

Chemoradiation in Ca Nasopharynx

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Ca Nasopharynx

Stage	T,N&M
Stage I	T1N0
Stage II	T1N1,T2 N0-N1
Stage III	T3N0-N1 or T-T3 N2
Stage IVa	T4N0-2or Any T,N3
Stage IVb	M1

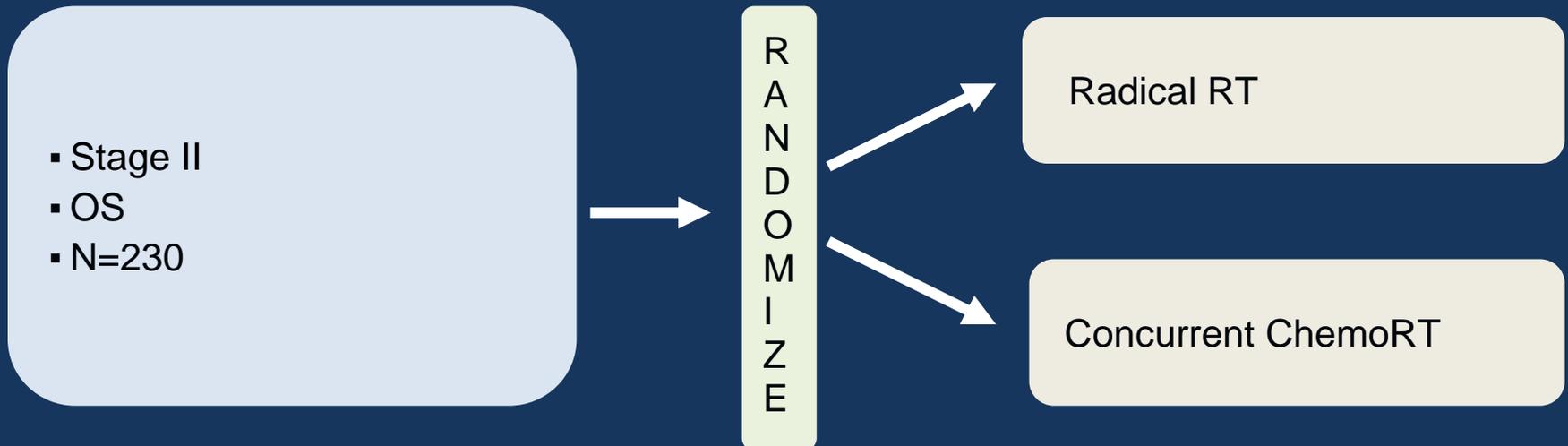
ARTICLE

Concurrent Chemoradiotherapy vs Radiotherapy Alone in Stage II Nasopharyngeal Carcinoma: Phase III Randomized Trial

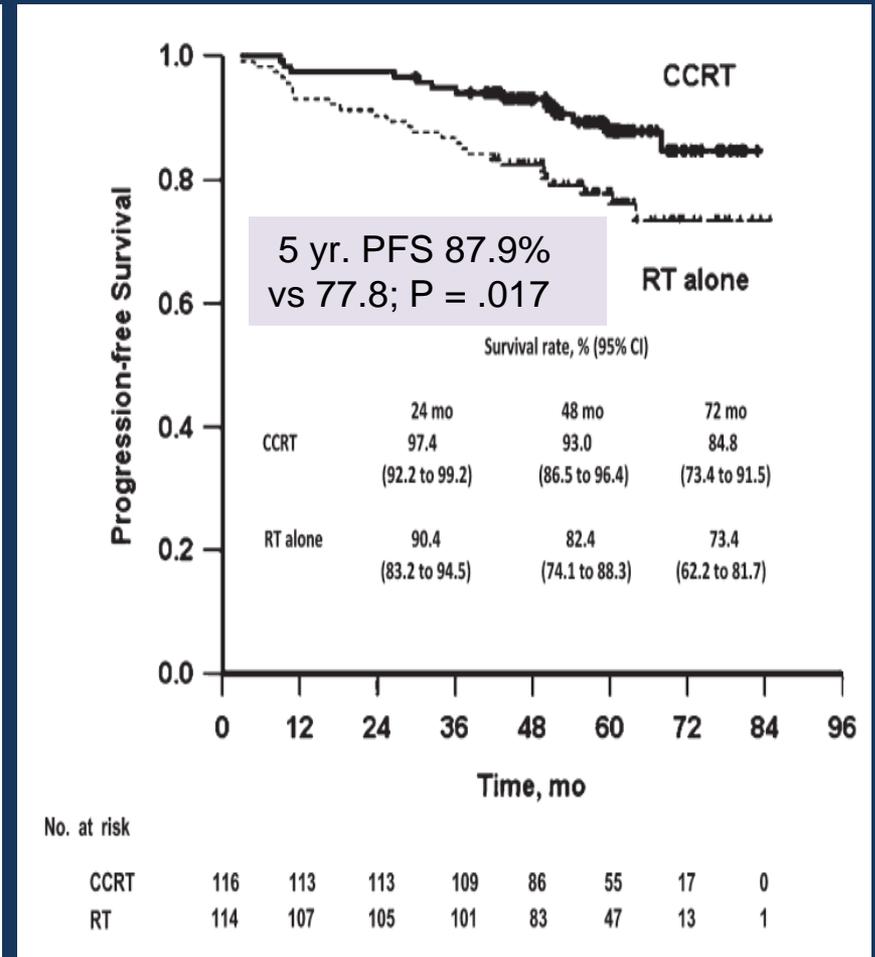
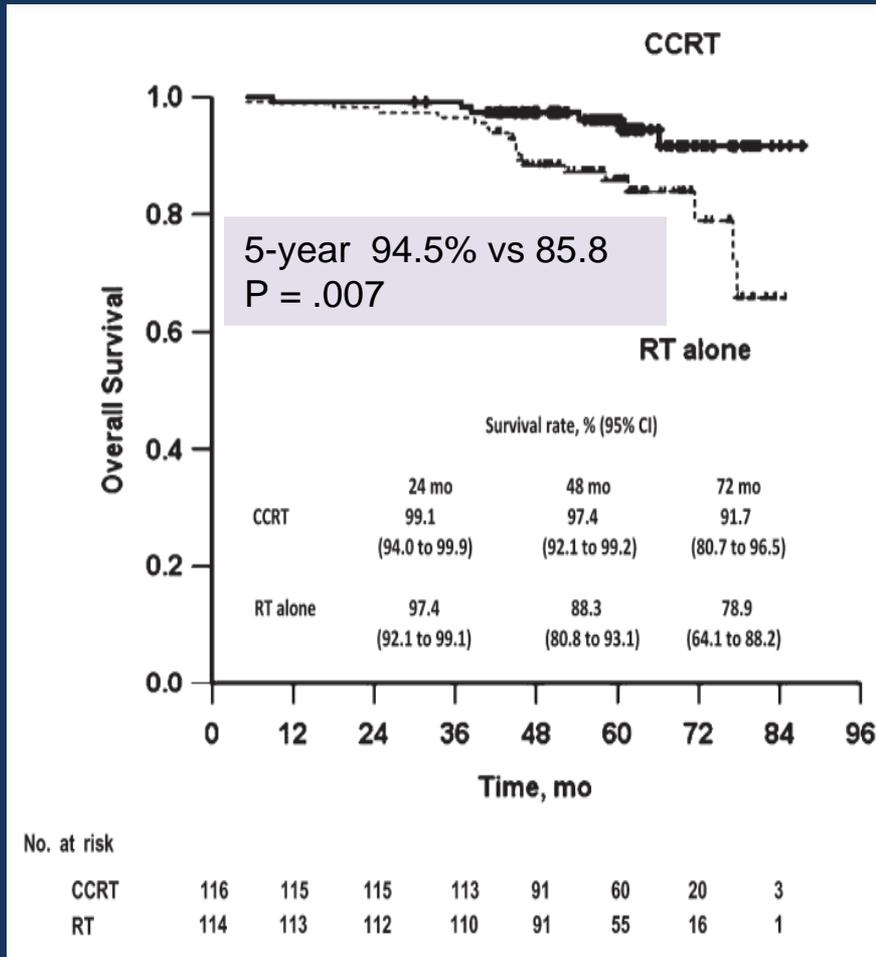
Qiu-Yan Chen, Yue-Feng Wen, Ling Guo, Huai Liu, Pei-Yu Huang, Hao-Yuan Mo, Ning-Wei Li, Yan-Qun Xiang, Dong-Hua Luo, Fang Qiu, Rui Sun, Man-Quan Deng, Ming-Yuan Chen, Yi-Jun Hua, Xiang Guo, Ka-Jia Cao, Ming-Huang Hong, Chao-Nan Qian, Hai-Qiang Mai

