TRANS RECTAL ULTRASOUND GUIDED HIGH DOSE RATE INTERSTITIAL BRACHYTHERAPY FOR CARCINOMA PROSTATE

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CARCINOMA PROSTATE

- Uncommon malignancy
- Implantable cases are rare

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22 / 1000 LDR 10 years
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- 21 / 500 HDR 5 years
- Inappropriate training
- TRUS & brachytherapy under one roof
- Dedicated systems are expensive

BRACHYTHERAPY

- Accessibility -- good
- Imaging -- amenable
- Invasiveness -- minimal
- Radiation tolerance -- good
- Dose escalation -- beneficial
- Dispensability -- high
- Critical organs -- close by

INDICATIONS

- Stage T1b to T3b
- Any Gleason score
- Any PSA level
- MO

- Low risk -- monotherapy
- Intermediate / high risk as boost

CONTRAINDICATOINS

ABSOLUTE

- M1 disease
- Medically unfit for anesthesia
- Life expectancy < 5 years
- Technically not feasible to implant whole gland

RELATIVE

- Gland > 80 cc → Pubic arch interference
- TURP last six months or large TURP defect
- Obstructive urinary symptoms IPSS > 14
- Prior pelvic radiotherapy

TURP

- Mostly unnecessary
- If IPSS >20 (shows obstruction)
- Complications
- Poor survival
- Delays RT by 3-6 months.
- Excessive urinary morbidity.

BRACHYTHERAPY - TYPES

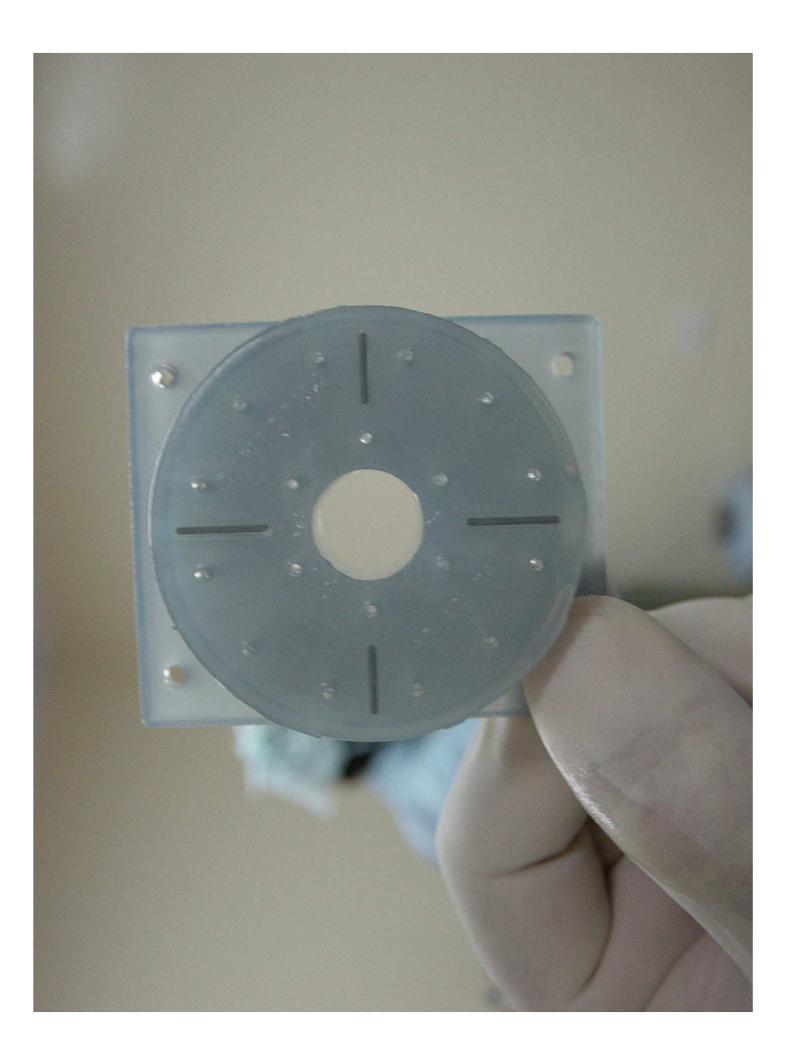
Permanent seed implants

Manually afterloaded low dose rate (LDR)

Remote afterloaded high dose rate (HDR)

SPECIAL CONSIDERATIONS

- Pre op. low residue diet
- Pre op. laxatives / enema
- Liquid diet intra op.
- Flatus tube
- Three way urinary catheter
- Epidural anaesthesia and analgesia



