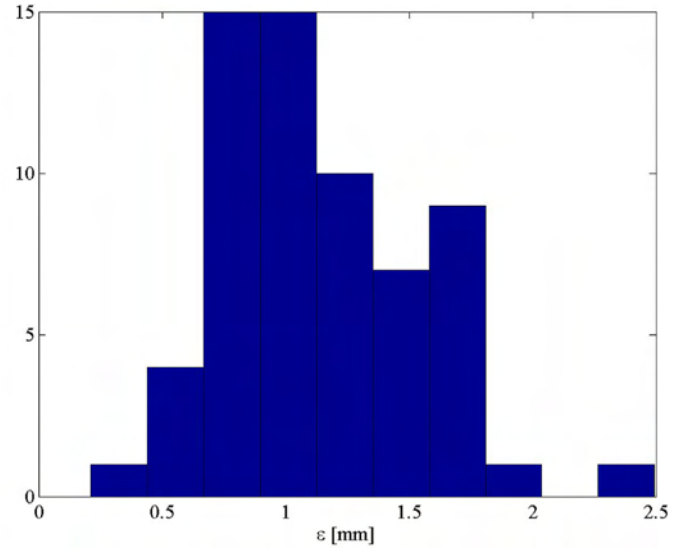


Fig. 7. Seed error placement distribution (gel tests).

TABLE I  
SEED PLACEMENT ACCURACY

	Spatial Components			Overall Error [mm]
	X[mm]	Y[mm]	Z[mm]	
Mean	0.42	0.17	0.79	1.14
Standard Deviation	0.30	0.38	0.64	0.41

allows for automated needle manipulation, insertion, and seed deployment. The position of the robot in MR coordinates is computed using a special registration marker and algorithm. The registration algorithm provides sub-millimeter registration accuracy. The mean seed placement accuracy in agar models was approximately 1.2 mm with a 0.4-mm standard deviation. The system shows remarkable accuracy and has the potential to improve the precision of radioactive seeds implantation. Currently, the system is evaluated in animal studies.

#### APPENDIX LINE ELLIPSE REGISTRATION ALGORITHM

The line is parameterized as

$$\mathbf{p} = \mathbf{v}_1 + t\mathbf{d}_1; t \in \mathbb{R}.$$

Parameters  $\mathbf{v}_1$  and  $\mathbf{d}_1$  are computed from  $\mathcal{P}_L$  using a classic line fitting method. Then, the registration is fixed up to a rotation about the line and a translation along the line. These two unknowns are solved by fitting the points  $\mathcal{P}_E$  over the ellipse. The ellipse points are parameterized as

$$\mathbf{u}(\theta) = \mathbf{v}_1 + \delta\mathbf{d}_1 + \mathbf{R}(\mathbf{e}_3, \mathbf{d}_1)\mathbf{R}_z(\alpha)(\mathbf{c}_e + \mathbf{u}_r(\theta))$$

where

- $\theta \in [0, 2\pi]$  is the ellipse parameter;
- $\delta$  is the translation along the direction  $\mathbf{d}_1$  such that  $\mathbf{v}_1 + \delta\mathbf{d}_1 = \mathbf{v}_1^i$ ;
- $\mathbf{R}(v_1, v_2) \in SO(3); \|\mathbf{v}_1\|_2 = \|\mathbf{v}_2\|_2 = 1$  is the rotation matrix that transform  $\mathbf{v}_1$  in  $\mathbf{v}_2$ ;
- $\mathbf{e}_3 = (0 \ 0 \ 1)^T$ ;

- $\mathbf{R}_z(\alpha) \in SO(3)$  is the matrix that implements a rotation about the z-axis with the angle  $\alpha$ ;
- $\mathbf{c}_e$  is the position of the ellipse center in a coordinate system parallel with RCS and centered at  $\mathbf{v}_1^r$ ;
- $\mathbf{u}^r(\theta) = D/2 \begin{pmatrix} \cos(\theta) \\ \sin(\theta) \\ \cos(\theta) \end{pmatrix}$ .

The ellipse is the result of the intersection between a plane and a cylinder of diameter  $D$ . The cylinder axis is the robot z-axis and the plane makes a  $45^\circ$  angle with the robot z-axis.