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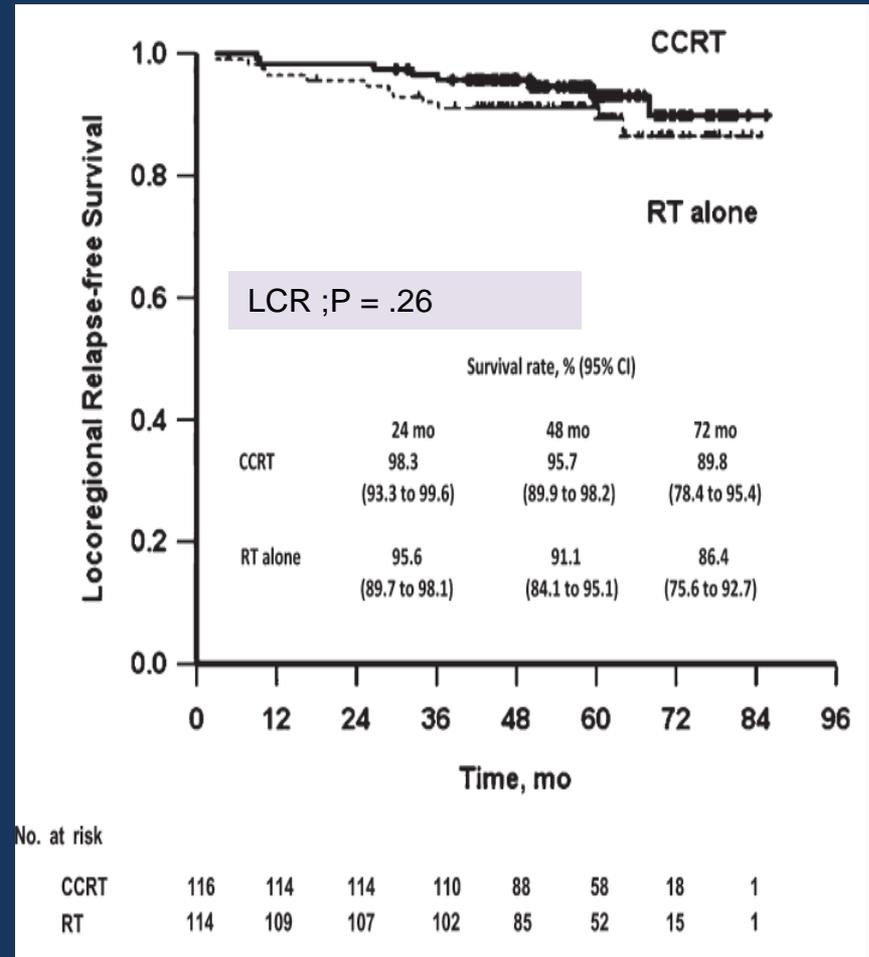
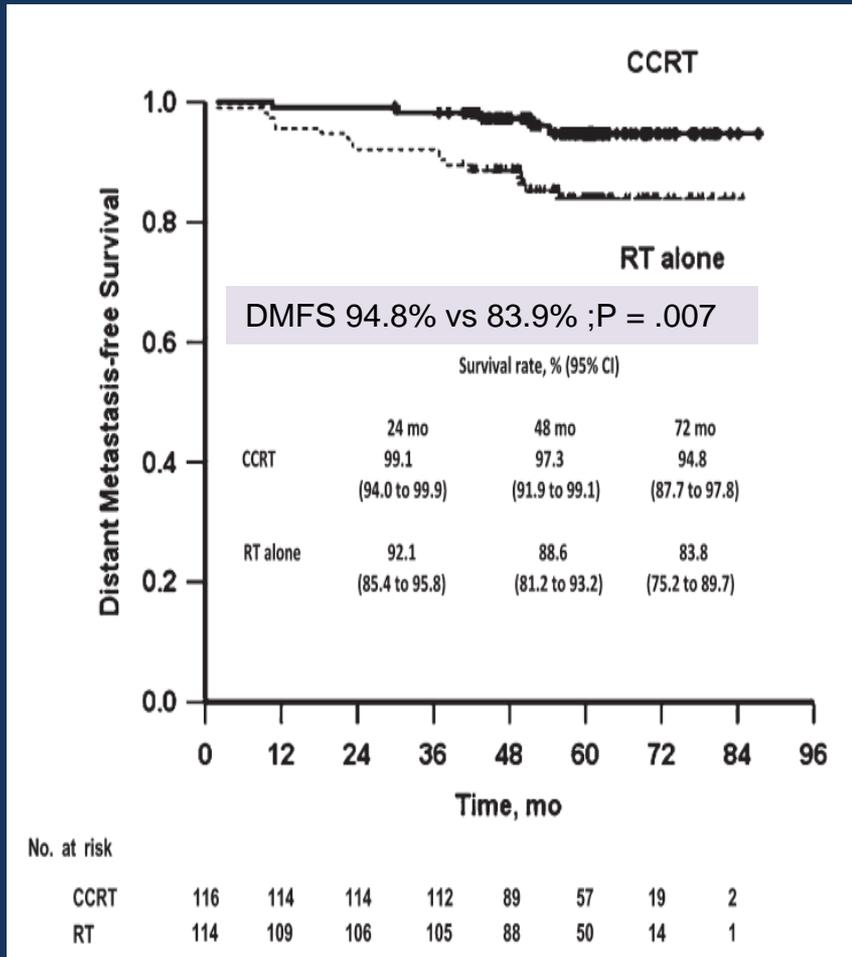
Correspondence to: Hai-Qiang Mai, MD, PhD, Department of Nasopharyngeal Carcinoma, Sun Yat-sen University Cancer Center, 651 Dongfeng Rd East, Guangzhou 510060, People's Republic of China (e-mail: maihq@mail.sysu.edu.cn).



Median follow up -60 months



Median follow up -60 months



Role of Chemo in Stage II

RESEARCH ARTICLE

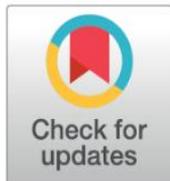
The role of concurrent chemotherapy for stage II nasopharyngeal carcinoma in the intensity-modulated radiotherapy era: A systematic review and meta-analysis

Fang Liu¹✉, Tao Jin²✉, Lei Liu¹✉*, Zhongzheng Xiang¹, Ruonan Yan¹, Hui Yang³*

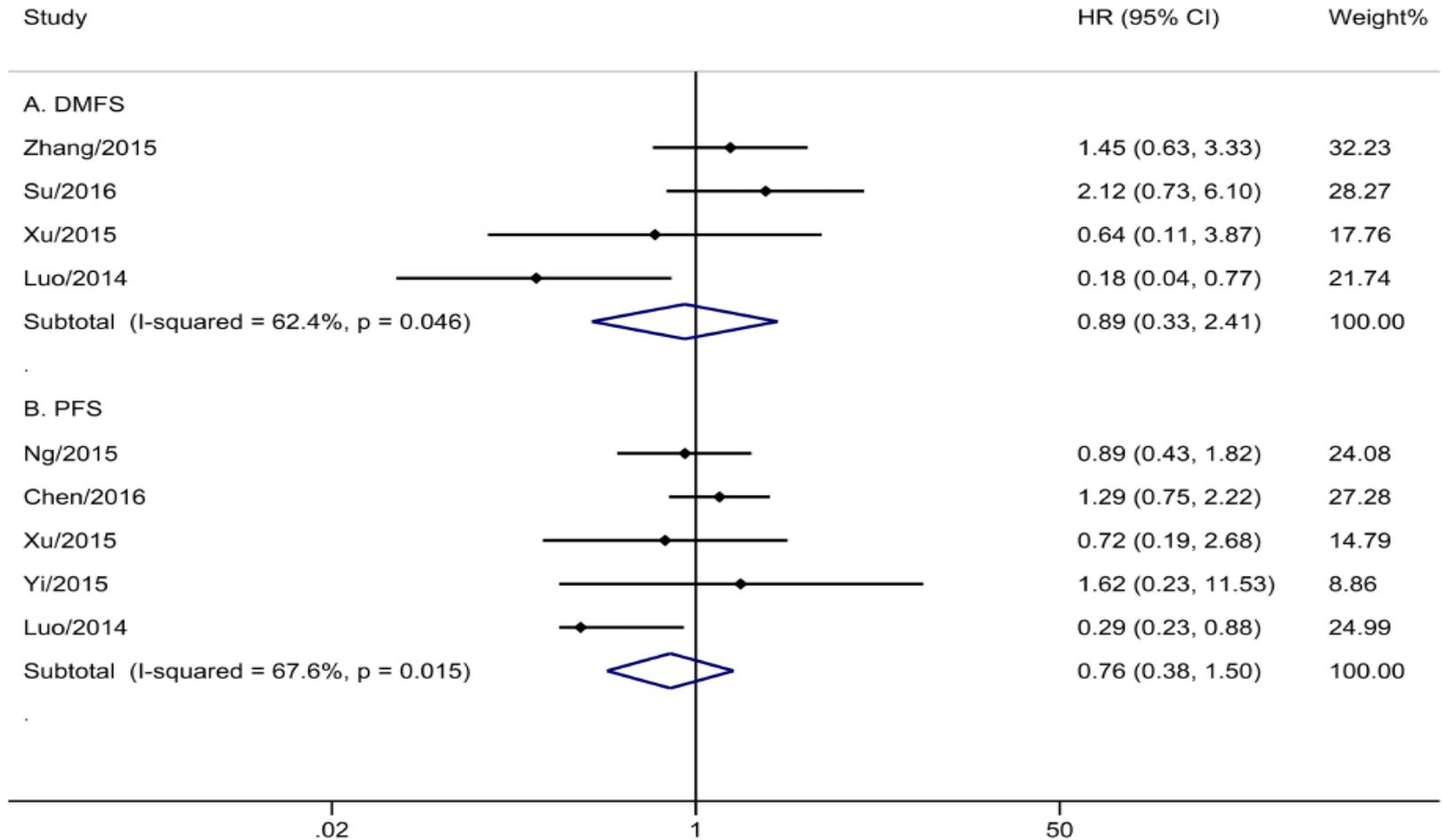
1 The Department of Medical Oncology, Cancer Center, State Key Laboratory of Biotherapy, West China Hospital, Sichuan University, Chengdu, Sichuan, China, **2** Department of Urology, Institute of Urology, West China Hospital, Sichuan University, Chengdu, Sichuan, China, **3** Department of Otolaryngology-Head and Neck Surgery, West China Hospital, Sichuan University, Chengdu, Sichuan, China

✉ These authors contributed equally to this work.