TRUS GUIDED BRACHYTHERAPY

Allows direct and continuous visualization of the relationship between

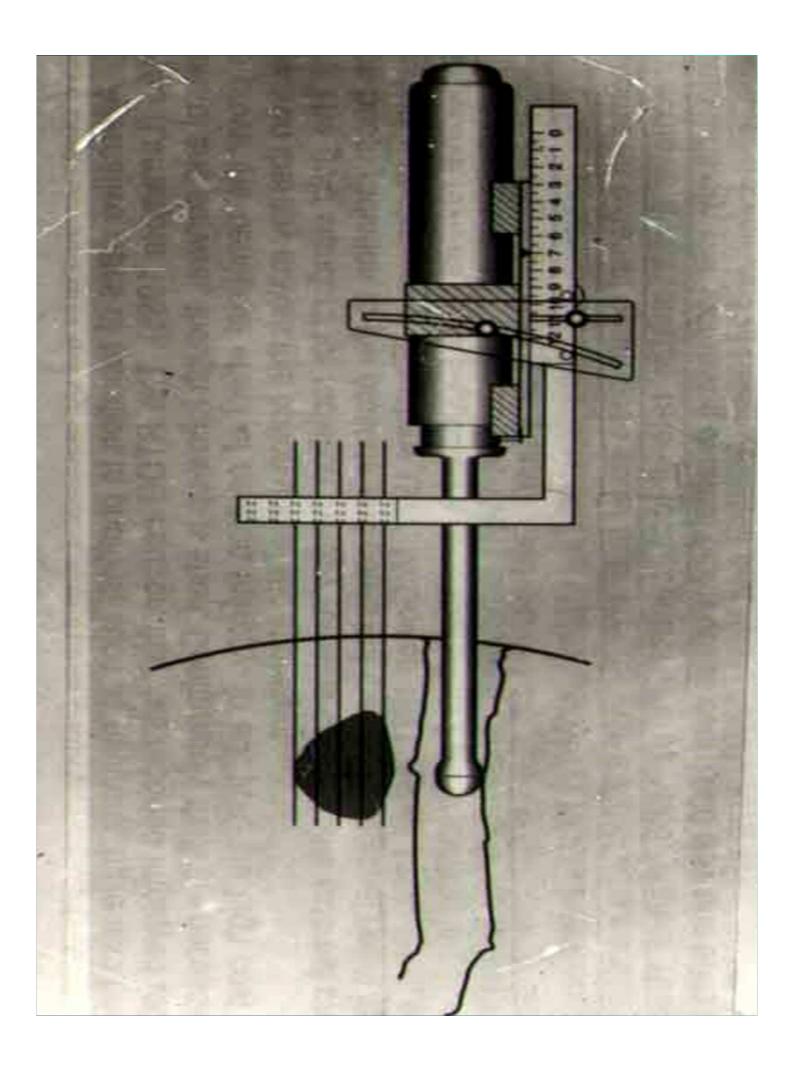
Rectal wall

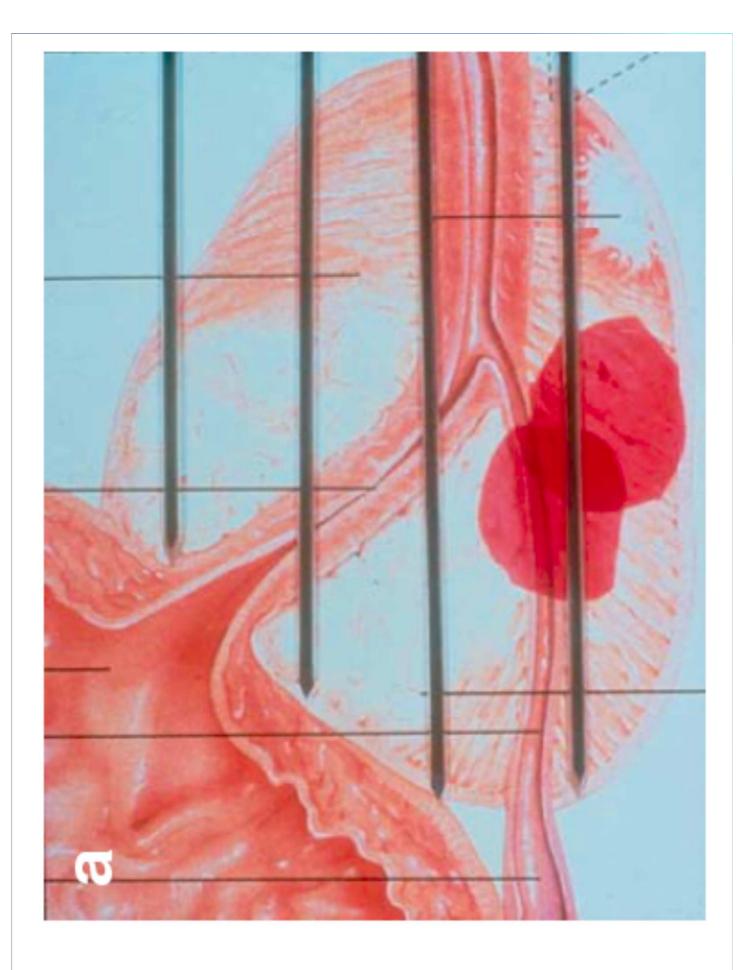
Urethra

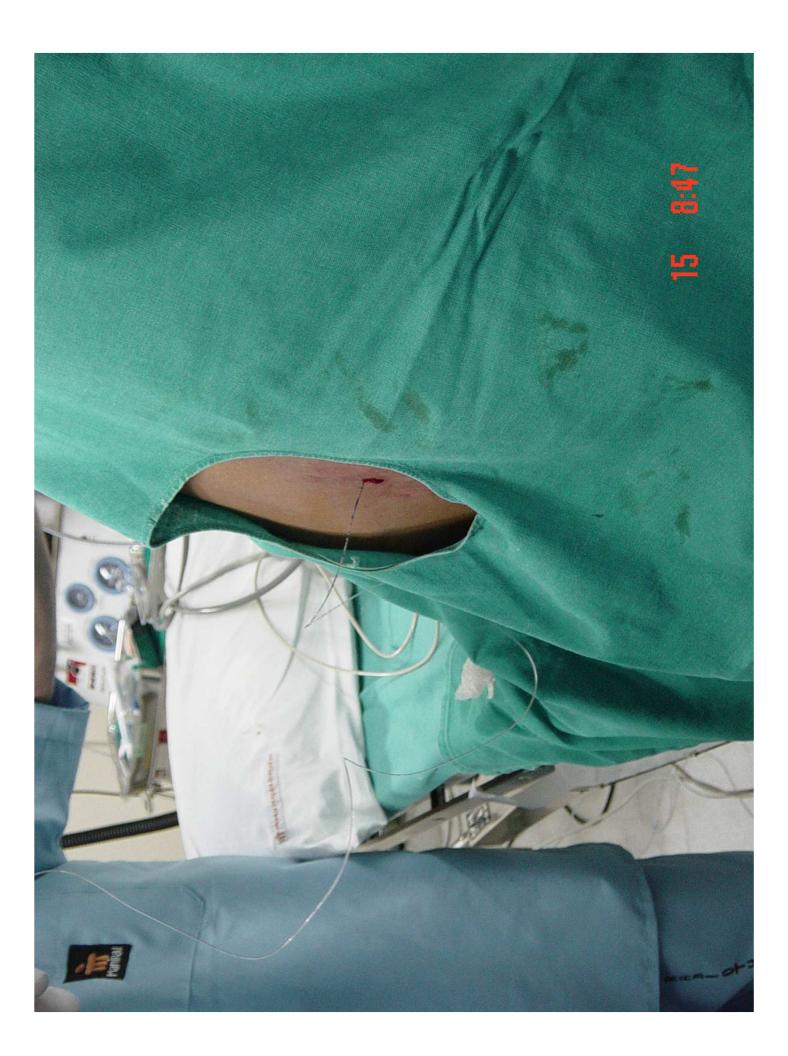
Urinary bladder

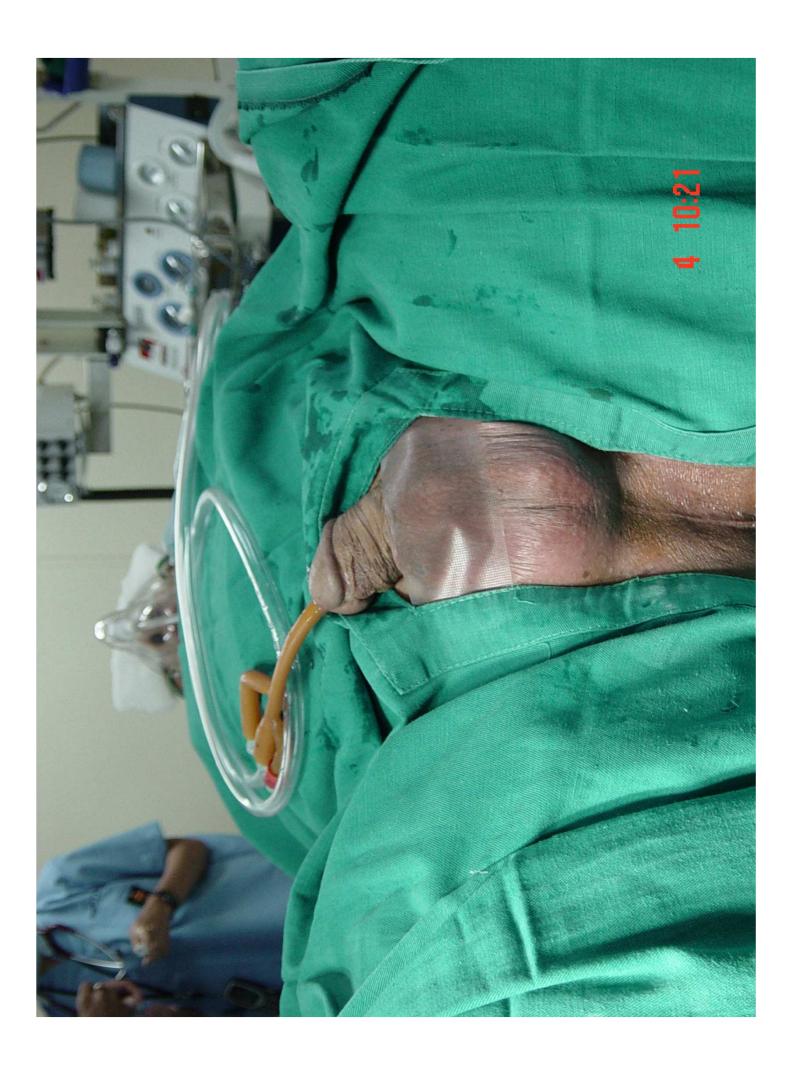
Prostate contour

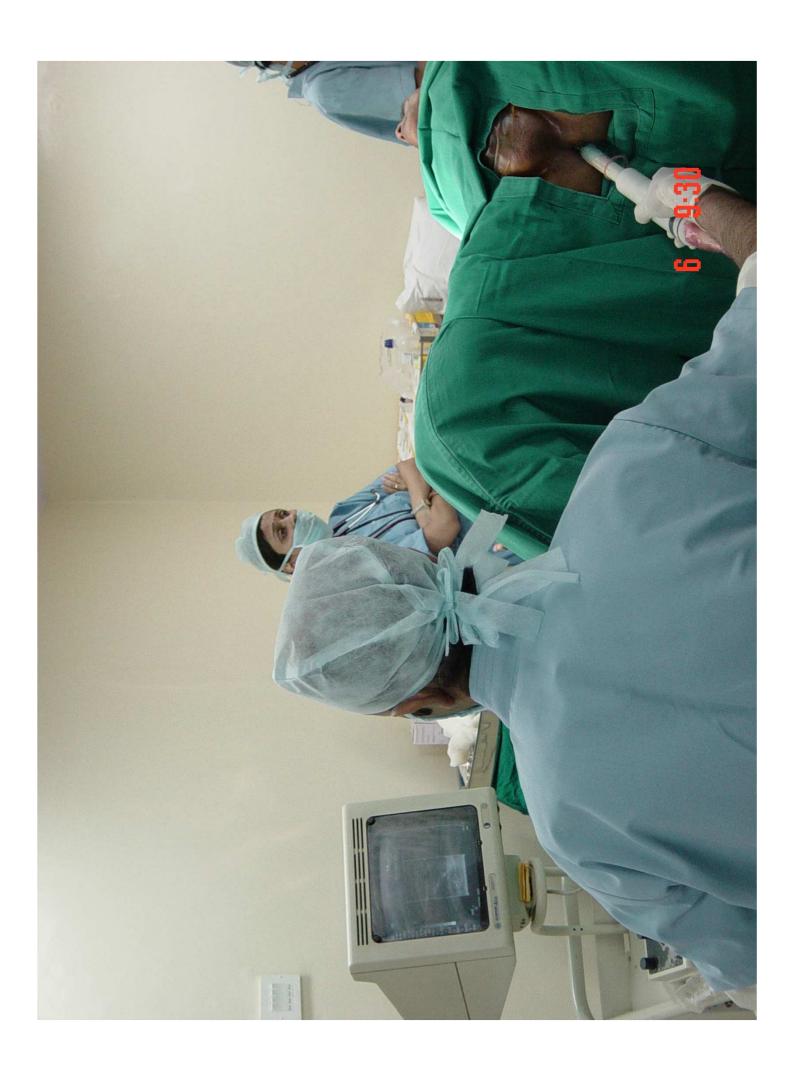
- Precise dose delivery system
- Very effective treatment