With the previous notations the rotation matrix from RCS to ICS is  $\mathbf{R}_r^i = \mathbf{R}(\mathbf{e}_3, \mathbf{d}_1) \mathbf{R}_z(\alpha)$ . Assuming that there are available initial approximations  $\delta_0$  and  $\alpha_0$  for  $\delta$ , respectively,  $\alpha$ , the following algorithm can be used to compute the registration transformation.

Step 1) For each point  $\mathbf{p}_j \in \mathcal{P}_E$  find the closest point on the ellipse. The vector from an ellipse point to  $\mathbf{p}_j$  is

$$\mathbf{C}_j(\theta) = \mathbf{v}_1 + \delta_0 \mathbf{d}_1 + \mathbf{R}_0(\mathbf{c}_e + \mathbf{u}^r(\theta)) - \mathbf{p}_j$$

where  $\mathbf{R}_0 = \mathbf{R}(\mathbf{e}_3, \mathbf{d}_1) \mathbf{R}_z(\alpha)$ . Then, the closest ellipse point to  $\mathbf{p}_j$  is  $\mathbf{u}(\theta_j)$  with

$$\theta_j = \arg \min_{\theta \in [0, 2\pi]} \|\mathbf{C}_j(\theta)\|_2^2.$$

Step 2) If  $\delta = \delta_0 + \delta_\varepsilon$ ,  $\alpha = \alpha_0 + \alpha_\varepsilon$  and  $\alpha_\varepsilon$  and  $\delta_\varepsilon$  are small then,

$$\mathbf{R}_z(\alpha) = \mathbf{R}_z(\alpha_0) (\mathbf{I}_3 + \text{Skew}(\mathbf{e}_3) \alpha_\varepsilon)$$

where  $\text{Skew}(\mathbf{v}); \mathbf{v} \in \mathbb{R}^3$  is the skew symmetric matrix associated with the vector  $\mathbf{v}$

$$\text{Skew}(\mathbf{v}) = \begin{pmatrix} 0 & -v_3 & v_2 \\ v_3 & 0 & -v_1 \\ -v_2 & v_1 & 0 \end{pmatrix}; \mathbf{v} = \begin{pmatrix} v_1 \\ v_2 \\ v_3 \end{pmatrix}.$$

Therefore, the following equation holds:

$$\mathbf{p}_j \approx \mathbf{v}_1 + (\delta_0 + \delta_\varepsilon) \mathbf{d}_1 + \mathbf{R}(\mathbf{e}_3, \mathbf{d}_1) \mathbf{R}_z(\alpha_0) (\mathbf{I}_3 + \text{Skew}(\mathbf{e}_3) \alpha_\varepsilon) (\mathbf{c}_e + \mathbf{u}_r(\theta_j)).$$

This equation can be rearranged in

$$\mathbf{p}_j - \mathbf{v}_1 - \delta_0 \mathbf{d}_1 - \mathbf{R}_0(\mathbf{c}_e + \mathbf{u}^r(\theta_j)) \approx \delta_\varepsilon \mathbf{d}_1 + \mathbf{R}_0 \text{Skew}(\mathbf{e}_3) (\mathbf{c}_e + \mathbf{u}^r(\theta_j)) \alpha_\varepsilon. \quad (1)$$

Equation (1) can be stacked in an over constrained linear system with 2 unknowns and  $3n_E$  equations

$$\begin{pmatrix} \dots & \dots \\ \mathbf{d}_1 & \mathbf{a}_j \\ \dots & \dots \end{pmatrix} \begin{pmatrix} \delta_\varepsilon \\ \alpha_\varepsilon \end{pmatrix} = \begin{pmatrix} \dots \\ \mathbf{b}_j \\ \dots \end{pmatrix} \quad (2)$$

where

$$\begin{aligned} \mathbf{a}_j &= \mathbf{R}_0 \text{Skew}(\mathbf{e}_3) (\mathbf{c}_e + \mathbf{u}^r(\theta_j)) \\ \mathbf{b}_j &= \mathbf{p}_j - \mathbf{v}_1 - \delta_0 \mathbf{d}_1 - \mathbf{R}_0 (\mathbf{c}_e + \mathbf{u}^r(\theta_j)). \end{aligned}$$

The system 2 is solved in a least-squares sense for  $\delta_\varepsilon$  and  $\alpha_\varepsilon$ .