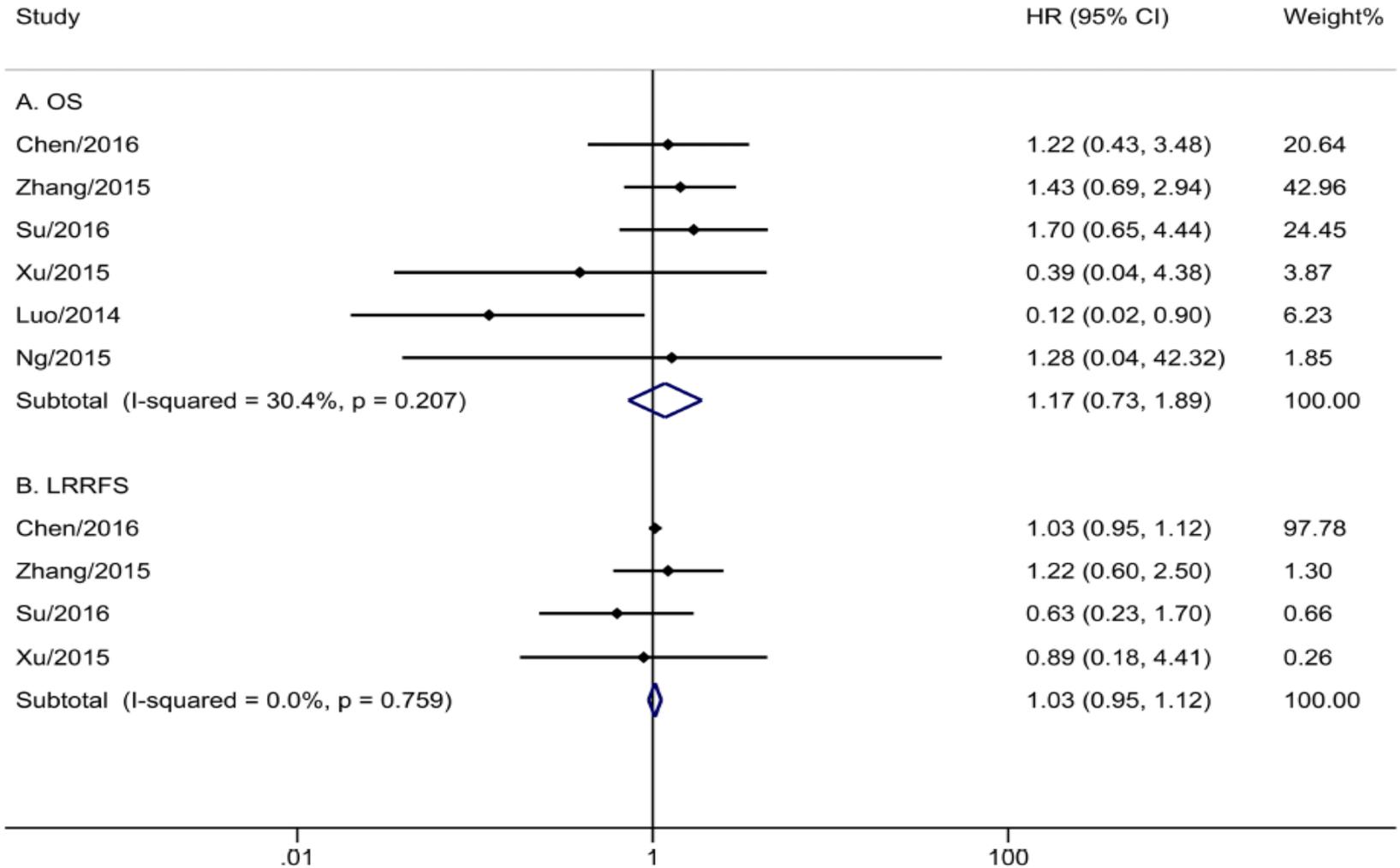
* yangfree1966@163.com (HY); liuleihx@gmail.com (LL)



PFS & DMFS



OS



Stage II & T3N0 Ca Nasopharynx(n=341)



RT alone (IMRT)

CCRT Cisplatin 100/m2

Median follow up ,41 months, 3-year FFS was 90.7% Vs92.1%, $p_{\text{non-inferiority}}=0.00017$)

	RT group(%)	CCRT group (%)	p Value
Intention-to-treat population	n = 172	n = 169	
3-yr overall survival	98.7	99.4	0.178
3-yr distant metastasis-free survival	95.6	98.1	0.136
3-yr locoregional recurrence-free survival	94.4	93.9	0.975

Exclusion criteria

- 1.lymph node $\geq 30\text{mm}$,
- 2.positive neck lymph node at level IV
3. plasma EBV DNA level ≥ 4000 copy/ml.

Toxicity less in RT group

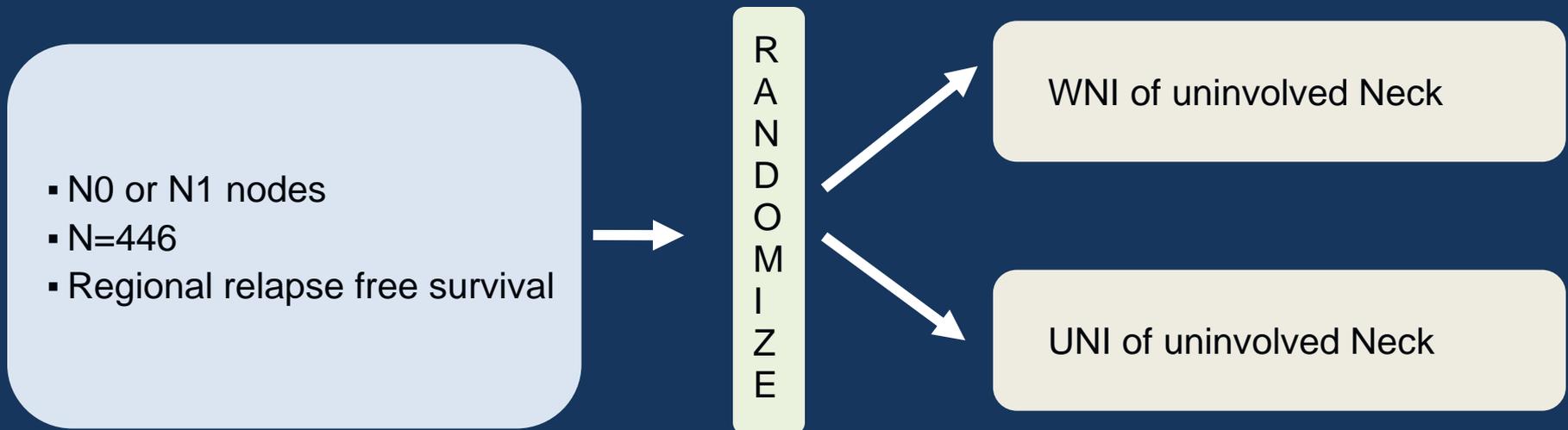
No difference in
OS
DMFS
LRRFS

ASCO 2022 Abstract 6000

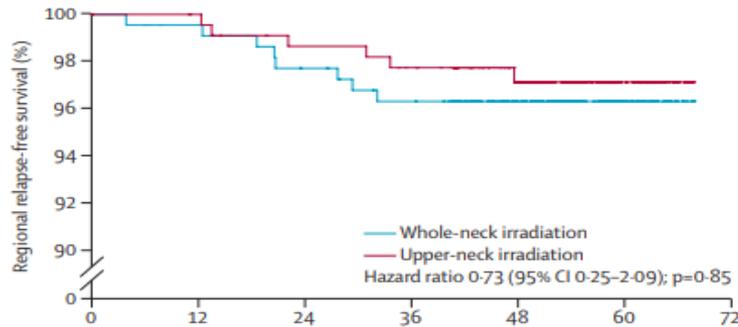
Elective upper-neck versus whole-neck irradiation of the uninvolved neck in patients with nasopharyngeal carcinoma: an open-label, non-inferiority, multicentre, randomised phase 3 trial