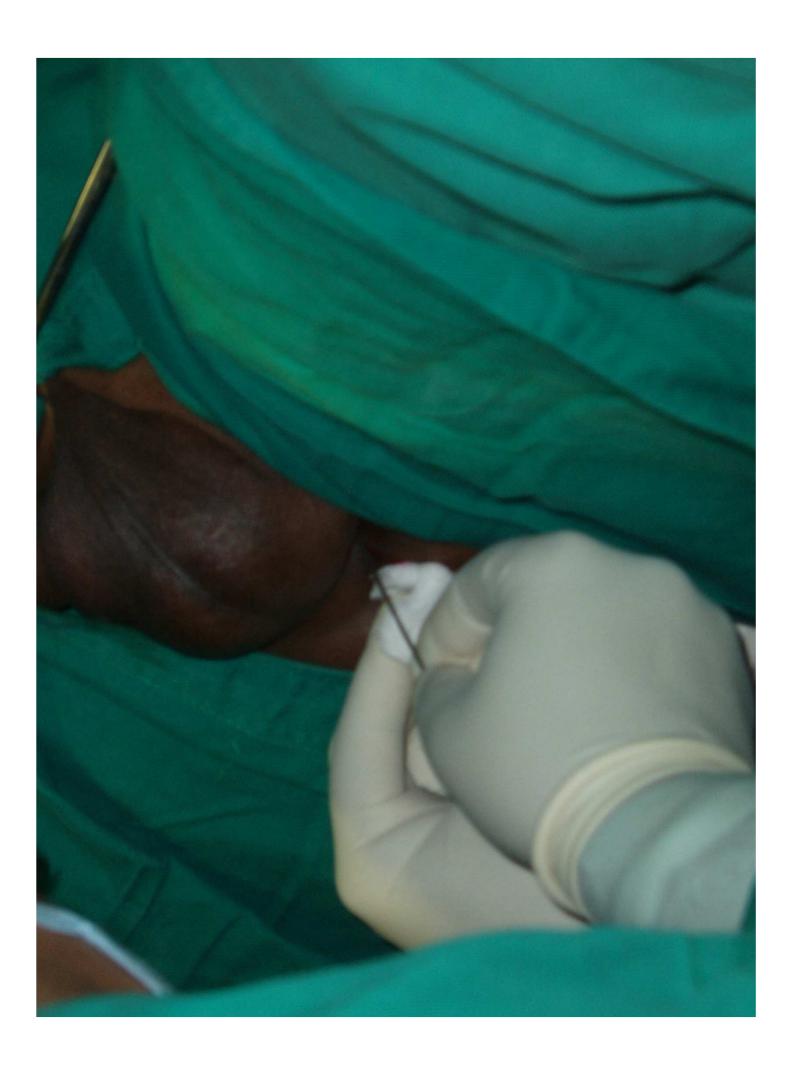


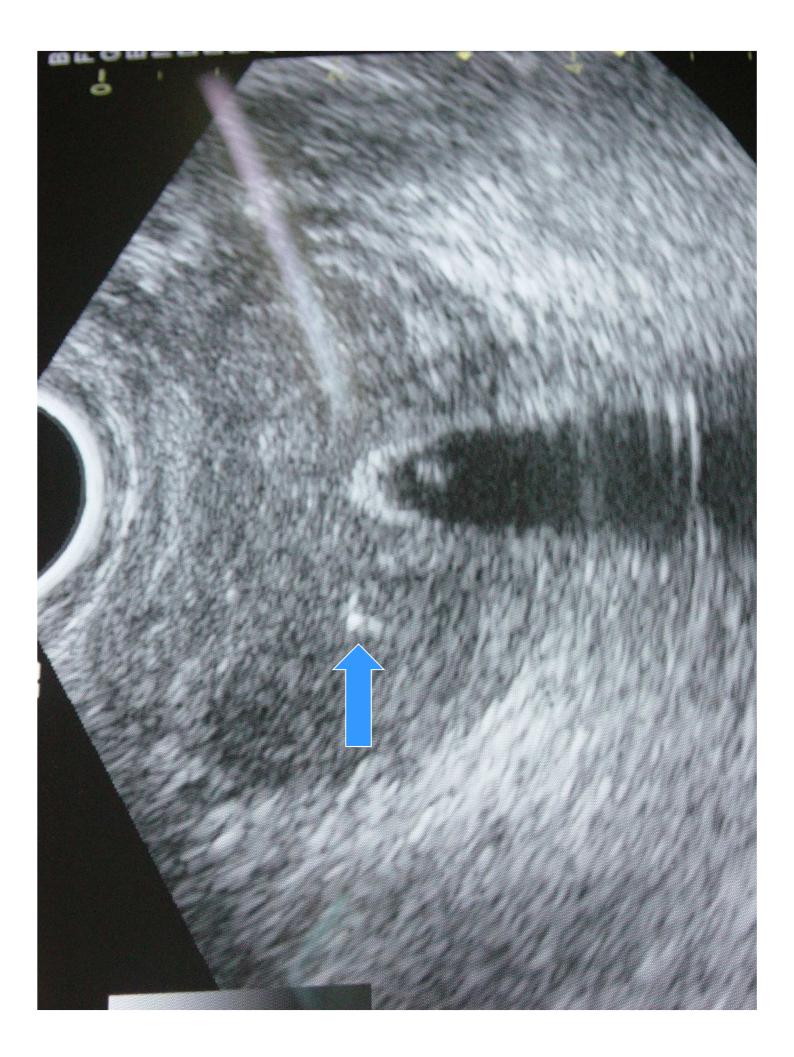




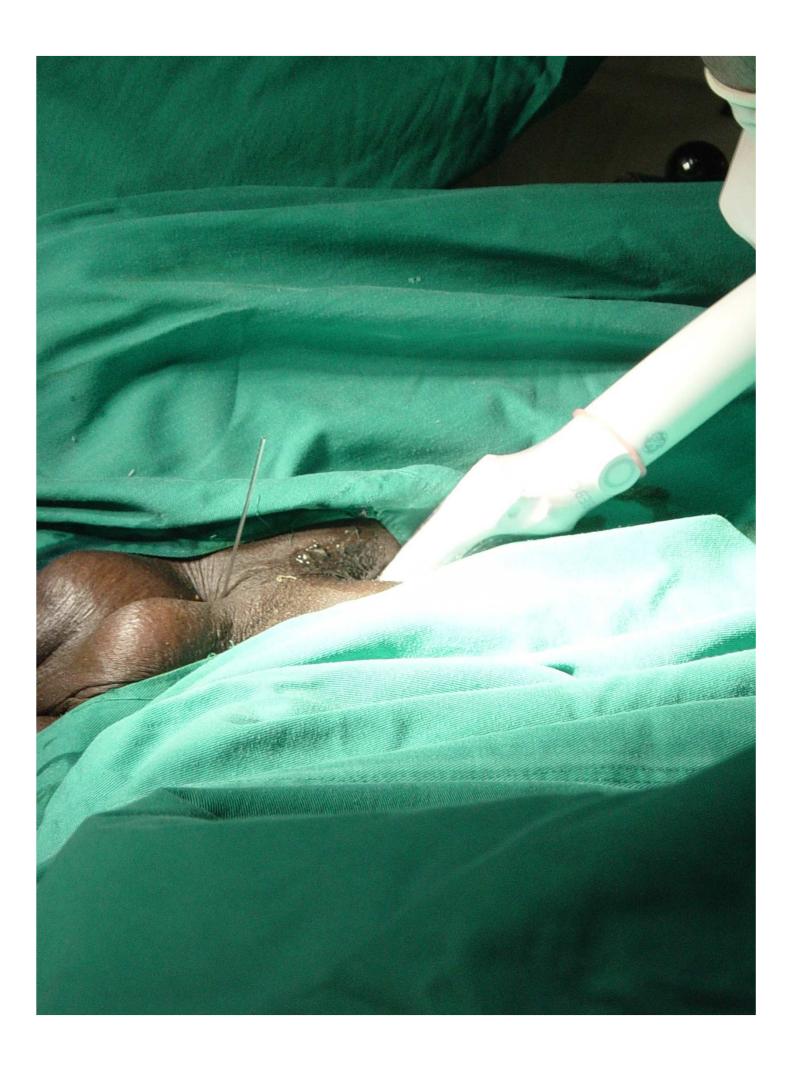


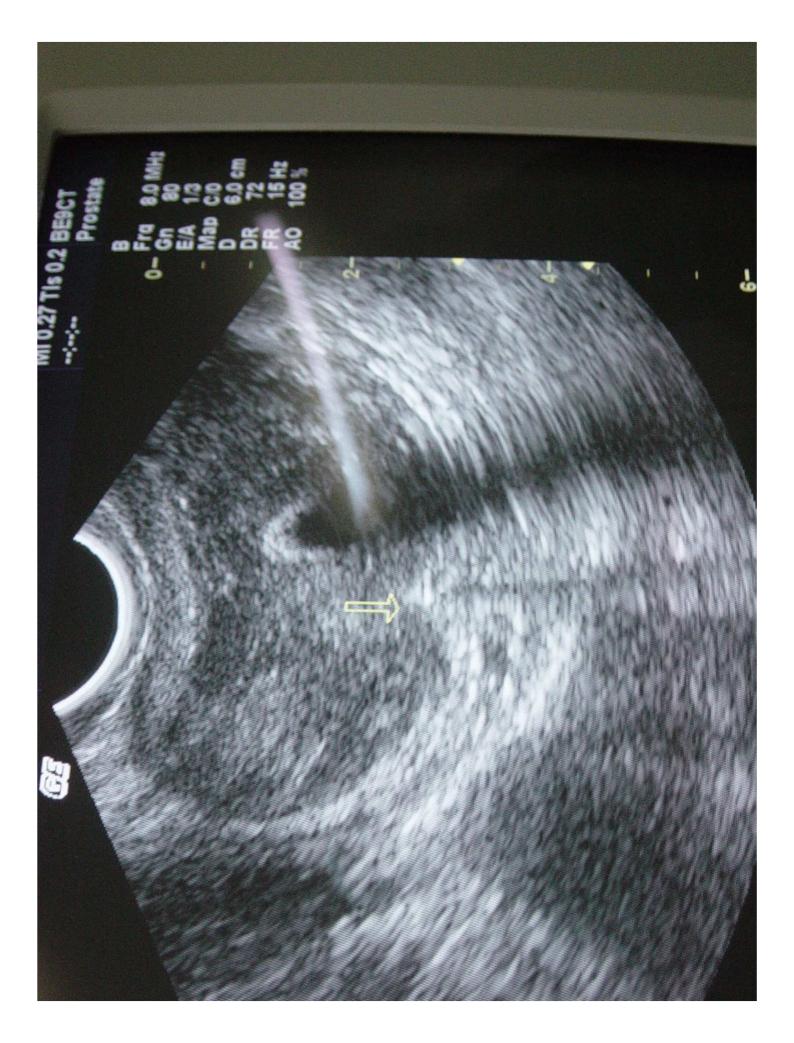


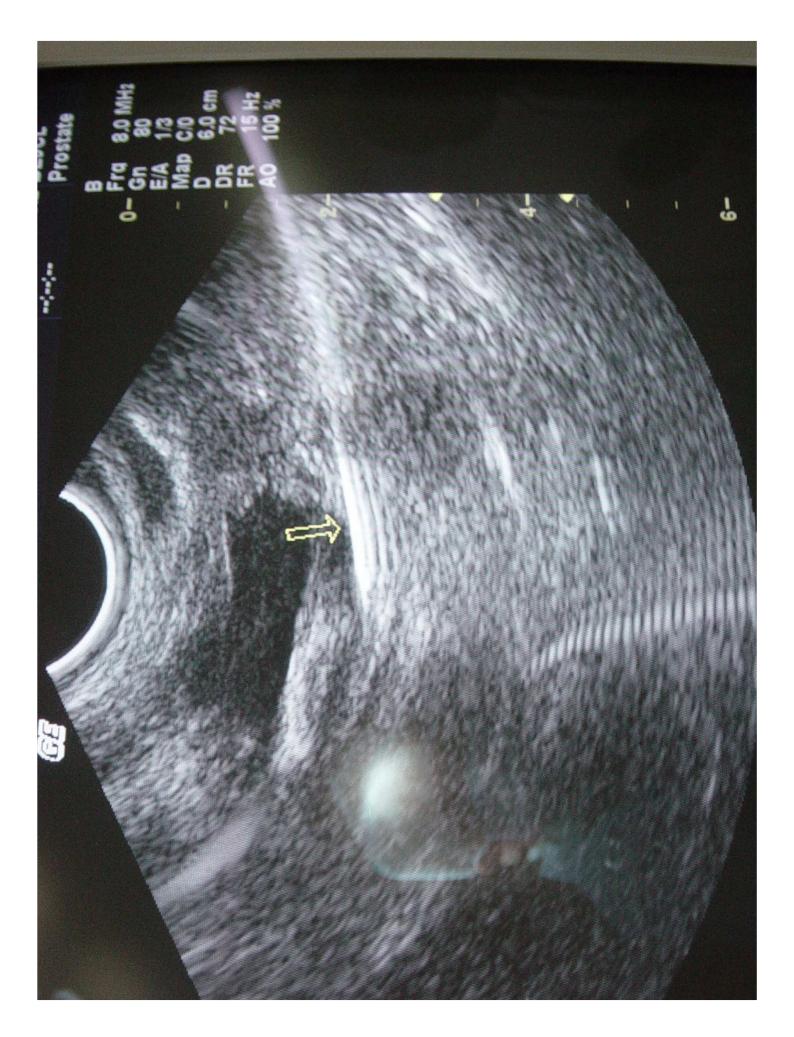


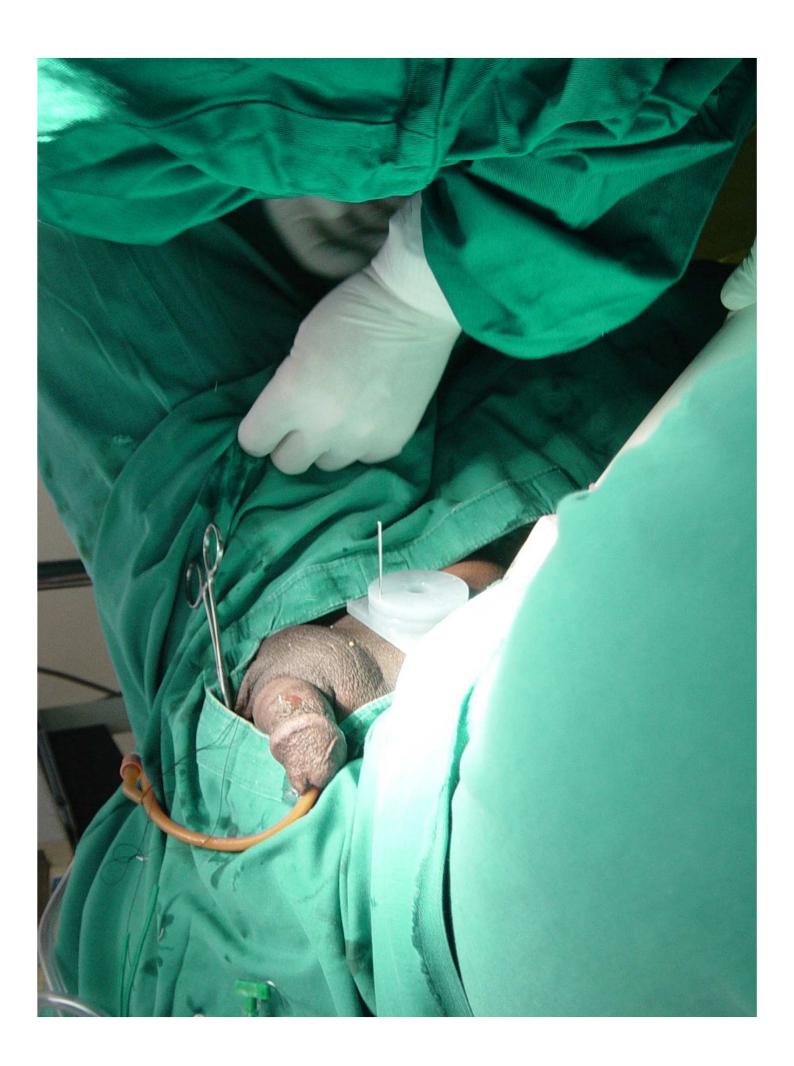


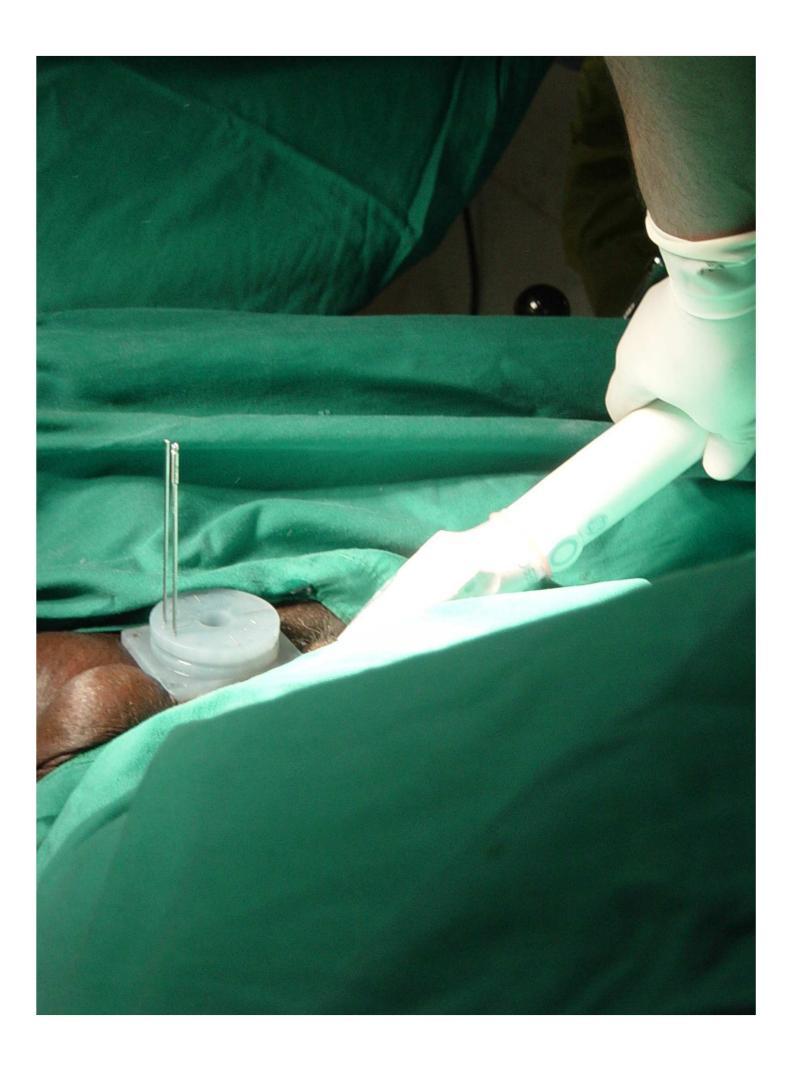








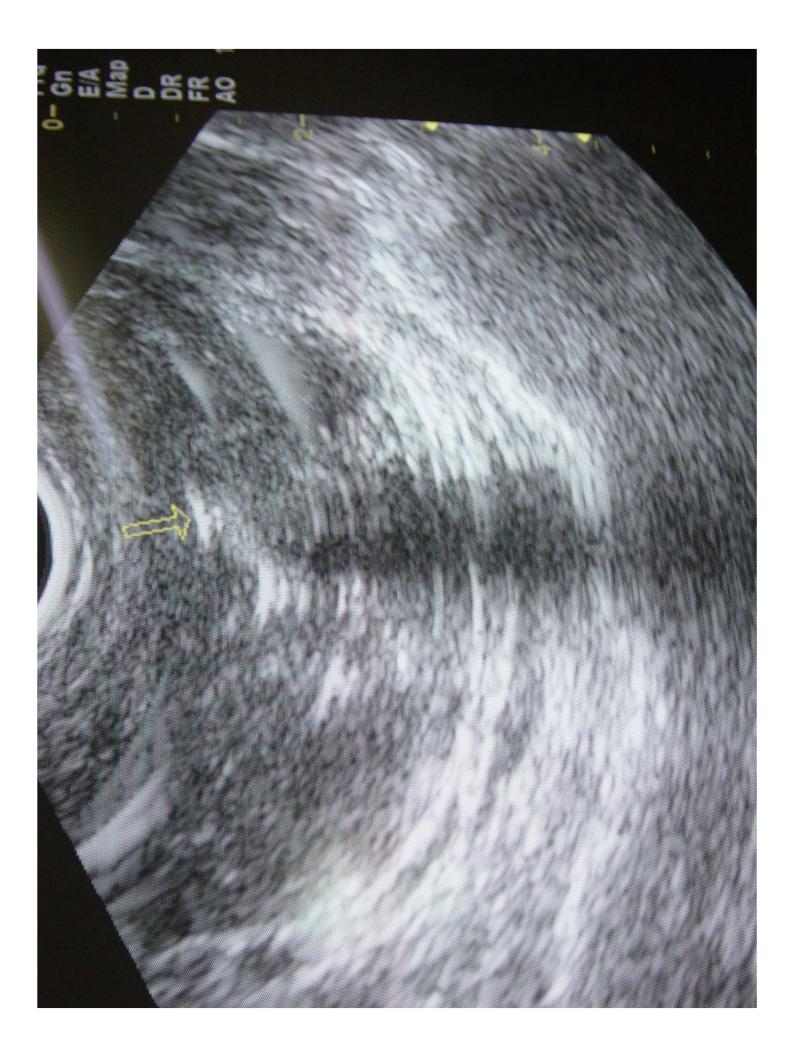








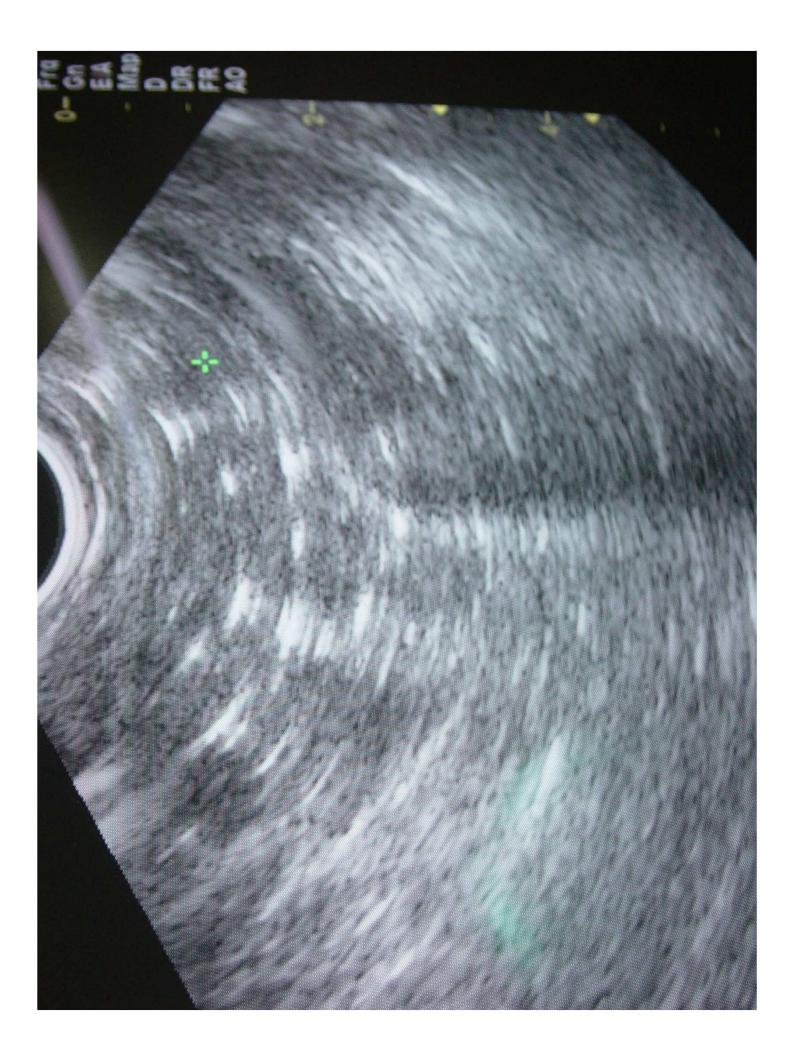


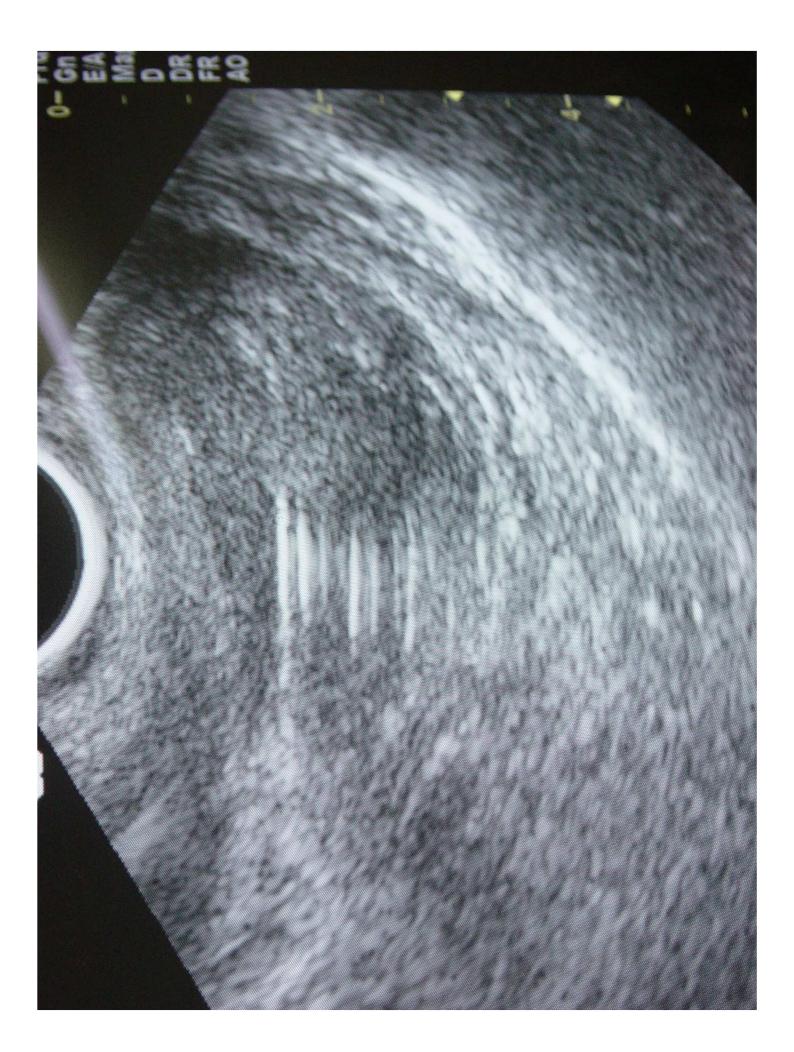












