Intraluminal Brachytherapy for Cancer Nasopharynx



Rationale For Brachytherapy In Nasopharynx

- Good local control is achieved with Radiotherapy or RT+ Chemotherapy.
- Higher the dose higher the local control. [Good Dose Response Relation]
 [Vikram et al, Marks et al]
- Several techniques have been tried- Transpalatal interstitial implantation

Several endocavitory applicator based tech.

NPx is secluded, midline surrounded by bones, vessels and nerves
 hence endocavitory procedure most suitable.

Indications

- Boost for persistent disease after radiotherapy or chemoradiotherapy.
 T1, T2a tumors
 T2b tumors with good response
- 2. Recurrent cases

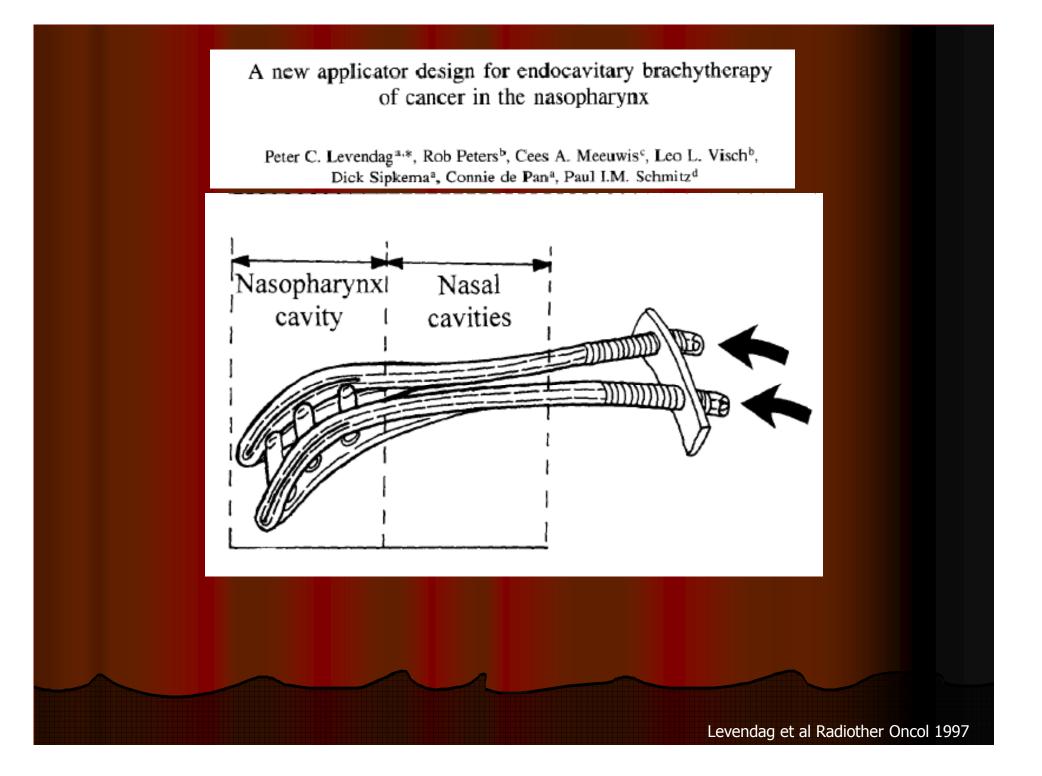
Suitable Candidate

1. Tumors restricted to Nasopharynx

with no involvement of nasal cavity or oropharynx

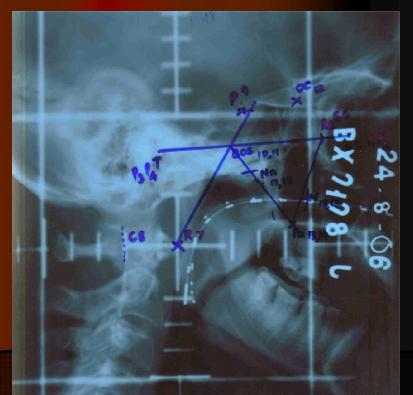
2. Thickness of CTV <10 mm-

superficial tumors/ Tx that have shrunk significantly well circumscribed, superficial local recurrences.