Ling-Long Tang*†, Cheng-Long Huang*, Ning Zhang*, Wei Jiang*, Yi-Shan Wu*, Shao Hui Huang, Yan-Ping Mao, Qing Liu, Ji-Bin Li, Shao-Qiang Liang, Guan-Jie Qin, Wei-Han Hu, Ying Sun, Fang-Yun Xie, Lei Chen†, Guan-Qun Zhou†, Jun Ma†

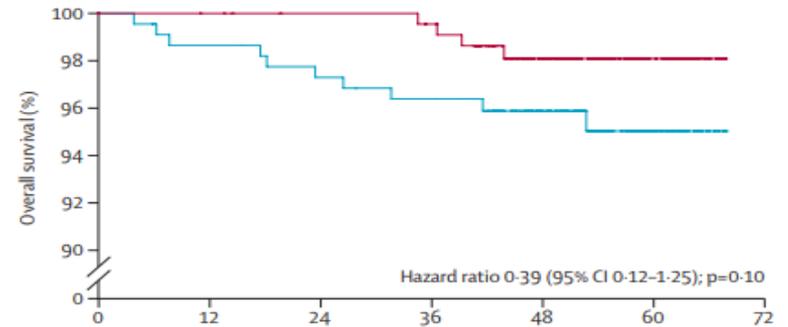


Median follow-up -53 months

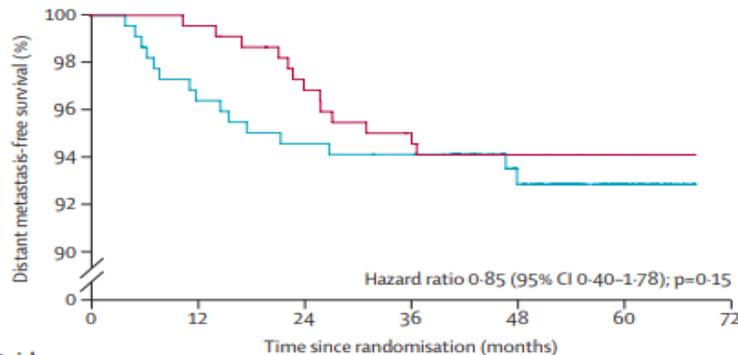


Number at risk (number censored)

Whole-neck irradiation	222 (0)	218 (3)	211 (3)	204 (4)	143 (61)	43 (100)	0 (43)
Upper-neck irradiation	224 (0)	223 (1)	217 (3)	213 (2)	150 (62)	47 (103)	0 (47)

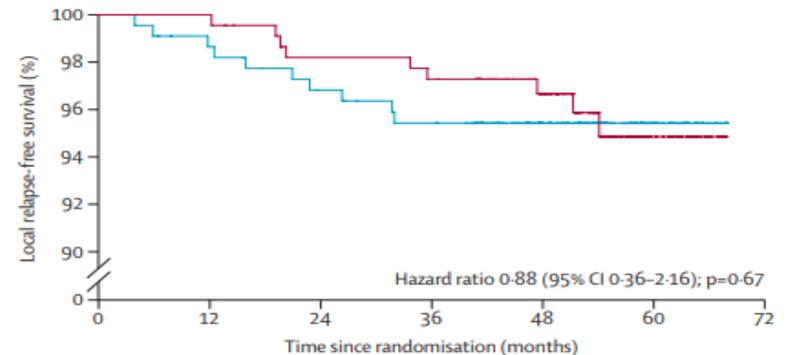


Whole-neck irradiation	222 (0)	218 (1)	214 (1)	210 (2)	146 (63)	45 (100)	0 (45)
Upper-neck irradiation	224 (0)	223 (1)	220 (3)	218 (1)	151 (64)	47 (104)	0 (47)



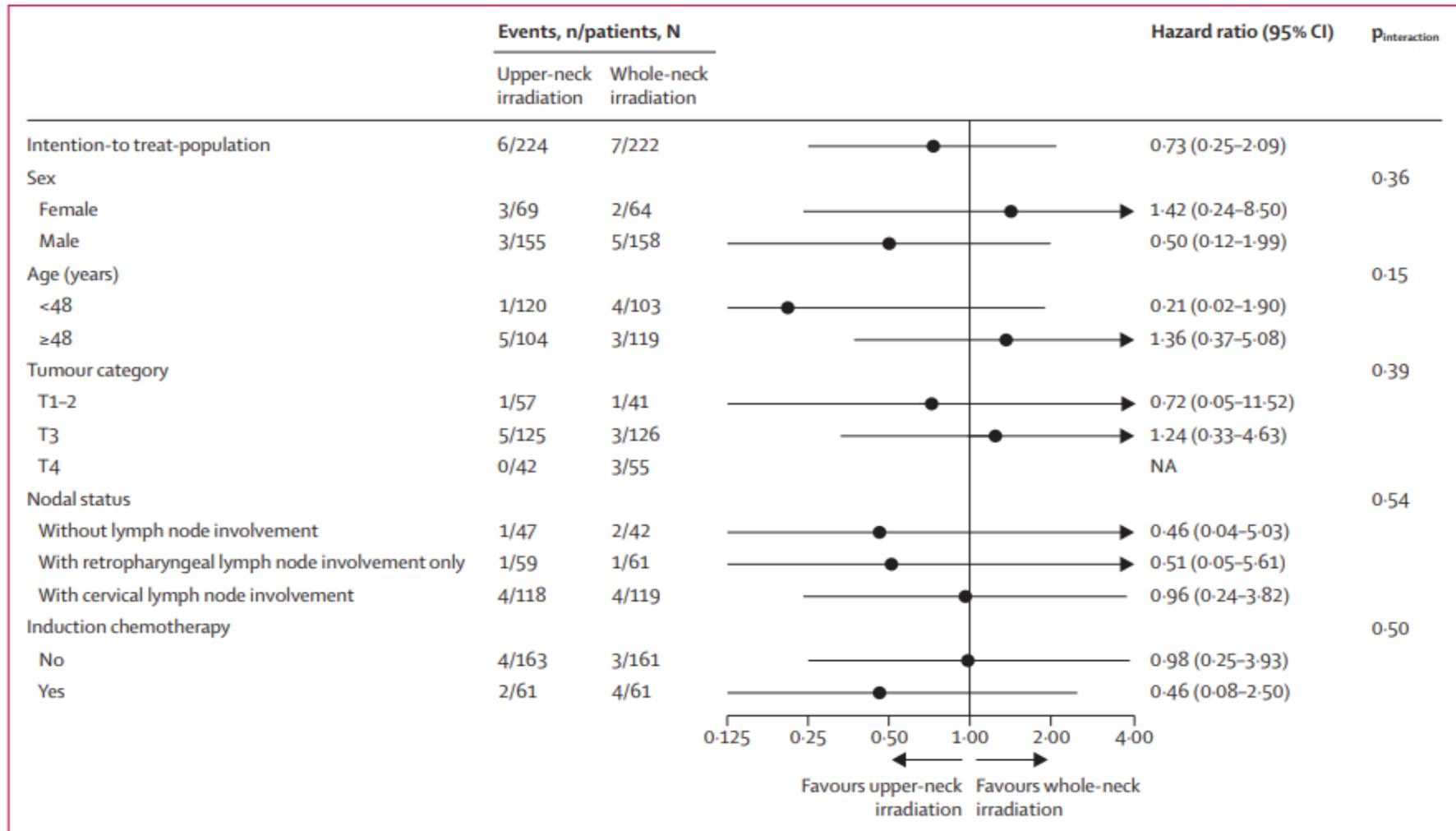
Number at risk (number censored)

Whole-neck irradiation	222 (0)	213 (1)	207 (2)	204 (2)	141 (61)	44 (97)	0 (44)
Upper-neck irradiation	224 (0)	222 (1)	214 (3)	208 (1)	145 (61)	47 (98)	0 (47)



Whole-neck irradiation	222 (0)	217 (2)	210 (3)	204 (3)	142 (62)	43 (99)	0 (43)
Upper-neck irradiation	224 (0)	223 (1)	216 (3)	212 (2)	148 (63)	44 (102)	0 (44)