Step 3) The registration transformation is updated using the values computed at Step 2)

$$\begin{aligned} \delta_0 &= \delta_0 + \delta_\varepsilon \\ \mathbf{R}_0 &= \mathbf{R}_0 (\mathbf{I}_3 + \text{Skew}(\mathbf{e}_3) \alpha_\varepsilon). \end{aligned}$$

If  $|\delta_\varepsilon| + |\alpha_\varepsilon| > \text{MinUpd}$  then go back to Step 1); otherwise, follow to Step 4).

Step 4) The registration transformation is

$$\begin{aligned} \mathbf{R}_r^i &= \mathbf{R}_0 \\ \mathbf{T}^i &= \mathbf{v}_1 + \delta_0 \mathbf{d}_1 - \mathbf{R}_r^i \mathbf{v}_1^r. \end{aligned}$$

## REFERENCES

- [1] A. Jemal, R. Siegel, E. Ward, T. Murray, J. Xu, C. Smigal, and M. J. Thun, "Cancer statistics, 2006," *CA Cancer J. Clin.*, vol. 56, pp. 106–130, 2006.
- [2] J. Naitoh, R. L. Zeiner, and J. B. Dekernion, "Diagnosis and treatment of prostate cancer," *Am. Family Physician*, vol. 57, pp. 1531–+, 1998.
- [3] D. Ash, D. M. Bottomley, and B. M. Carey, "Prostate brachytherapy," *Prostate Cancer Prostatic Dis.*, vol. 1, pp. 185–188, 1998.
- [4] S. E. M. Langley and R. Laing, "Prostate brachytherapy has come of age: A review of the technique and results," *BJU Int.*, vol. 89, pp. 241–249, 2002.
- [5] F. A. Vicini, V. R. Kini, G. Edmundson, G. S. Gustafson, J. Stromberg, and A. Martinez, "A comprehensive review of prostate cancer brachytherapy: Defining an optimal technique," *Int. J. Radiation Oncol. Biol. Phys.*, vol. 44, pp. 483–491, 1999.
- [6] J. E. Sylvester, J. C. Blasko, P. D. Grimm, R. Meier, and J. A. Malmgren, "Ten-year biochemical relapse-free survival after external beam radiation and brachytherapy for localized prostate cancer: The seattle experience," *Int. J. Radiation Oncol. Biol. Phys.*, vol. 57, pp. 944–952, 2003.
- [7] R. G. Stock, J. A. Cesaretti, and N. N. Stone, "Disease-specific survival following the brachytherapy management of prostate cancer," *Int. J. Radiation Oncol. Biol. Phys.*, vol. 64, pp. 810–816, 2006.
- [8] H. Ragde, A. A. A. Elgamil, P. B. Snow, J. Brandt, A. A. Bartolucci, B. S. Nadir, and L. J. Korb, "Ten-year disease free survival after transperineal sonography-guided iodine-125 brachytherapy with or without 45-gray external beam irradiation in the treatment of patients with clinically localized, low to high Gleason grade prostate carcinoma," *Cancer*, vol. 83, pp. 989–1001, 1998.
- [9] L. Potters, C. Morgenstern, E. Calugaru, P. Fearn, A. Jassal, J. Presser, and E. Mullen, "12-year outcomes following permanent prostate brachytherapy in patients with clinically localized prostate cancer," *J. Urol.*, vol. 173, pp. 1562–1566, 2005.
- [10] K. K. Yu and H. Hricak, "Imaging prostate cancer," *Radiologic Clin. N. Am.*, vol. 38, pp. 59–85, 2000.
- [11] J. Kurhanewicz, M. G. Swanson, S. J. Nelson, and D. B. Vigneron, "Combined magnetic resonance imaging and spectroscopic imaging approach to molecular imaging of prostate cancer," *J. Magn. Reson. Imag.*, vol. 16, pp. 451–463, 2002.
- [12] S. D. Pathak, P. D. Grimm, V. Chalana, and Y. Kim, "Pubic arch detection in transrectal ultrasound-guided prostate cancer therapy," *IEEE Trans. Med. Imag.*, vol. 17, no. 5, pp. 762–771, Oct. 1998.
- [13] Z. P. Wei, G. Wan, L. Gardi, G. Mills, D. Downey, and A. Fenster, "Robot-assisted 3D-TRUS-guided prostate brachytherapy: System integration and validation," *Med. Phys.*, vol. 31, pp. 539–548, 2004.
- [14] K. Chinzai, R. Kikinis, and F. A. Jolesz, "MR compatibility of mechatronic devices: Design criteria," in *Proc. Medical Image Computing and Computer-Assisted Intervention—Miccai'99*, 1999, vol. 1679, Lecture Notes in Computer Science, pp. 1020–1030.
- [15] D. Stoianovici, "Multi-Imager compatible actuation principles in surgical robotics," *Int. J. Med. Robot. Comput. Assist. Surg.*, vol. 1, pp. 86–100, 2005.
- [16] K. Chinzai, N. Hata, F. A. Jolesz, and R. Kikinis, "MR compatible surgical assist robot: System integration and preliminary feasibility study," in *Proc. Medical Image Computing and Computer-Assisted Intervention—Miccai'2000*, 2000, vol. 1935, Lecture Notes in Computer Science, pp. 921–930.