Treatment Planning

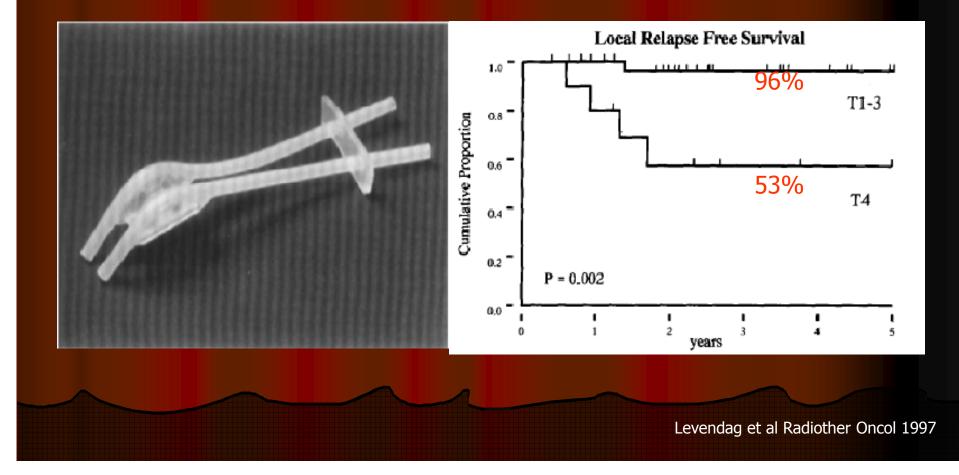
- Treatment planning based on orthogonal radiographs
- Patient positions representing target and critical structures are depicted on lateral and AP simulation film.
- Obtained dose distribution is optimized such that
- Target receives a dose of 3Gy(reference dose)
- Normal tissue points receive a dose as low as reasonably achievable.
 - 1. Nasopharynx point
 - 2. Base of skull point
 - 3. Node of Rouviere point
 - 4. Optic chiasm
 - 5. Retina
 - 6. Pituitary
 - 7. Spinal cord



Peter Levendag et al. [Radiotherapy Oncol 1997]

Protocol followed

T1,T3 -60Gy/30# EBRT, 3GyX6# HDR (2#/day, 6hrs apart) T4- 70Gy/35# EBRT, 3GyX4# HDR (2#/day, 6hrs apart).



Wang et al [Massachusetts General Hospital 25 years Experience]

5 acturial survival rates after RT related to Boost technique at MGH

	Local C	ontrol Rate (%)	Disease-Specific	Survival Rate (%)	
rapy therapy rapy		92 56 20	$82 \\ 53 \\ p - 0.0001 \\ 74$	p = 0.0001 78 43 $p = 0.0001$ 74	
therapy		26	p = 0.25		
		82	$p \approx 0.0001$	p = 0.0001	
n	Local Control Rate (%)	Disease-Specific Survival Rate (%)			
38 30 8	40 52 21 p - 0.31	59 55 71 p = 0.33			irradiati
	therapy therapy therapy therapy therapy <i>n</i> 38 30	rapy therapy therapy therapy therapy <i>Local Control</i> <i>n Rate</i> (%) 38 40 30 52 8 21	therapy 56 rapy 20 therapy 26 rapy 112 therapy 82 Local Control Disease-Specific n Rate (%) Survival Rate (%) 38 40 59 30 52 55 8 21 71	rapy 92 82 therapy 56 53 $p = 0.0001$ $p = 0.0001$ rapy 20 74 therapy 26 60 $p = 0.25$ $p = 0.25$ rapy 112 81 therapy 82 56 $p = 0.0001$ $p = 0.0001$ Local Control Disease-Specific n Rate (%) Survival Rate (%) 38 40 59 30 52 55 8 21 71 $p = 0.31$ $p = 0.33$ 5 year-actu	rapy928278therapy565343 $p = 0.0001$ $p = 0.0001$ $p = 0.0001$ rapy207474therapy2660 $p = 0.25$ $p = 0.08$ rapy1128178therapy8256 $p = 0.0001$

Managing nasopharyngeal carcinoma with intracavitary brachytherapy: one institution's 45-year experience

Nancy Lee*, Rex Hoffman, Theodore L. Phillips, Ping Xia, Jeanne M. Quivey, Vivian Weinberg, I-Chow J. Hsu

Overall treatment outcome			
	Primary	Recurrent	
	disease	disease	EE patiente received ILPT
Variable	(n = 43)	(<i>n</i> = 12)	55 patients received ILBT
Follow-up (months)			EPDT doco E4 72Cy primany
Median	36	.50	EBRT dose-54-72Gy-primary
Range	1-278	6-204	20,420
Local control			30-42Gy-recurrer
3-year estimated	94%	75%	
5-year estimated	89%	64%	Brachytherapy- 5-7Gy-HDR
Failed	6	4	, , , , ,
Distant metastasis-free rate			10-54Gy-LDR
3-year estimated	79%	100%	
5-year estimated	75%	100%	
Failed	9	2	
Overall survival			
3-year estimated	90%	91%	
5-year estimated	86%	91%	
Died	9	2	

EBRT+ Brachytherapy were well tolerated.