MI 0.30 TIS 0.2 BE9CT MANIPAL HOSPITAL 06 9:09:56 AM Urology

O- Fra

Bladder(0.7) Prostate Renal

eneric

Bladder

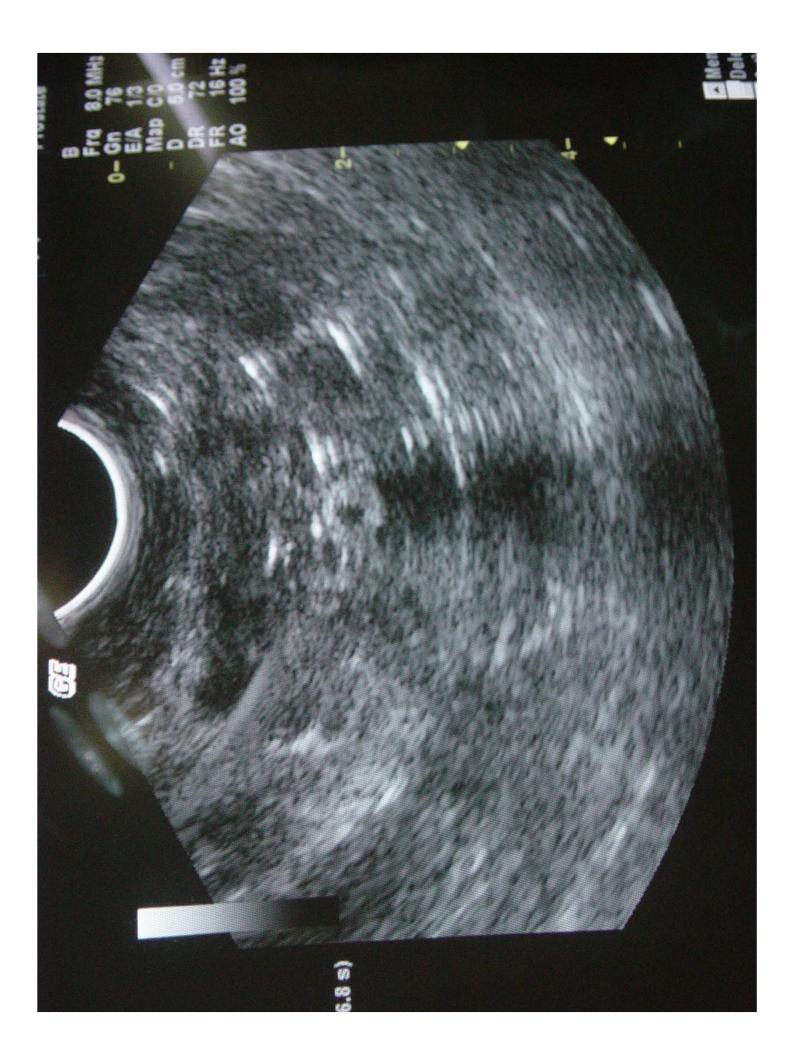
Bladder Vol(0.7) Prostate Vol

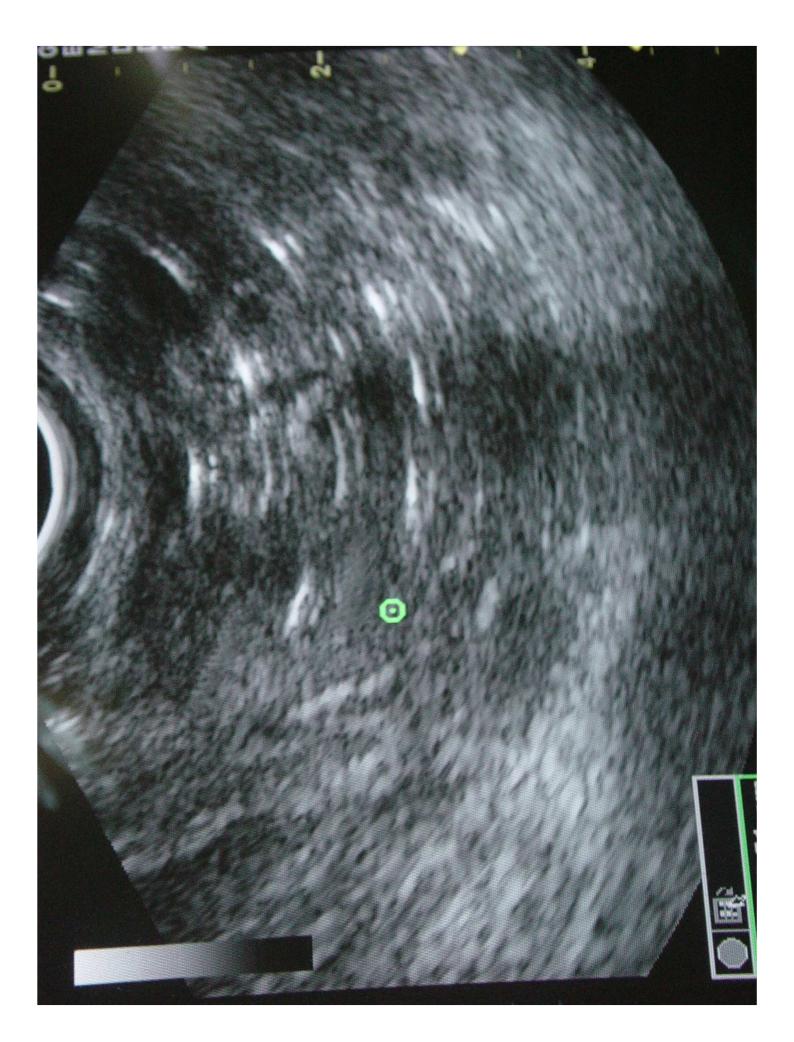
Renal Vol

Bladder Vol

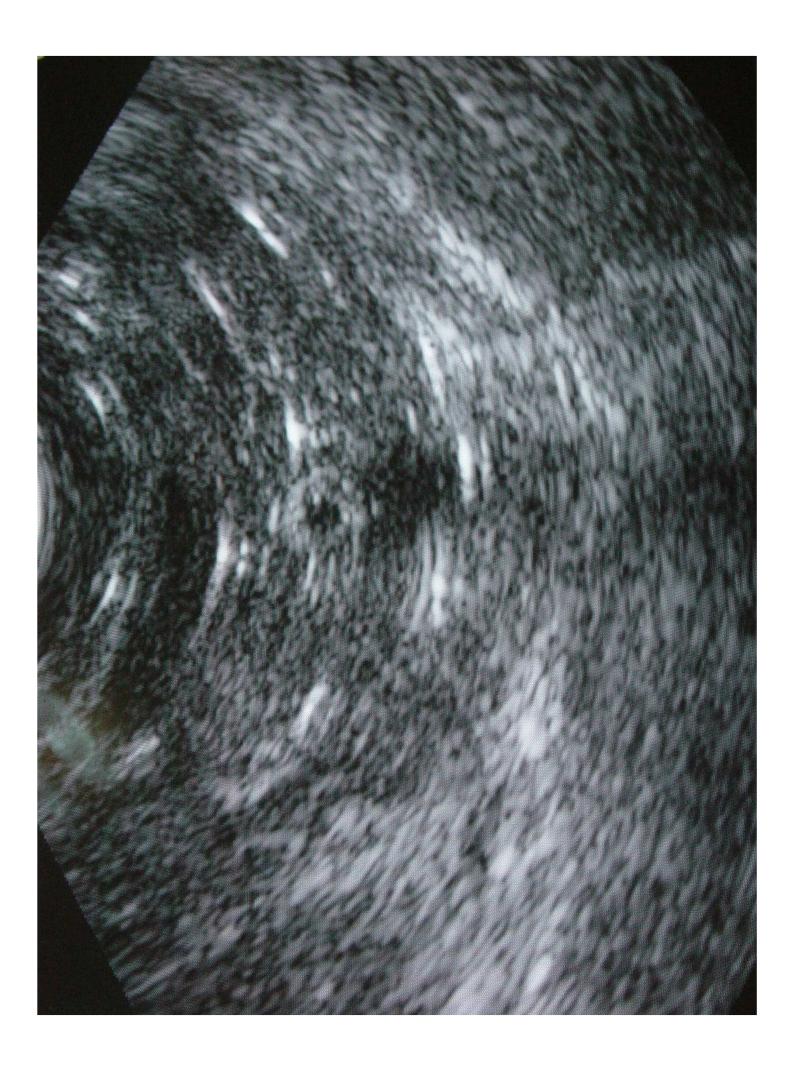
Worksheet Display

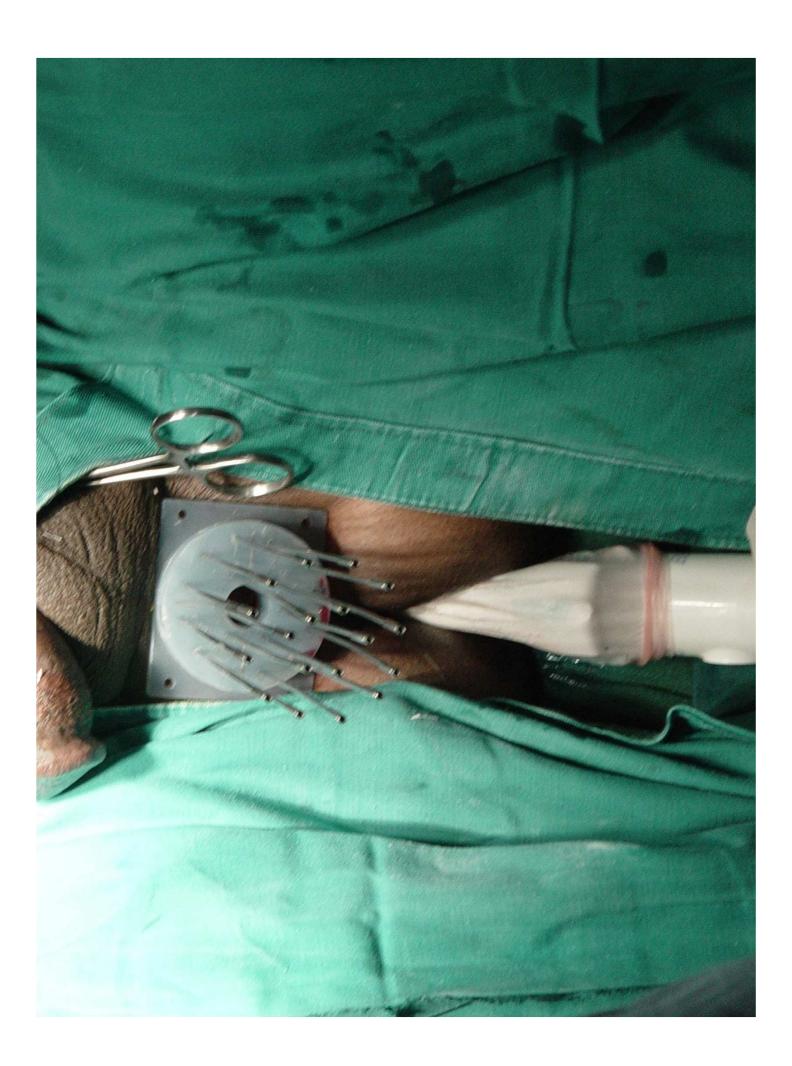


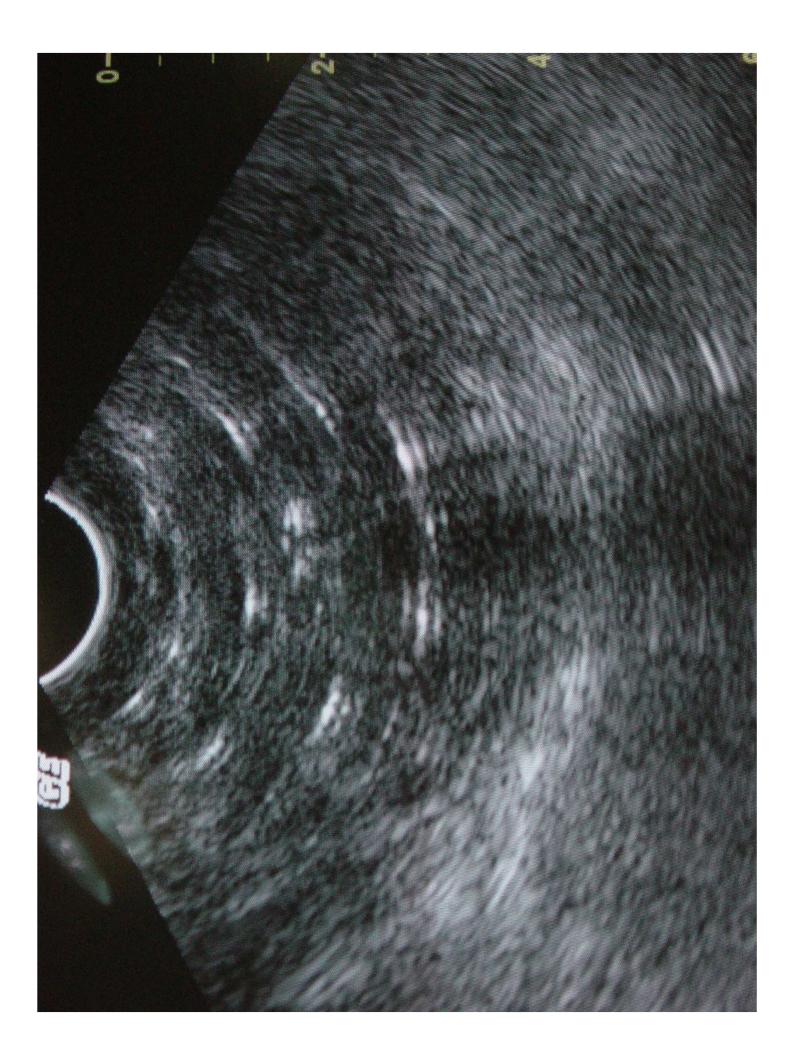




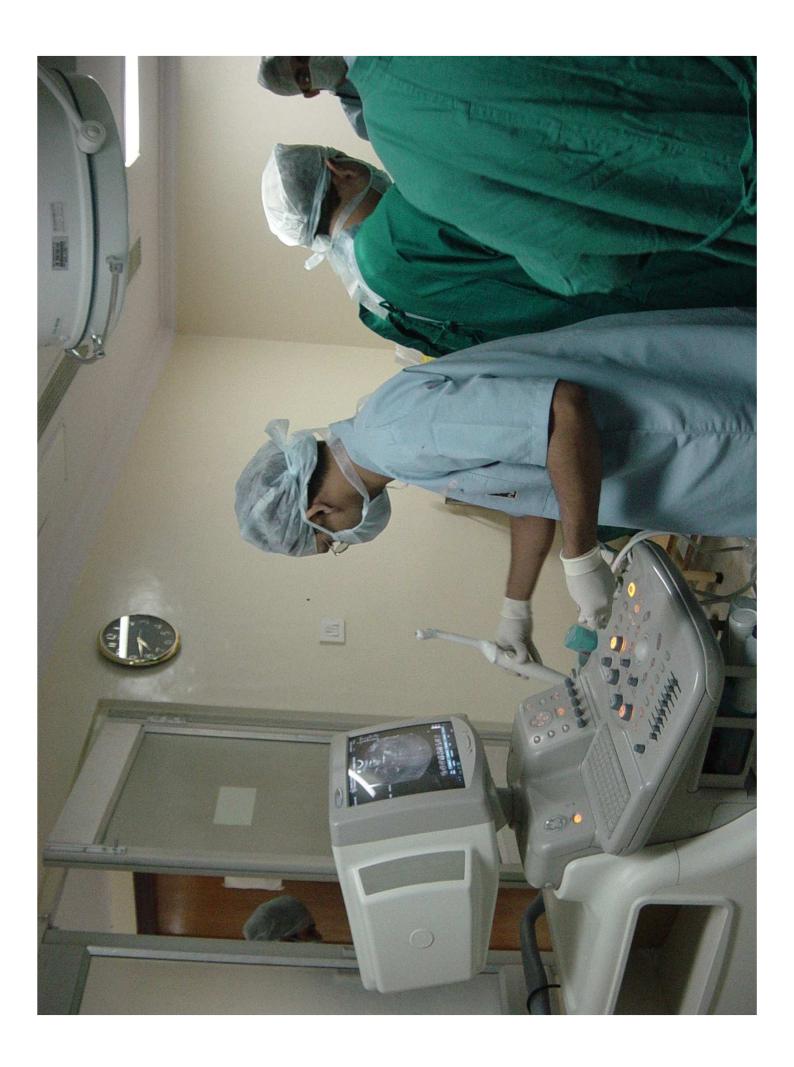