- [17] B. T. Larson, A. G. Erdman, N. V. Tsekos, E. Yacoub, P. V. Tsekos, and L. G. Koutlas, "Design of an MRI-compatible robotic stereotactic device for minimally invasive interventions in the breast," *J. Biomech. Eng.—Trans. ASME*, vol. 126, pp. 458–465, 2004.
- [18] N. V. Tsekos, A. Ozcan, and E. Christoforou, "A prototype manipulator for magnetic resonance-guided interventions inside standard cylindrical magnetic resonance imaging scanners," *J. Biomech. Eng.—Trans. ASME*, vol. 127, pp. 972–980, 2005.
- [19] E. Hempel, H. Fischer, L. Gumb, T. Hohn, H. Krause, U. Voges, H. Breitwieser, B. Gutmann, J. Durke, M. Bock, and A. Melzer, "An MRI-compatible surgical robot for precise radiological interventions," *Comput. Aided Surg.*, vol. 8, pp. 180–191, 2003.
- [20] A. Krieger, R. C. Susil, C. Menard, J. A. Coleman, G. Fichtinger, E. Atalar, and L. L. Whitcomb, "Design of a novel MRI compatible manipulator for image-guided prostate interventions," *IEEE Trans. Biomed. Eng.*, vol. 52, no. 2, pp. 306–313, Feb. 2005.
- [21] M. Muntener, A. Patriciu, D. Petrisor, D. Mazilu, H. Bagga, L. R. Kavoussi, K. Cleary, and D. Stoianovici, "MRI compatible robotic system for fully automated brachytherapy seed placement," *Urology*, 2006, to be published.
- [22] D. Stoianovici, A. Patriciu, D. Mazilu, D. Petrisor, and L. Kavoussi, "A new type of motor: Pneumatic step motor," *IEEE/ASME Trans. Mechatronics*, vol. 12, no. 1, pp. 98–106, Feb. 2006.
- [23] M. P. R. Van Gellekom, M. A. Moerland, J. J. Battermann, and J. J. W. Lagendijk, "MRI-guided prostate brachytherapy with single needle method – A planning study," *Radiother. Oncol.*, vol. 71, pp. 327–332, 2004.
- [24] V. Lagerburg, M. A. Moerland, M. K. Konings, R. E. van de Vosse, J. J. W. Lagendijk, and J. J. Battermann, "Development of a tapping device: A new needle insertion method for prostate brachytherapy," *Phys. Med. Biol.*, vol. 51, pp. 891–902, 2006.
- [25] R. C. Susil, J. H. Anderson, and R. H. Taylor, "A single image registration method for CT-guided interventions," in *Proc. Medical Image Computing and Computer-Assisted Intervention—Miccai'99*, 1999, vol. 1679, Lecture Notes in Computer Science, pp. 798–808.
- [26] S. Lee, G. Fichtinger, and G. S. Chirikjian, "Numerical algorithms for spatial registration of line fiducials from cross-sectional images," *Med. Phys.*, vol. 29, pp. 1881–1891, 2002.
- [27] A. Krieger, G. Metzger, G. Fichtinger, E. Atalar, and L. L. Whitcomb, "A hybrid method for 6-DOF tracking of MRI-compatible robotic interventional devices," presented at the 2006 IEEE International Conf. Robotics and Automation, Orlando, FL, 2006.
- [28] L. Ibanez, W. Schroeder, L. Ng, and J. Cates, *The ITK Software Guide*. Clifton Park, NY: Kitware Inc., 2003.
- [29] J. M. Fitzpatrick, J. B. West, and C. R. Maurer, "Predicting error in rigid-body point-based registration," *IEEE Trans. Med. Imag.*, vol. 17, no. 5, pp. 694–702, Oct. 1998.



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