Nancy Lee et al Brachytherapy 2002

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THE ROLE OF BRACHYTHERAPY IN EARLY-STAGE NASOPHARYNGEAL CARCINOMA

JOSEPH T. CHANG, M.D.,* LAI-CHU SEE, M.S.,[†] SIMON G. TANG, M.D., M.S.,* STEVE P. LEE, M.D., M.S.,[‡] CHUN-CHIEH WANG, M.D.* AND JI-HONG HONG, M.D., PH.D.*

150 patients 100 pts.- 64.8-68.4Gy EBRT, 5-16Gy/1-3# HDR 50 patients- 68.4-72.8Gy EBRT, no brachytherapy 3 prognostic groups- group I- <72.5Gy, No brachy group II- 72.5-75Gy, 1-2# brachy

					Consultantions	
	n	Disease free 5-year rate (%)	Local control 5-year rate (%)	Disease specific 5-year rate (%)	Complications Group I- 1.9%	
Treatment group*	50	71.6	73.7	77.0	Group II-4.2%	
II III p-value	71 58	92.4 77.0 0.0039	93.9 79.5 0.0064	95.5 82.4 0.0194	Group III- 13.8	
Pairwise comparison I vs. II I vs. III II vs. III		0.0006 [†] 0.4489 [†] 0.0078 [†]	0.0007 [†] 0.4120 [†] 0.0154 [†]	$\begin{array}{c} 0.0106^{\dagger} \\ 0.7459^{\dagger} \\ 0.1020^{\dagger} \end{array}$	P=0.03	

group III- >75Gy, 3# brachy

1.9% 4.2% 13.8%

Chang et al IJROBP 2002

Tata Memorial Hospital Experience

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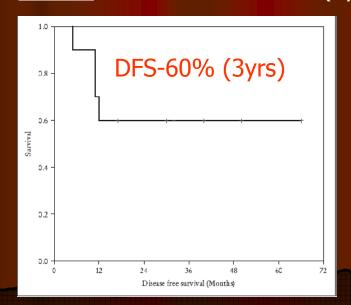
Bull Cancer 2005 ; 92 (7-8) : E45-50

High dose rate brachytherapy boost for primary nasopharyngeal carcinoma: preliminary results of an ongoing prospective study

1998-2003, 10 patients of primary NPC

Median EBRT dose-66Gy

Median HDR-Brachy Boost dose-12Gy/1-4# <u>Results</u> Local control- 90%(3yrs)



Rotterdam Silicone Nasopharyngeal applicator

Patient cha	Patients (nb)	
Age (years)	\leq 50 years	6
	> 50 years	4
Sex	Male	7
	Female	3
Tumor Status	T1-2	6
	T3-4	4
Nodal Status	Node positive	8
	Node negative	2