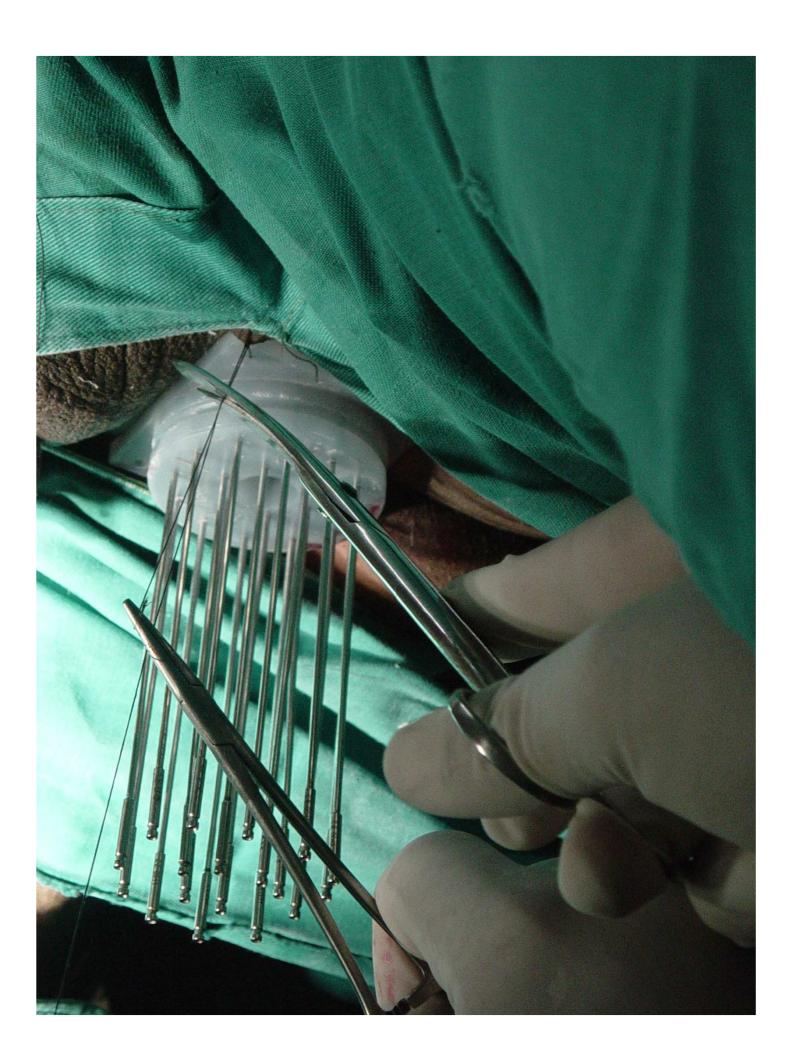


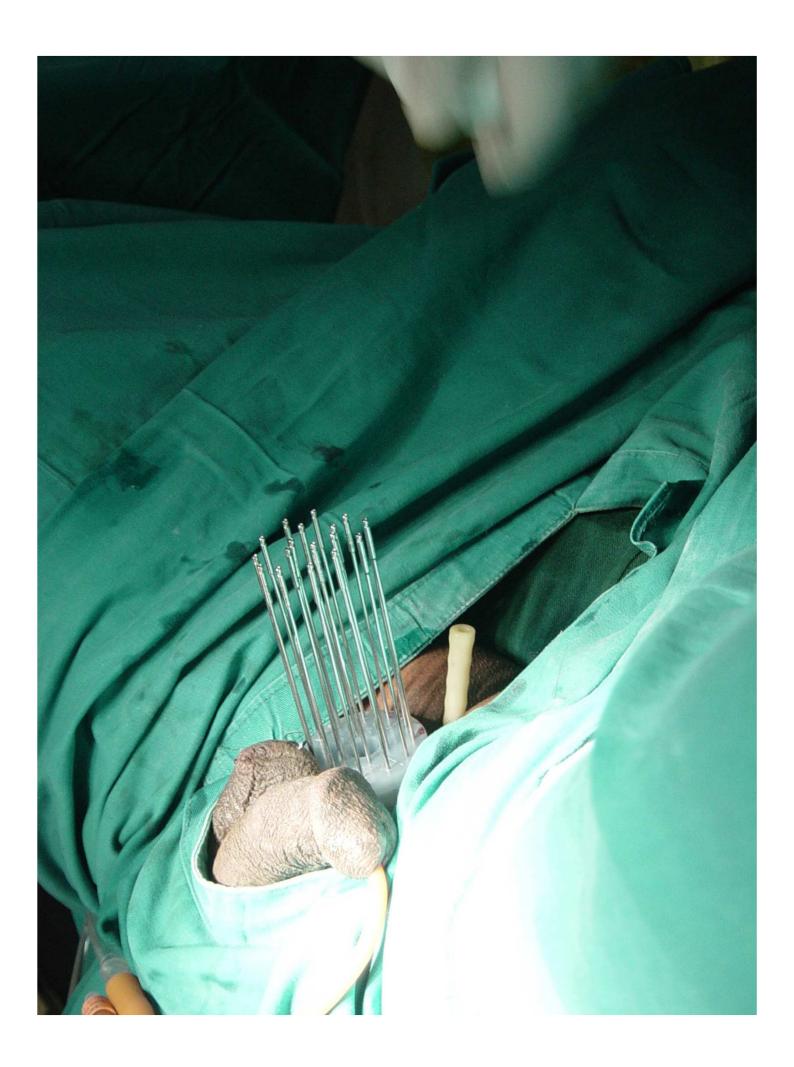


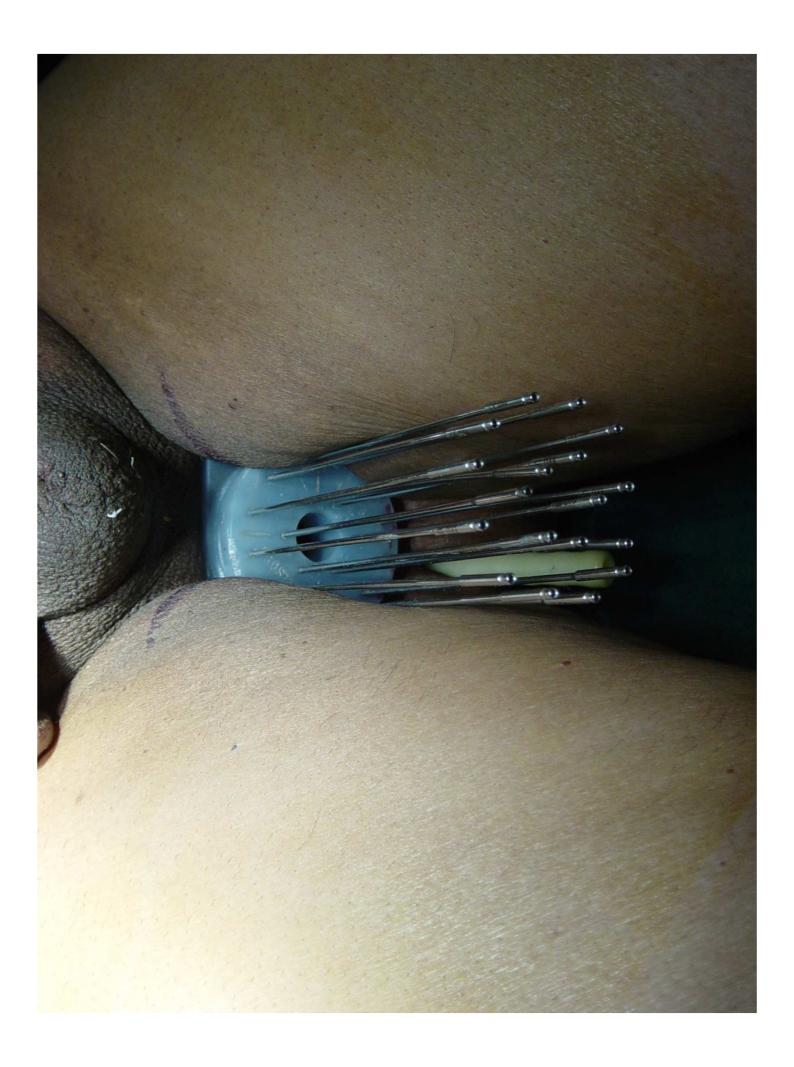


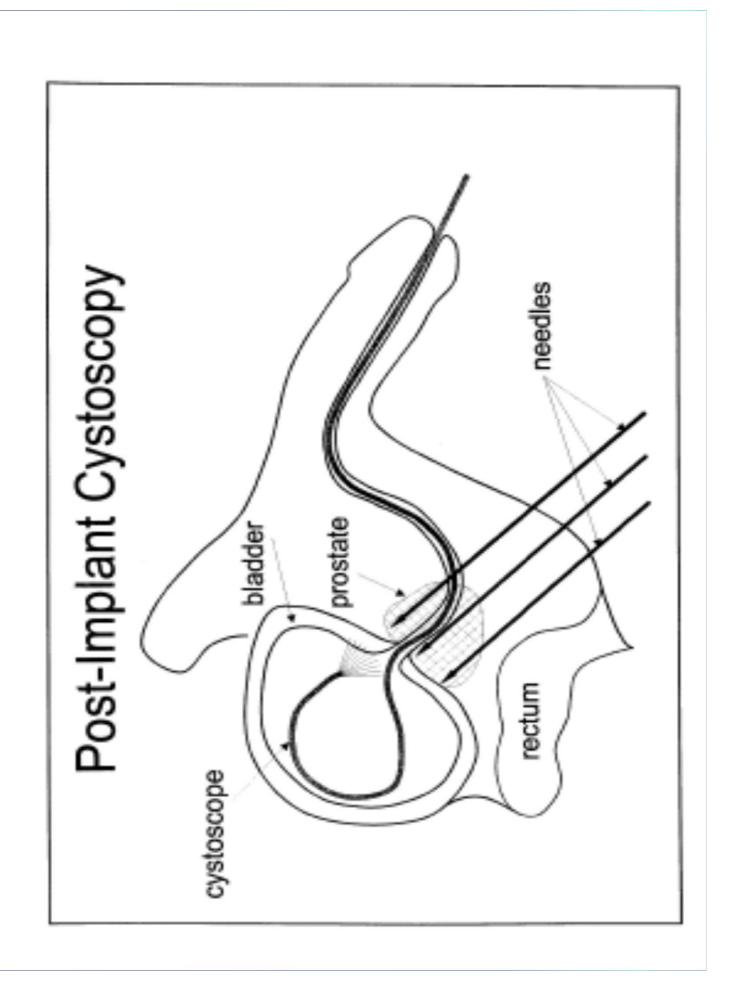


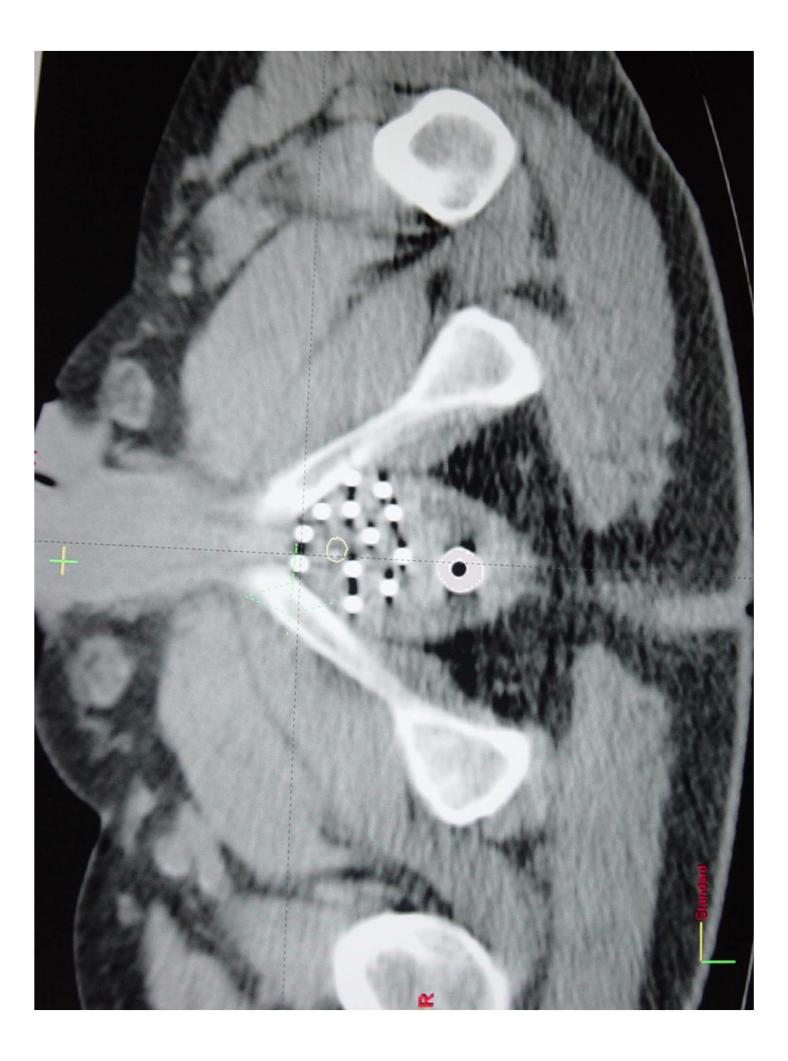








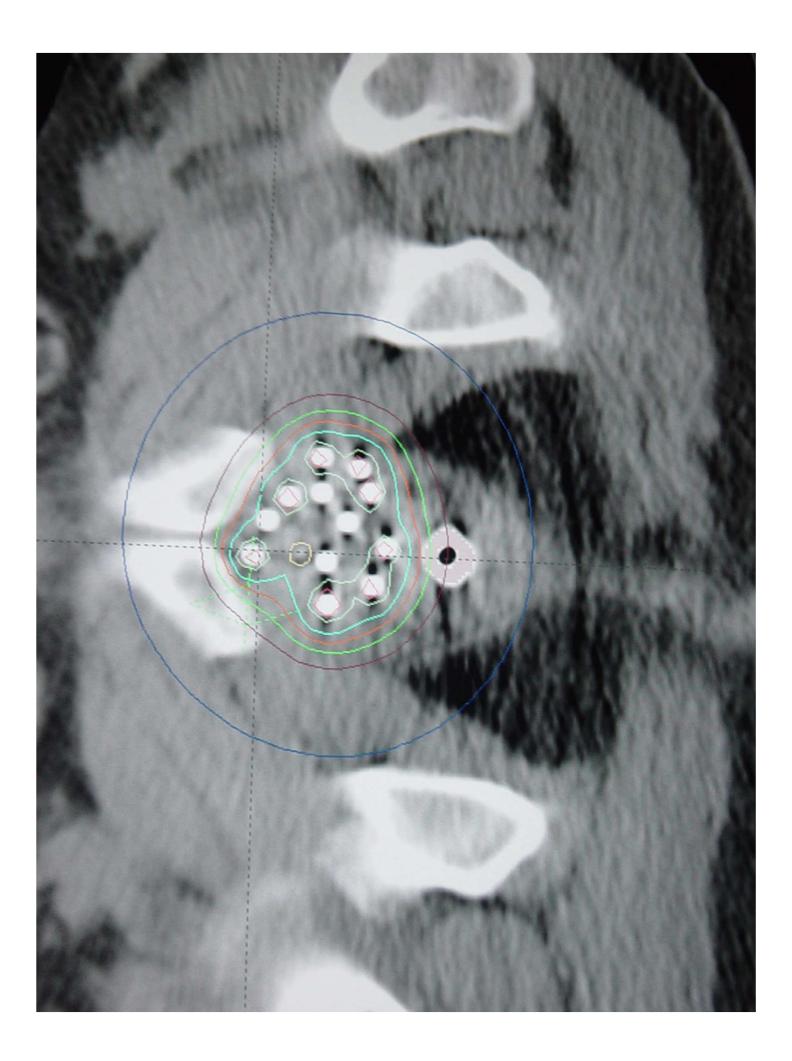






OPTIMIZATION CONSTRAINTS

- Max. urethral dose </= 125% MPD
- MPD allowed to indent few mm. anteriorly but still covered by 80% isodose
- Higher doses to posterolateral portions (anatomic rationale) 150-200%