Subset analysis



Toxicity

	Upper-neck irradiation group (n=222)			Whole-neck irradiation group (n=222)		
	Grade 1-2	Grade 3	Grade 4	Grade 1-2	Grade 3	Grade 4
Any acute toxicities						
Dermatitis	114 (51%)	1 (<1%)	0	123 (55%)	1 (<1%)	0
Mucositis	125 (56%)	20 (9%)	0	131 (59%)	22 (10%)	1 (<1%)
Dry mouth	159 (72%)	0	0	161 (73%)	0	0
Dysphagia	7 (3%)	0	0	14 (6%)	0	0
Weight loss	114 (51%)	0	0	125 (56%)	0	0
Trismus	0	0	0	1 (<1%)	0	0
Subcutaneous oedema	1 (<1%)	0	0	0	0	0
Any late toxicities*						
Skin†	32 (14%)	0	0	55 (25%)	0	0
Neck tissue damage	50 (23%)	0	0	86 (39%)	2 (1%)	0
Hypothyroidism	63 (28%)	3 (1%)	0	84 (38%)	3 (1%)	0
Dysphagia	38 (17%)	0	0	69 (31%)	2 (1%)	0
Hoarseness	3 (1%)	0	0	1 (<1%)	0	0
Dry mouth	153 (69%)	11 (5%)	0	160 (72%)	15 (7%)	0
Trismus	2 (1%)	0	0	5 (2%)	0	0
Auditory	110 (50%)	0	0	137 (62%)	2 (1%)	0
Temporal lobe injury	17 (8%)	0	0	21 (10%)	0	0

Data are n (%). Safety analyses were done in the safety population, comprising all patients who commenced the randomly assigned treatment. *One patient in the whole-neck irradiation group died within 3 months after radiotherapy and thus the late toxicity analysis included 221 patients in the whole-neck irradiation group. †Grade 1 skin toxicity included slight atrophy, pigmentation change, and some hair loss; grade 2, included patch atrophy, moderate telangiectasia, and total hair loss; grade 3 or 4 skin toxicity was not observed in this trial.

Stage III& IVa

Intergroup Trial N=193

Concurrent Chemo RT
70 Gy/ 35# + 3 cycles of CDDP

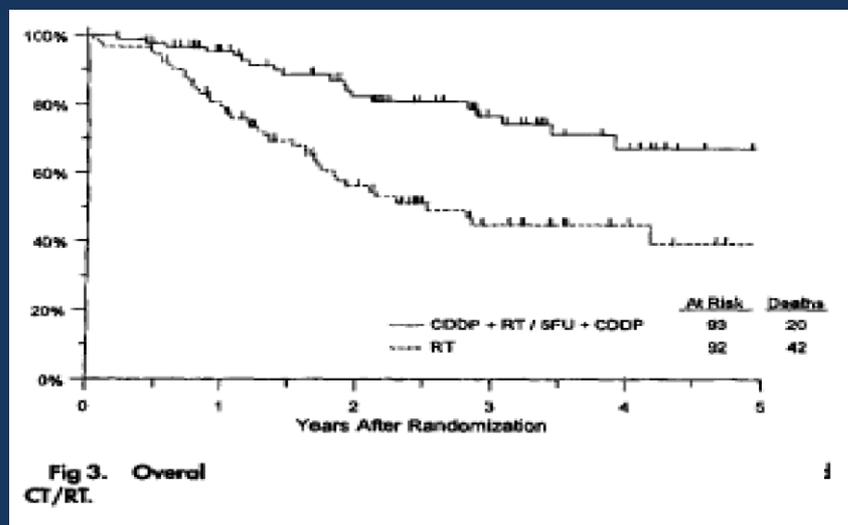
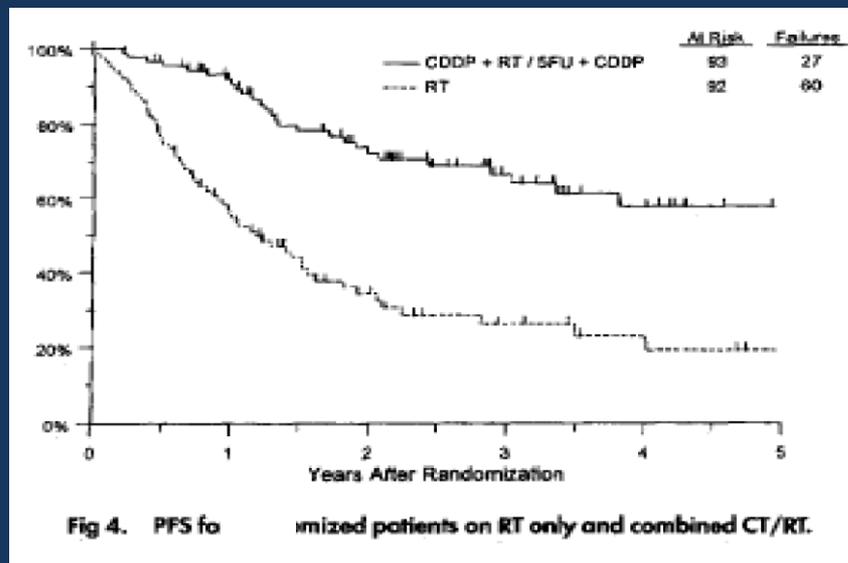
RT alone

Adjuvant Chemotherapy
(CDDP + 5 FU)

x 3

U.S. Intergroup 0099

- 3Y PFS 69% (CRT) vs. 24% (RT alone), $p < 0.001$
- 3Y OS 76% (CRT) vs. 46% (RT alone), $p = 0.005$



U.S. Intergroup 0099

Issues

- Flawed study design
 - Are the benefits from chemo due to concurrent administration, adjuvant, or both?
- Terminated early after interim analysis showed survival benefit
- RT alone arm performed worse than expected
- Old RT techniques
- Many patients enrolled had WHO type I NPC (not EBV-associated)
- Adjuvant PF chemotherapy only feasible in some patients

Randomized Trial of Radiotherapy Versus Concurrent Chemoradiotherapy Followed by Adjuvant Chemotherapy in Patients With American Joint Committee on Cancer/International Union Against Cancer Stage III and IV Nasopharyngeal Cancer of the Endemic Variety

Joseph Wee, Eng Huat Tan, Bee Choo Tai, Hwee Bee Wong, Swan Swan Leong, Terence Tan, Eu Tiong Chua, Edward Yang, Khai Mun Lee, Kam Weng Fong, Hoon Seng Khoo Tan, Kim Shang Lee, Susan Loong, Vijay Sethi, Eu Jin Chua, and David Machin

Preliminary Results of a Randomized Study on Therapeutic Gain by Concurrent Chemotherapy for Regionally-Advanced Nasopharyngeal Carcinoma: NPC-9901 Trial by the Hong Kong Nasopharyngeal Cancer Study Group

Anne W.M. Lee, W.H. Lau, Stewart Y. Tung, Daniel T.T. Chua, Rick Chappell, L. Xu, Lillian Siu, W.M. Sze, T.W. Leung, Jonathan S.T. Sham, Roger K.C. Ngan, Stephen C.K. Law, T.K. Yau, Joseph S.K. Au, Brian O'Sullivan, Ellie S.Y. Pang, S.K. O, Gordon K.H. Au, and Joseph T. Lau

Subsequent Asian Trials Contradictory

			3Y OS	Rate of DM
Wee, JCO, 2005 (Singapore)	221 pts WHO type II/II Mostly T3-4 +/-or N2-3	Cis/RT → PF X3	80%	18%
		RT alone	65%	38%
			<i>p=0.0061</i>	<i>p=0.0029</i>
Lee, JCO, 2005 (Hong Kong)	348 pts WHO type II/II Mostly N2-3	Cis/RT → PF X3	78%	24%
		RT alone	78%	27%
			<i>p=0.97</i>	<i>p=0.96</i>

J Clin Oncol 23:6730-6738.

J Clin Oncol 23:6966-6975.



Meta-analysis in NPC MAC-NPC Collaborative Group

CLINICAL INVESTIGATION

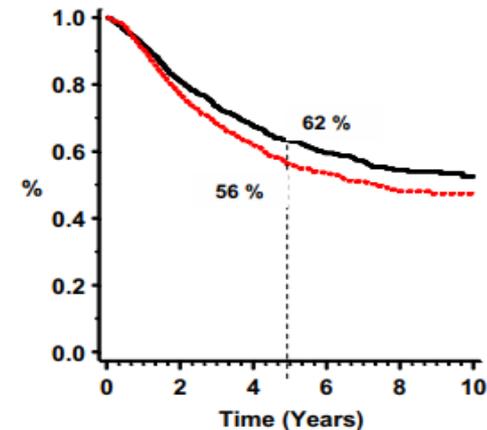
Head and Neck

CHEMOTHERAPY IN LOCALLY ADVANCED NASOPHARYNGEAL CARCINOMA: AN INDIVIDUAL PATIENT DATA META-ANALYSIS OF EIGHT RANDOMIZED TRIALS AND 1753 PATIENTS

BERTRAND BAUJAT, M.D.,* HÉLÈNE AUDRY, M.SC.,* JEAN BOURHIS, M.D., PH.D.*
ANTHONY T. C. CHAN, M.D.,† HALUK ONAT, M.D.,‡ DANIEL T. T. CHUA, M.D.,§ DORA L. W. KWONG,
M.D.,§ MUHYI AL-SARRAF, M.D.,|| KWAN-HWA CHI, M.D.,¶ MASATO HAREYAMA, M.D.,#
SING F. LEUNG, M.D.,† KULLATHORN THEPHAMONGKHOL, M.D.,* AND
JEAN-PIERRE PIGNON, M.D., PH.D.,* ON BEHALF OF THE MAC-NPC COLLABORATIVE GROUP

*Institut Gustave-Roussy, Villejuif, France; †Department of Clinical Oncology, Prince of Wales Hospital, Hong-Kong, China;
‡Istanbul University, Institute of Oncology, Istanbul, Turkey; §Department of Clinical Oncology, Queen Mary Hospital,
Hong-Kong, China; ||Wayne State University, Detroit, MI; ¶Taiwan Cooperative Oncology Group, Taipei, Taiwan;
#Department of Radiology, Sapporo Medical University, Sapporo, Japan

- HR for death=0.82 (95% CI 0.71-0.95)
- 6% absolute survival benefit at 5 years
- Greatest benefit from concurrent chemo
HR=0.60 (concurrent)
HR=0.97 (adjuvant)
HR=0.99 (induction)



Patients at risk

RT+CT	990	730	502	281	120	46
RT alone	985	683	443	237	100	41

CCRT Vs CCRT followed by Adjuvant chemo?