<u>Toxicity</u>

No patient had significant late toxicities except

Mild Xerostomia-8/10

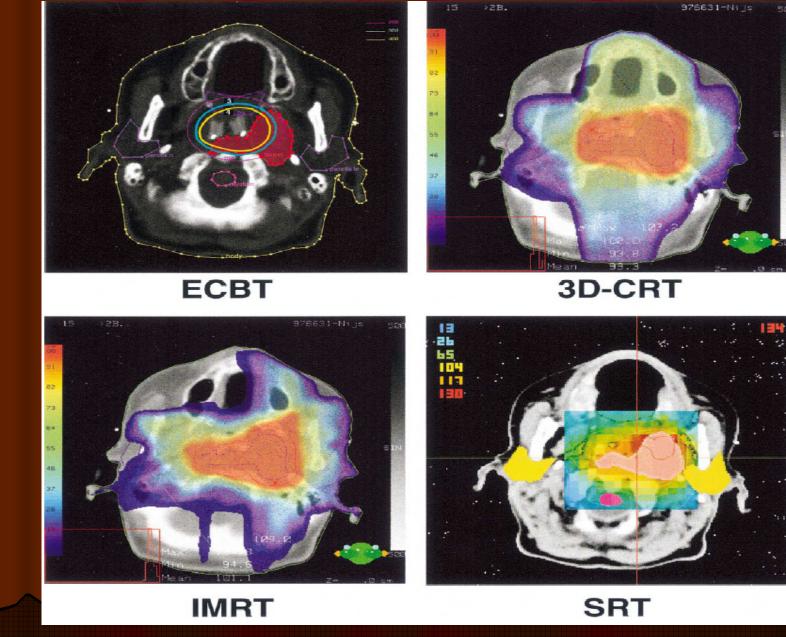
Persistent crust formation- 1/10

R Malde et al Bull Cancer 2005

Literature Review of studies using ILBT as BOOST

Author [Ref] T- Stage		Dose (Gy)		Chemo-	5-yr local	5-yr survival
		EBRT	Brachytherapy	therapy	control	
Chang [20] 1996	T1-2 (133)	64.8-68.4 Gy	HDR: 5-16.5 Gy/ 1-3 # @ 2 cm off axis	Nil	< 72.5 Gy : 73% 72.5-75Gy: 94% > 75 Gy : 79%	72% 92% 77%
Slevin [22] 1997	T1 = 1, T2 = 4 T3 = 3	45-60 Gy	HDR: 5-7.5Gy/ 1#	Nil	87% (3y)	37% (3y DFS) 75% (3y OS)
Levendag [7] 1998	T1 = 3, T2 = 9 T3 = 17, T4 = 13	T1-3 = 60 Gy T4 = 70 Gy	T1-3: 18Gy/6 # T4: 16Gy/4 # @ 1 cm off-axis	1 (2.5%)	86% (3y)	71% (3y DFS)
Syed [12] 2000	T1 = 1, T2 = 4 T3 = 6, T4 = 4	50-60 Gy	HDR Implant: 33-37 Gy	5 (33%)	59%	74% (5y DFS) 61% (5y OS)
Teo [18] 2000	T1 = 74 T2 = 89	60 Gy	HDR:18-24 Gy/3# @1cm off-axis	10 (6%)	93%	88% (5y DFS)
De Nittis [23] 2002	T1-T3 = 11	64-70 Gy (66 Gy median)	HDR: 6-15 Gy / 1-2 # @ 0.5 cm	11 (100%)	100% (3y)	100% (3y OS)
Lee [19] 2002	T1 = 21, T2 = 18 T3 = 4	54-72 Gy	HDR 5-7Gy/1-2 # LDR: 10-54 Gy.	17 (40%)	89%	75% (5y DFS) 86% (5y OS)
Levendag [17] 2002	T1 = 7, T2 = 39 T3 = 11, T4 = 14	60-70 Gy	HDR: 11-18 Gy / 4-6 # @ 1 cm off axis	20 (41%)	I-IIB: 100% (2y) III-IVB: 86% (2y)	I-IIB: 90% (2y DFS) 61% (2y OS) III-IVB:74% (2y DFS) 66% (2y OS)
Ozyar [21] 2002	T1 = 45, T2 = 32 T3 = 13, T4 = 16	58-71 Gy (65.4 Gy median)	HDR: 12 Gy/3 # @ 1 cm off-axis	55 (51%)	86% (3y)	76% (3y CSS) 67% (3y DFS)
Lu [26] 2004	T1 = 22 T = 11	70 Gy	HDR: 10 Gy/2 # @1 cm off axis	33 (100%)	93.6% (2y)	74% (2y DFS) 82% (2 y OS)
TMH Present study	T1-2 = 6 T3-4 = 4	60-70 Gy	HDR: 5-14 Gy / 1-4 # @ 1 cm off axis	8 (80%)	90% (3y)	60% (3y DFS)

Malde et al Bull Cancer2005



Complications

- 1. Skin/soft tissue fibrosis
- 2. Hearing deficit
- 3. Xerostomia
- 4. Trismus
- 5. Taste alteration
- 6. Otitis media
- 7. Neuroendocrine dysfunction
- 8. Choanal stenosis/atresia
- 9. Olfactory dysfunction
- 10. Palatal and sphenoidal bone necrosis

Conclusions

- 1. ILRT has a definite role in patients with early stage nasopharyngeal cancer as boost after EBRT.
- 2. ILRT is also effective in cases of recurrent nasopharyngeal cancer specially for re-irradiation.
- 3. With the advent of 3DCRT and IMRT, the role of nasopharyngeal brachytherapy needs to be redefined.
- 4. In limited resource settings ILRT nasopharynx continues to be important.