- Rectal dose </= 75%

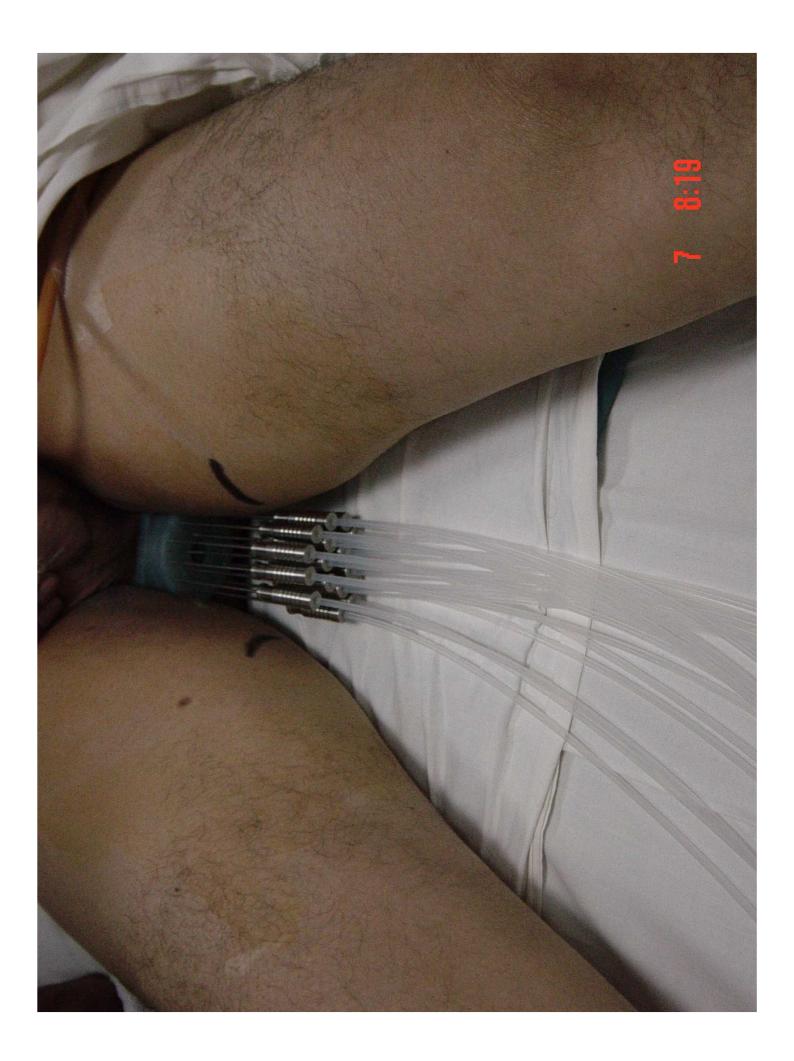












HDR EXPERIENCE

- August 2003 May 2008 22 cases
 - 1 -- Low risk
 - 10 -- Intermediate risk
 - 11 High risk
- T2b T3b
 Gl. 6- 8
 PSA .13 191
- XRT 50.4 Gy 3DCRT / 54Gy by IMRT
- ISBT minimum peripheral dose (MPD)
 - 5.5Gy x 3 -- 5
 - 5.5 Gy x 4 -- 5
 - 7.5 Gy x 2 -- 7
 - 8.5 Gy X 2 -- 4
 - 8.5 Gy x 4 -- 1
- Follow up 5 51 m. LC 100%
- One dead lung mets.