Concurrent chemoradiotherapy plus adjuvant chemotherapy versus concurrent chemoradiotherapy alone in patients with locoregionally advanced nasopharyngeal carcinoma: a phase 3 multicentre randomised controlled trial



Lancet Oncol 2012; 13: 163-71

Lei Chen, * Chao-Su Hu, * Xiao-Zhong Chen, * Guo-Qing Hu, Zhi-Bin Cheng, Yan Sun, Wei-Xiong Li, Yuan-Yuan Chen, Fang-Yun Xie, Shao-Bo Liang, Yong Chen, Ting-Ting Xu, Bin Li, Guo-Xian Long, Si-Yang Wang, Bao-Min Zheng, Ying Guo, Ying Sun, Yan-Ping Mao, Ling-Long Tang, Yu-Ming Chen, Meng-Zhong Liu, Jun Ma

European Journal of Cancer 75 (2017) 150–158



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ScienceDirect

journal homepage: www.ejcancer.com



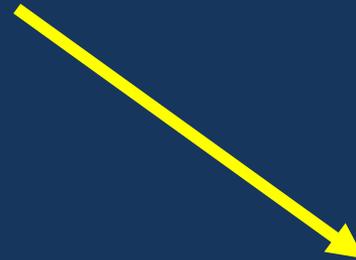
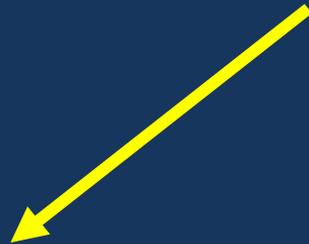
Original Research

Adjuvant chemotherapy in patients with locoregionally advanced nasopharyngeal carcinoma: Long-term results of a phase 3 multicentre randomised controlled trial



Lei Chen ^{a,1}, Chao-Su Hu ^{b,1}, Xiao-Zhong Chen ^{c,1}, Guo-Qing Hu ^{d,1}, Zhi-Bin Cheng ^{e,1}, Yan Sun ^{f,1}, Wei-Xiong Li ^g, Yuan-Yuan Chen ^c, Fang-Yun Xie ^a, Shao-Bo Liang ^h, Yong Chen ^a, Ting-Ting Xu ^b, Bin Li ^c, Guo-Xian Long ^d, Si-Yang Wang ^e, Bao-Min Zheng ^f, Ying Guo ⁱ, Ying Sun ^a, Yan-Ping Mao ^a, Ling-Long Tang ^a, Yu-Ming Chen ^j, Meng-Zhong Liu ^a, Jun Ma ^{a,*}

**Stage III-Stage IVb
N=508**



**Concurrent Chemo RT
70 Gy/ 35# + CDDP Weekly 40mg/m²**

Concurrent Chemo RT

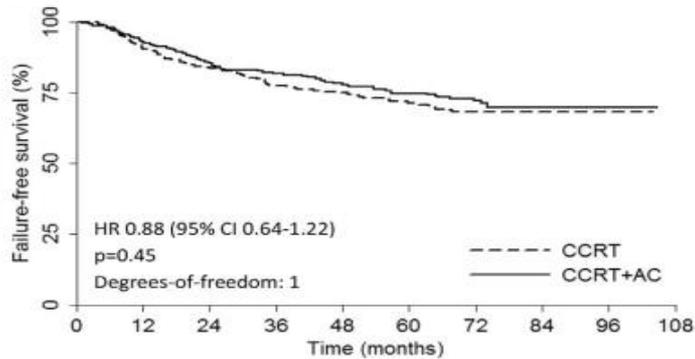


**Adjuvant Chemotherapy
(CDDP 80mg/m²+ 5 FU 800mg/m² D1-D4
q4weeks)**

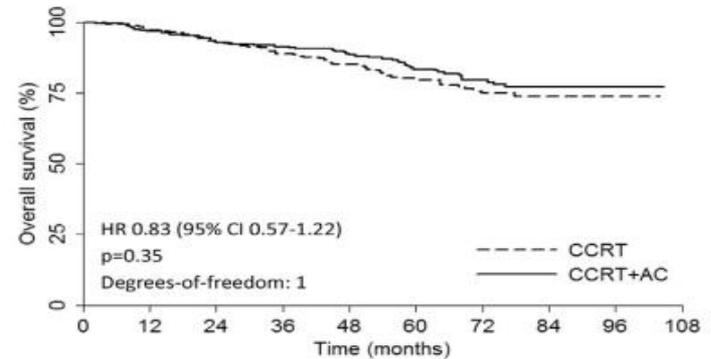
x 3

Lancet Oncol 2012; 13: 163–71

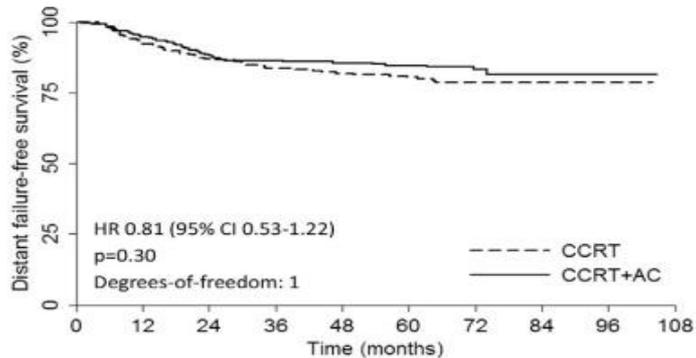
Median follow up 68.4 months



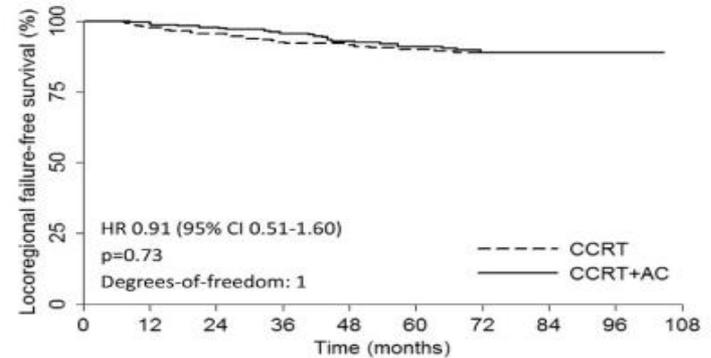
Number at risk		0	12	24	36	48	60	72	84	96	108
CCRT	257	231	212	187	176	150	91	28	8	0	
CCRT+AC	251	230	207	195	182	163	105	28	10	0	



Number at risk		0	12	24	36	48	60	72	84	96	108
CCRT	257	248	236	215	200	168	99	30	8	0	
CCRT+AC	251	239	224	218	203	178	112	31	10	0	



Number at risk		0	12	24	36	48	60	72	84	96	108
CCRT	257	234	216	195	183	159	94	29	8	0	
CCRT+AC	251	231	209	202	189	170	108	30	10	0	



Number at risk		0	12	24	36	48	60	72	84	96	108
CCRT	257	243	230	206	191	158	96	29	8	0	
CCRT+AC	251	238	221	210	196	171	108	29	10	0	

Review Article

Current Role of Chemotherapy in Nonmetastatic Nasopharyngeal Cancer

Tapesh Bhattacharyya ¹, Geethu Babu,² and Cessal Thommachan Kainickal ²

¹*Division of Radiation Oncology, Tata Medical Center, Kolkata, India*

²*Division of Radiation Oncology, Regional Cancer Centre, Trivandrum, India*

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Further progress

CCRT

Risk Adapted Adjuvant
Chemotherapy

Induction Chemotherapy

Adjuvant Chemotherapy
Capecitabine

Analysis of Plasma Epstein-Barr Virus DNA in Nasopharyngeal Cancer After Chemoradiation to Identify High-Risk Patients for Adjuvant Chemotherapy: A Randomized Controlled Trial