No significant morbidity

MORBIDITY

	LDR	HDR
<u>ACUTE</u>		
Haematuria	22/22	3/22
Retention	0/22	2/22
GU gr. III	2/22	0/22
<u>CHRONIC</u>		
Proctitis	1/22	0/22
Stricture	0/22	0/22

HIGH DOSE RATE BRACHYTHERAPY

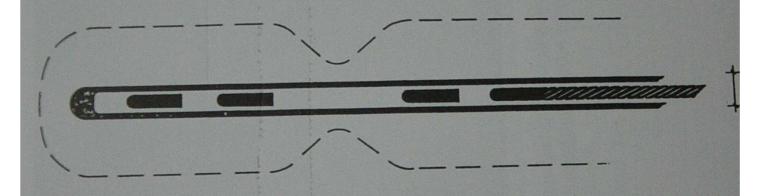
- No organ motion concerns
- Optimization allows IMRT
- High dose / fraction suits radiobiology of prostate cancer
- Short treatment time
- Minimum isolation → Better nursing care
- Minimal hospitalization → Better patient compliance
- Significantly reduced cost
- No second malignancy concerns

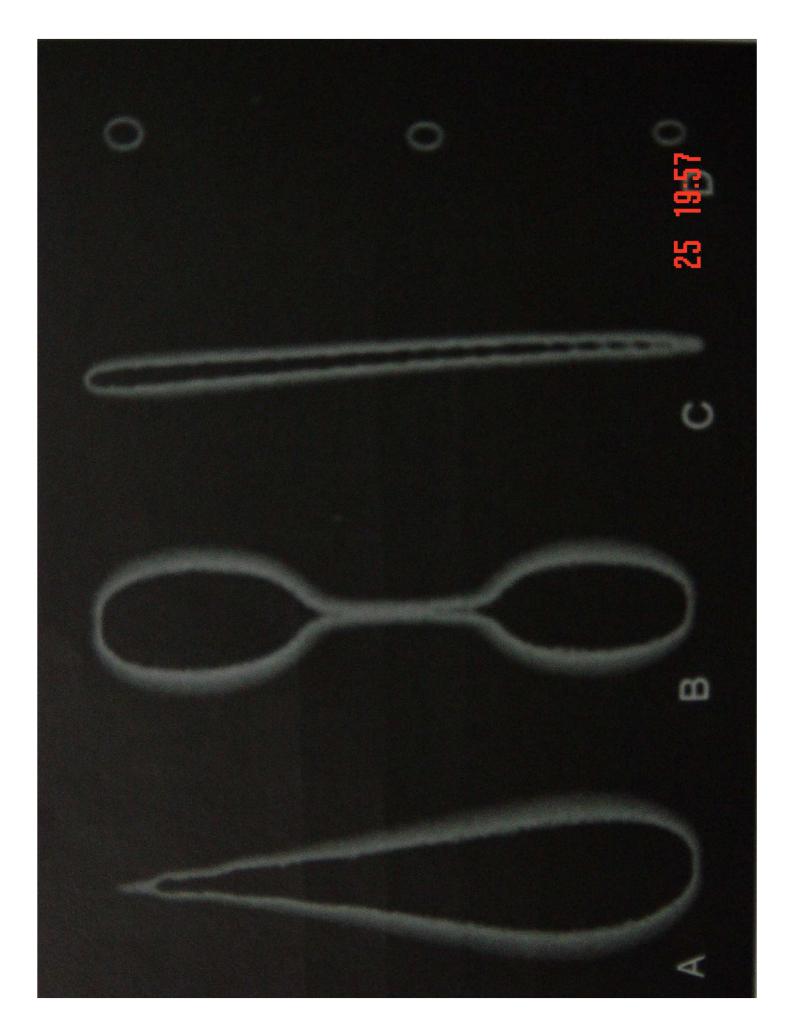
HIGH DOSE RATE BRACHYTHERAPY

 Infinite optimization possibilities due to more number of channels, dwell positions and dwell times

ALLOWS INTENSITY MODULATION WITHIN AND IMMEDIATELY AROUND PROSTATE

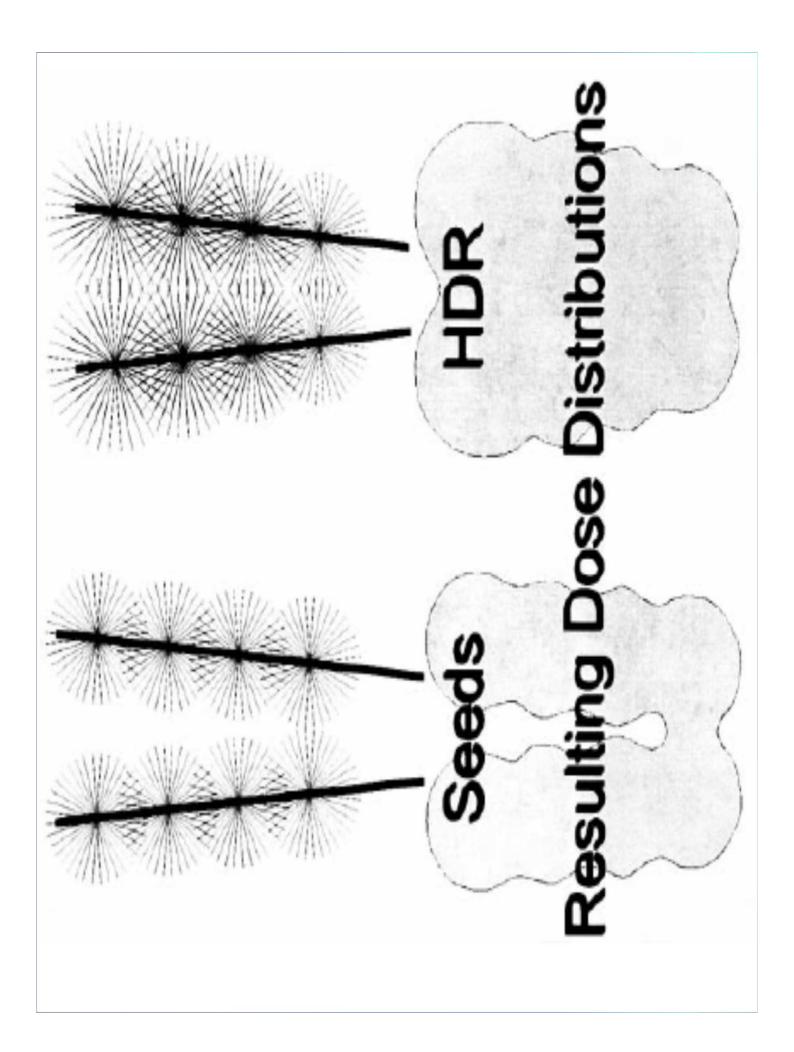
Better integration of XRT and BT may yield better cure rates

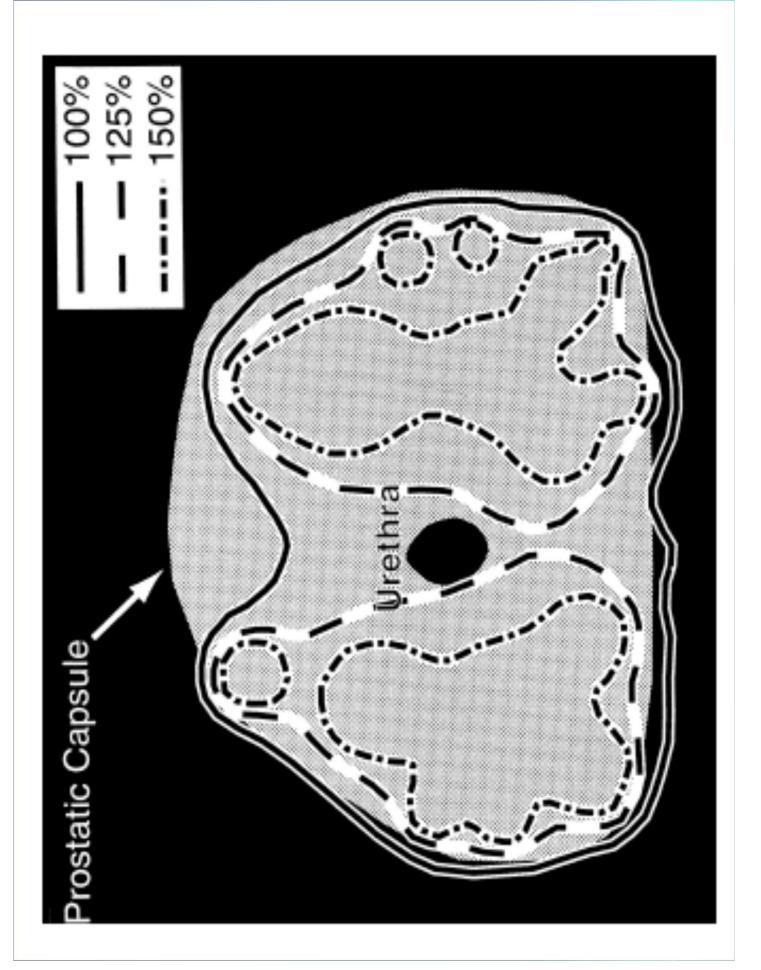




INTENSITY MODULATION USING HDR

- Correction of suboptimal needle placement makes it a forgiving type of procedure
- Possible to treat bigger size prostates with lesser needles by increasing dwell times in lateral or anterior needles
- Boost to areas of known gross disease
- Lesser dose to rectum by decreasing dwell times in posterior needles

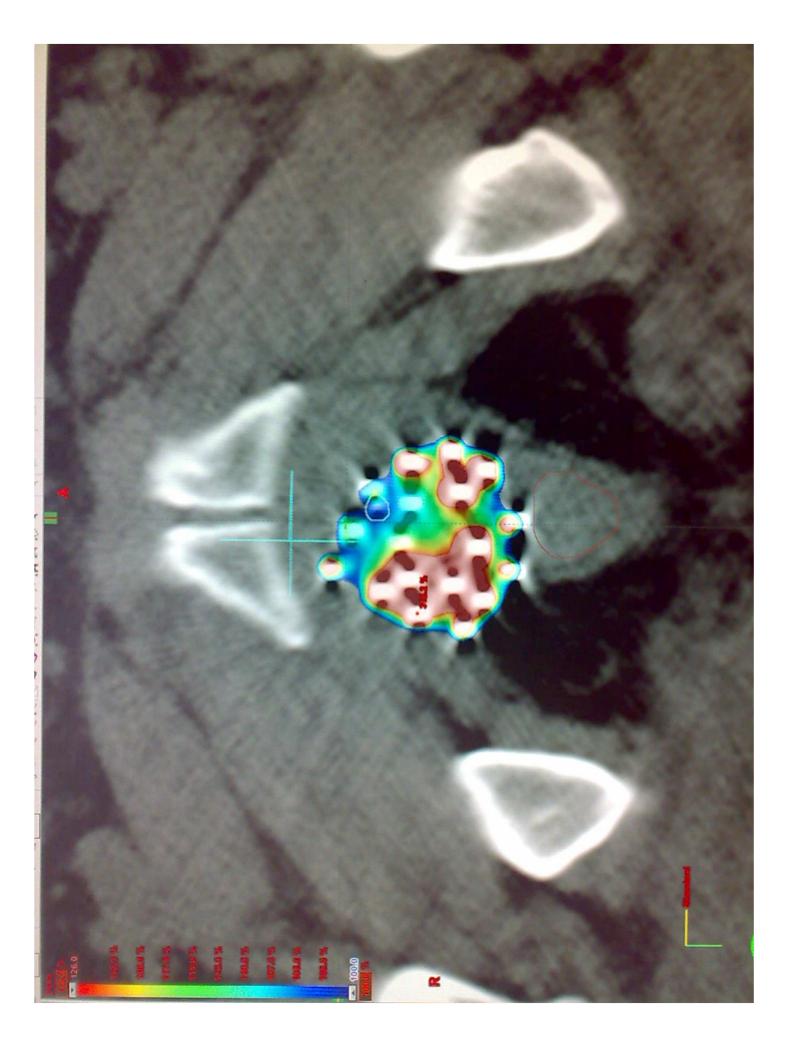




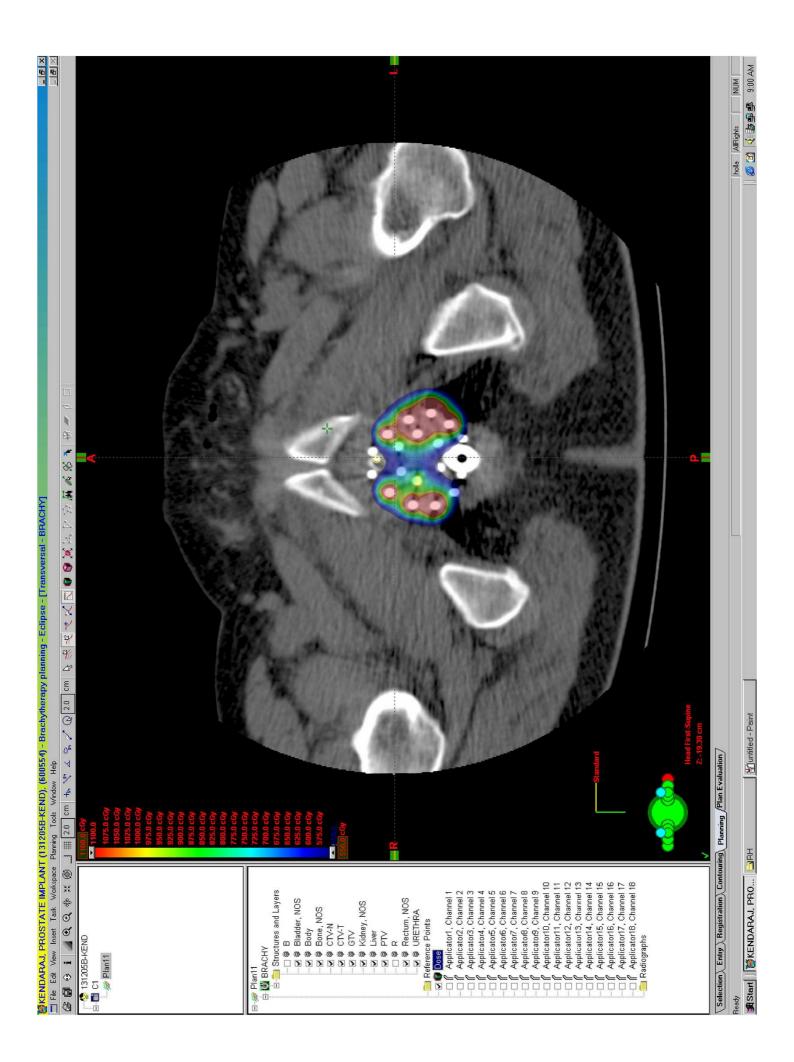
INTENSITY MODULATION UNIQUE TO HDR

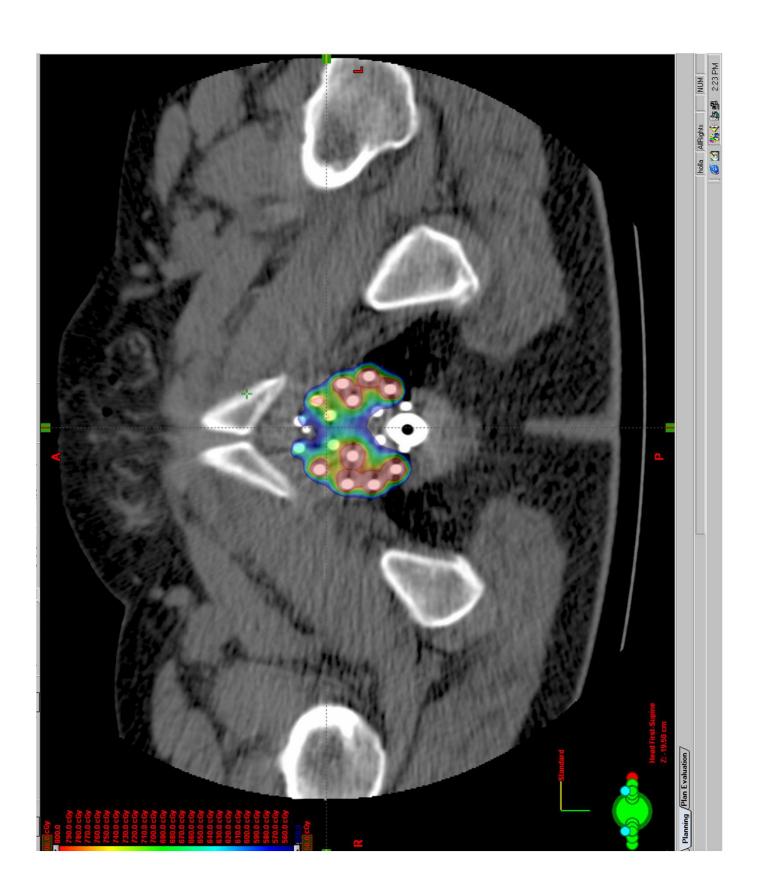
Decreased dose to urethra

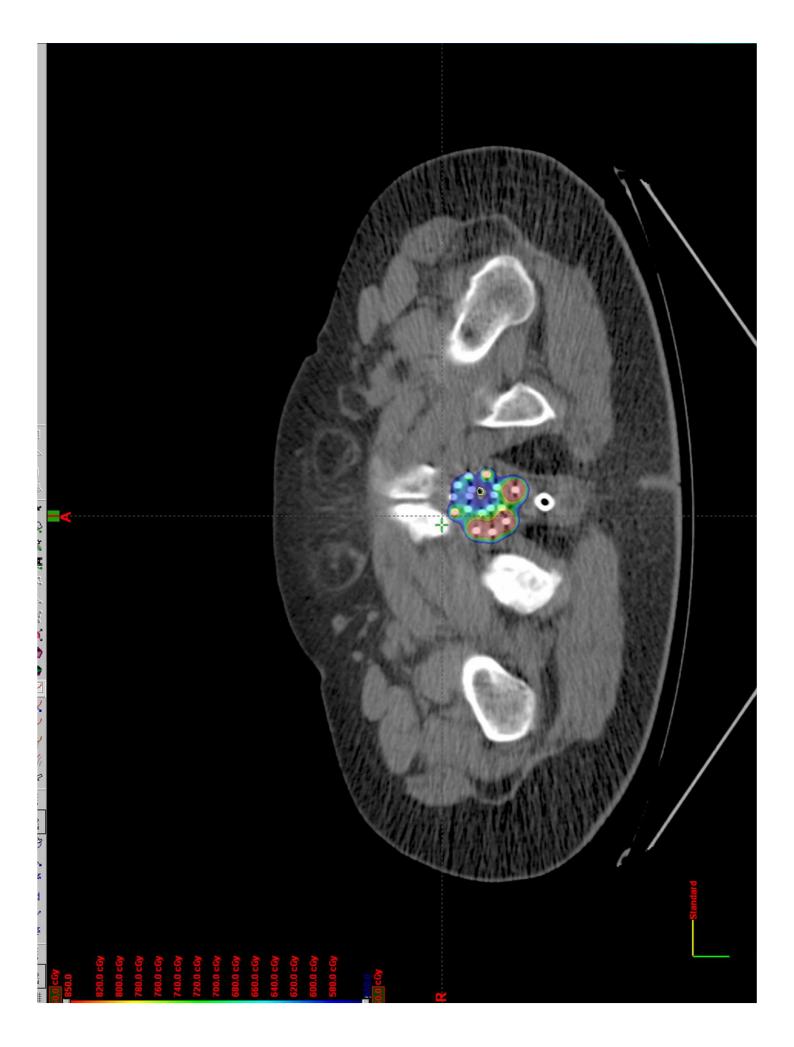
 Major limitation of IMRT is inability to do this (No significant reduction of urinary morbidity)











BC STRATIFIED BY RISK FACTOR GROUPS

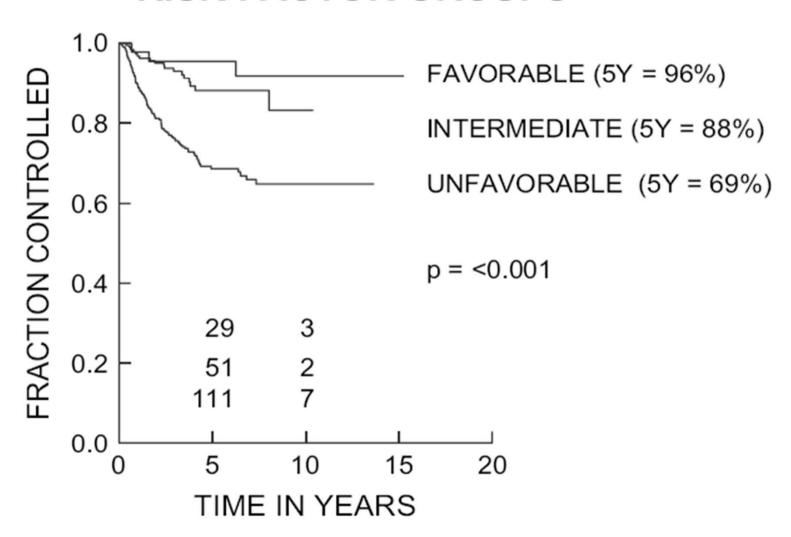


Fig. 2. Biochemical control (BC) stratified by risk factor groups.

BRACHYTHERAPY VERSUS 3D-CRT

5 YEAR BRFS (%)

	FR	IR	HR
3DCRT	90	70	47
SEEDS	94	82	65
SEED + XRT	85	77	45
HDR + XRT	96	87	69

Dose escalation with HDR prostate brachytherapy • A. A. MARINEZ et al.

Table 1. Equivalent dose per brachytherapy dose level

			BED* (Gy)	
Dose level	Brachytherapy dose	$\alpha/\beta = 10$	$\alpha/\beta = 5$	$\alpha/\beta = 1.2^{\dagger}$
Low dose	5.50 Gy × 3	67.1	70.7	80.2
	$6.00 \text{ Gy} \times 3$	70.0	74.3	86.1
	$6.50 \text{ GV} \times 3$	72.6	78.1	92.5
High dose	8.25 $G_{\rm V} \times 2$	72.0	78.8	94.2
,	8.75 Gv × 2	74.2	82.1	6.66
	9.50 Gv × 2	78.0	87.1	108.9
	$10.50 \text{ Gv} \times 2$	82.9	94.4	122.0
	11.50 Gy \times 2	87.0	8.66	136.3

Abbreviation: BED = biologically equivalent dose. * To 2 Gy per fraction, 70 Gy total external beam dose. † α/β ratio of 1.2 derived from our clinical trial (39).

HDR AS MONOTHRAPY

Favorable risk patients

• 5 year BRFS 98%

GRILLS et al, J. Urol., 2004

Longer follow up needed

HDR AS MONOTHERAPY

- 297 patients
- 8.5Gy x 4 in one implant Eq. 75.6 Gy in 1.8 Gy/ fr
 7 Gy x 6 in two implants Eq. 76 Gy in 2.0 Gy Gy/ fr
- 5 year results

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OAS – 94.5% DM – 0%
CSS – 100% BRFS – 91% (Phoenix)
LC – 98.9% GU toxicity -- < 1%
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Martinez et al, Brachytherapy, 7(2), 2008

HDR monotherapy vs. 103Pd monotherapy Table 1

Toxicities	HDR (%)	$^{103}\mathrm{Pd}~(\%)$	
Acute dysuria (Grades 1–3)	36	<i>L</i> 9	p < 0.001
Acute urinary frequency/urgency	54	92	p < 0.001
Urethral stricture			
Chronic urinary frequency/urgency	32	56	p < 0.001
Urethral stricture	∞	3	
Three-year impotency rate	16	45	

Most of above toxicities were Grade 1.

No difference in chronic dysuria, incontinence retention, and hematuria.

BRACHYTHERAPY

- Highly conformal dose to prostate (viz. 7) field 3D-CRT/IMRT)
- Radiobiologically appropriate
- Better normal tissue sparing
- No set up / organ motion and localization errors
- Convenient
- Quick

ADVANTAGES OF BRACHYTHERAPY OVER IMRT

- Significantly less investment
- Negligible recurring costs
- Cheap therapy
- Even best form of IMRT is still an XRT only.
- Radiobiologically superior
- Clinically and financially more relevant to Indian conditions

IMAGE BASED OPTIMIZED HIGH DOSE RATE CONFORMAL BRACHYTHERAPY IS THE BEST FORM OF IMRT