Anthony T.C. Chan, Edwin P. Hui, Roger K.C. Ngan, Stewart Y. Tung, Ashley C.K. Cheng, Wai T. Ng, Victor H.F. Lee, Brigitte B.Y. Ma, Hoi C. Cheng, Frank C.S. Wong, Herbert H.F. Loong, Macy Tong, Darren M.C. Poon, Anil T. Ahuja, Ann D. King, Ki Wang, Frankie Mo, Benny C.Y. Zee, K.C. Allen Chan, and Y.M. Dennis Lo

Hong Kong NPC Study Group 0502 Trial (NCT00370890)



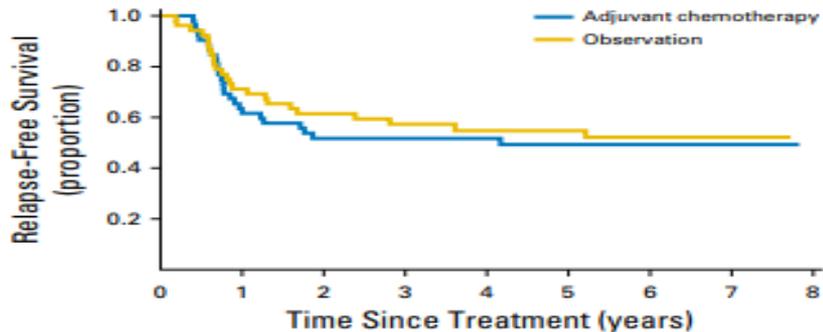
- UICC stage IIB, III, IVA or IVB NPC
- No clinical and radiological evidence of distant metastasis at diagnosis (M0)
- detectable plasma EBV-DNA at 6-8 weeks after completion of RT or CRT
- No clinical evidence of persistent loco-regional disease after primary treatment
- ECOG 0 or 1

R
A
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D
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E

Stratification:
- RT vs CRT
- Stage II/III vs IV

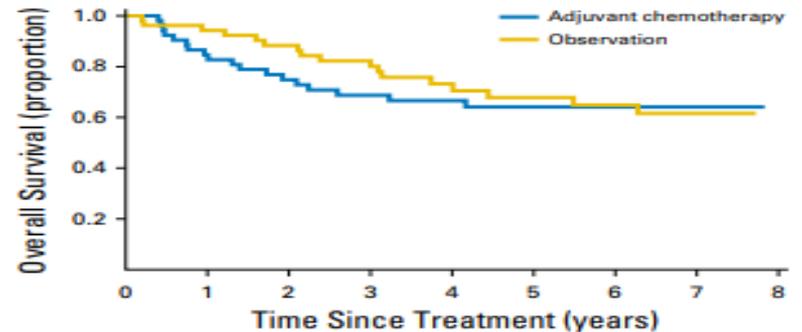
EBV-DNA PET/CT (0 month)	Adjuvant Chemotherapy	EBV-DNA PET/CT (6 months)
Arm A		
✓ ✓	Chemotherapy (Cisplatin-gemcitabine x 6)	✓ ✓
Arm B		
✓ ✓	No chemotherapy	✓ ✓

Efficacy



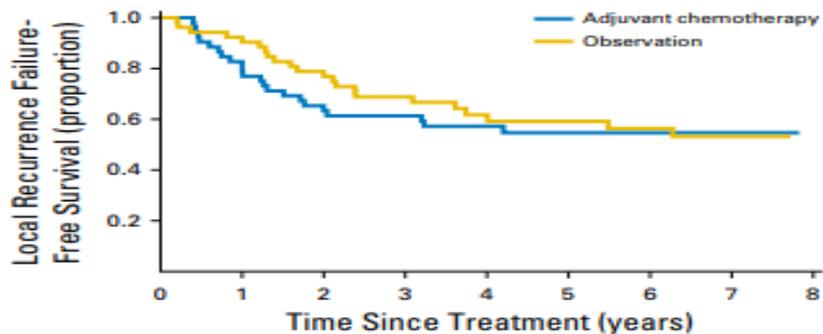
No. at risk

Adjuvant chemotherapy	52	34	27	26	23	19	16	12	6
Observation	52	38	32	27	21	21	18	14	9



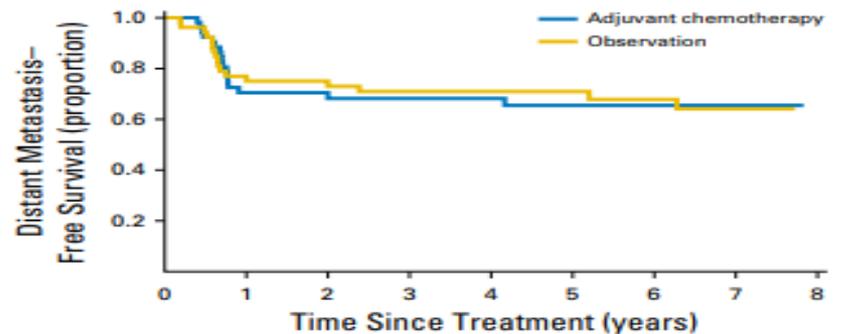
No. at risk

Adjuvant chemotherapy	52	45	38	34	28	24	20	14	8
Observation	52	50	45	39	28	25	22	14	9



No. at risk

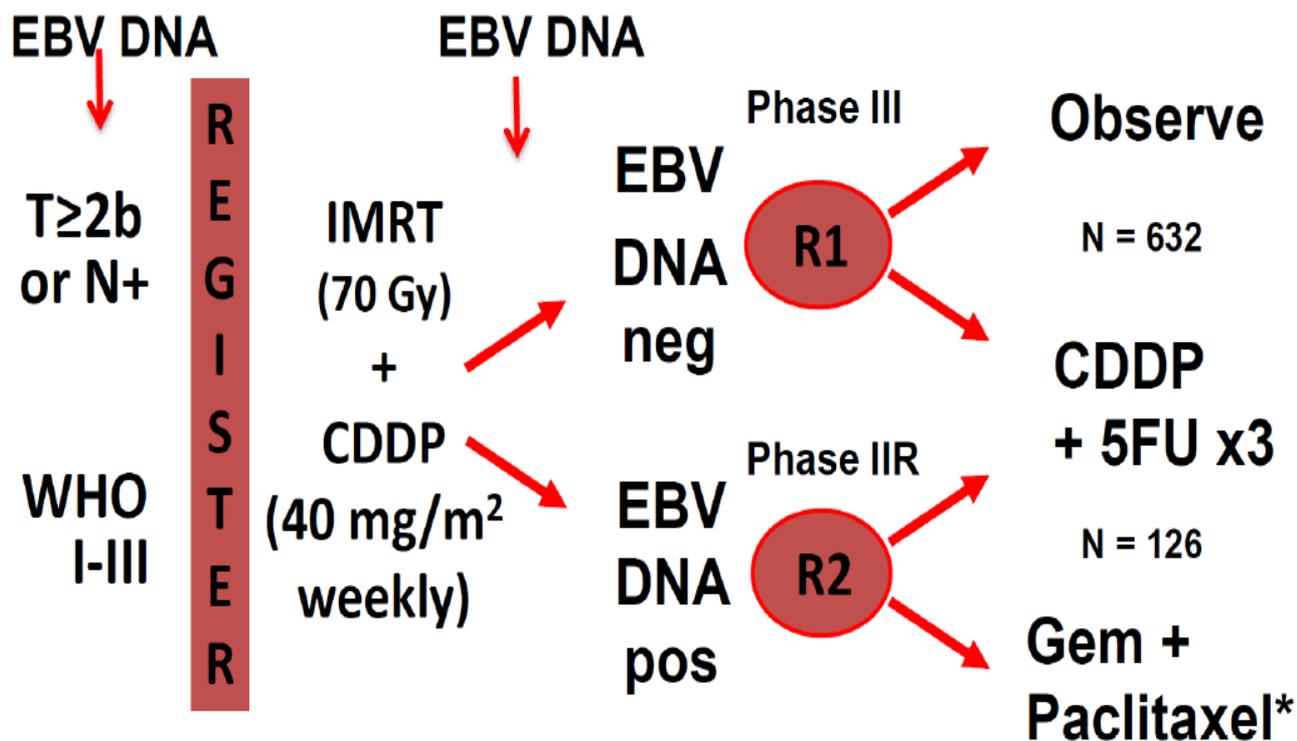
Adjuvant chemotherapy	52	43	33	30	24	20	17	12	6
Observation	52	48	40	33	25	23	20	14	9



No. at risk

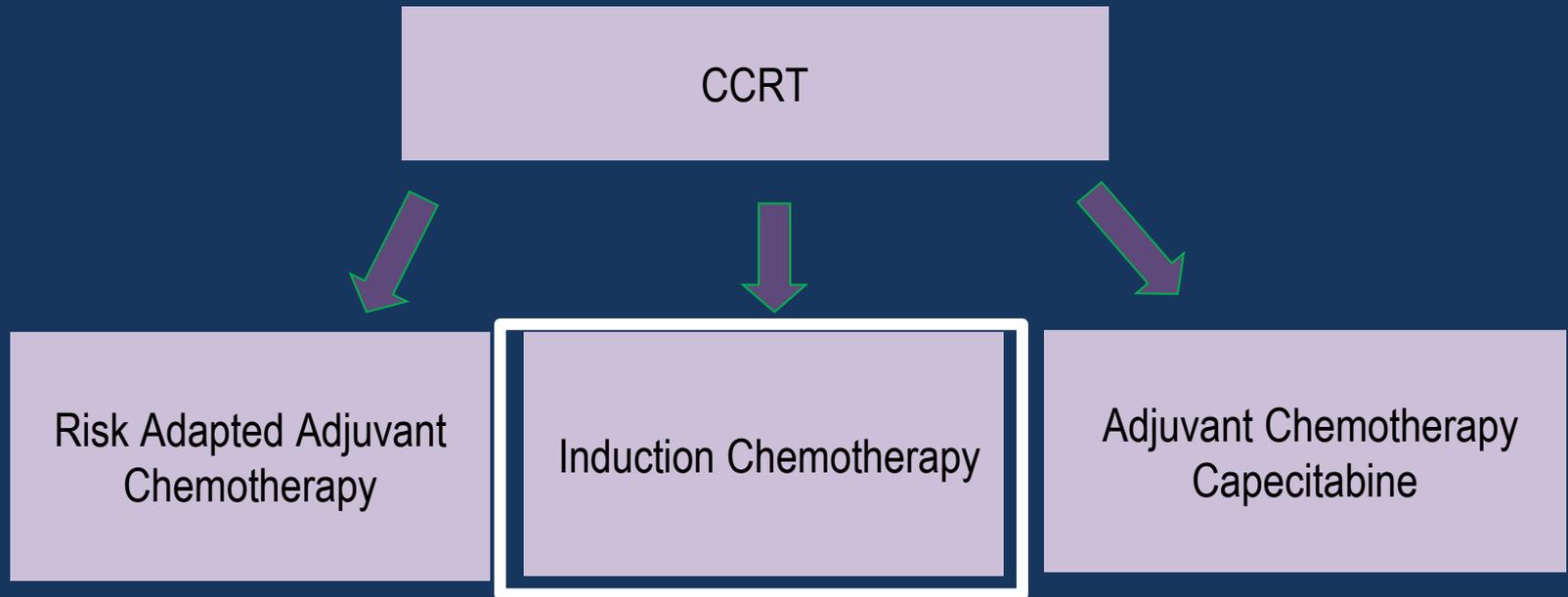
Adjuvant chemotherapy	52	35	31	30	27	23	19	14	8
Observation	52	40	36	32	23	22	19	14	9

NRG HN001- NPC Phase II-IIIR



*Gem 1000 m/m² d1,8 + Paclitaxel 80 mg/m² d1,8 every 21 d X 4 cycles
87 patients enrolled

Further progress



NACT in Ca Nasopharynx

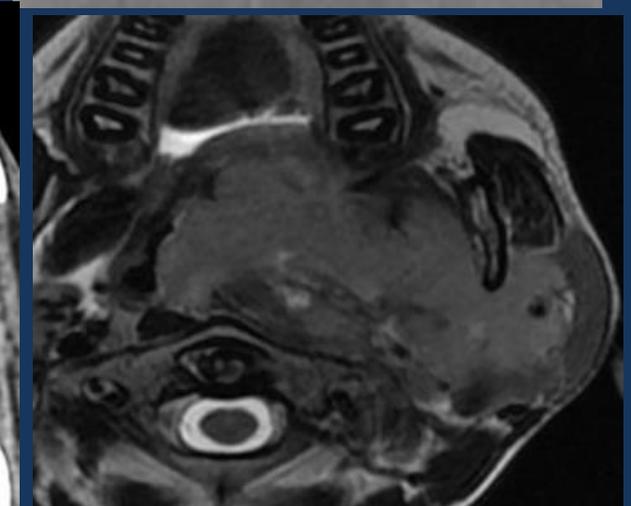
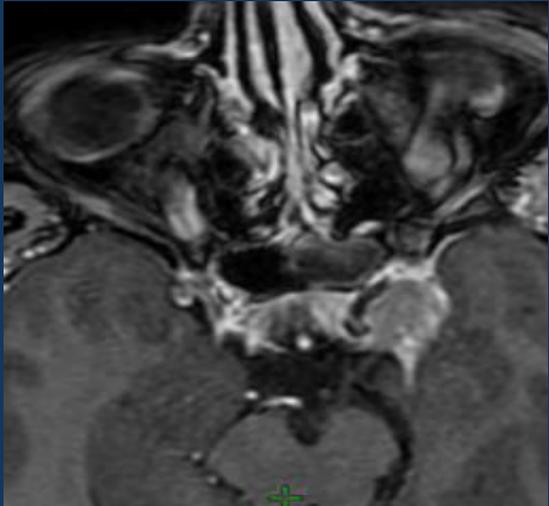
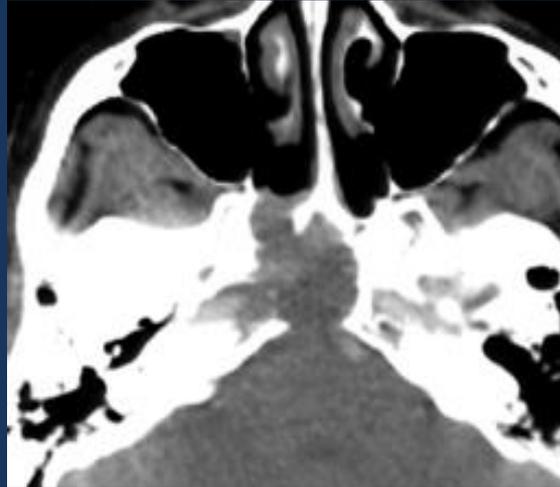
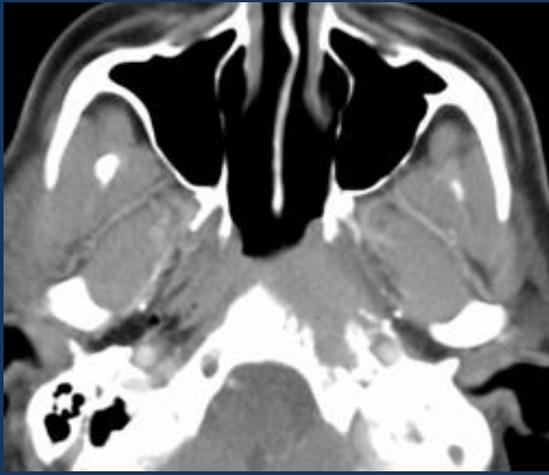
Advantages

- Early eradication of micro metastatic disease
- Facilitate RT planning
- Better tolerated compared to adjuvant chemotherapy

Disadvantages

- Delay in starting the definitive treatment
- Accelerated re population
- Poor compliance to subsequent definitive Chemo RT
- Concerns regarding toxicities

Advanced Ca Nasopharynx



Stage III-Stage IVa Ca Nasopharynx



Concurrent Chemo RT

NACT- 2-3 Cycles



Concurrent Chemo RT

Chapter

Chemotherapy in Nasopharyngeal Carcinoma

Lekha Madhavan Nair, Rejnish Ravi Kumar, Malu Rafi, Farida Nazeer, Kunnambath Ramadas and Kainickal Cessal Thommachan*

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Induction chemotherapy in nasopharyngeal carcinoma- A systematic review of phase III clinical trials

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Outcomes

Author (year)	Control	Intervention	Median follow up	DFS/FFS (Intervention vs. control arm in %)	OS	DMFS
Yang et al [22] (2019) N = 476	CCRT (cisplatin 80 mg/m ²)	CDDP+5FU infusion X 2 cycles →CCRT	82.6 months	5 year DFS 73.4 vs. 63.1 P = 0.007	5 year OS 80.8 vs. 76.8 P = 0.040	5 year DMFS 82.8 vs. 73.1 P = 0.014
Li et al [24] (2019) N = 480	CCRT (Cisplatin 100 mg/m ²)	Docetaxel 60 mg/m ² D1, cisplatin 60 mg/m ² D1, 5FU 600 mg/m ² D1-5 X 3 cycles →CCRT	71.5 months	5 year FFS 77.4 vs. 66.4 P = 0.019	5 year OS 85.6 vs 77.7 P = 0.042	5 year DMFS 88 vs 79.8 P = 0.030
Zhang et al. [25] (2019) N = 480	CCRT (cisplatin 100 mg/m ²)	Cisplatin 80 mg/m ² D1, gemcitabine 1 g/m ² D1, D8 X 3 cycles → CCRT	42.7 months	3 year Recurrence Free Survival 85.3 vs. 76.5 P = 0.001	3 year OS 94.6 vs. 90.3 HR 0.43(0.24–0.77)	3 year DMFS 91.1 VS 84.4 HR 0.43(0.25–0.73)
Frikha et al [26] (2018) N = 83	CCRT Weekly cisplatin 40 mg/m ²	Docetaxel 75 mg/m ² , cisplatin 75 mg/m ² , 5FU 750 mg/m ² D 1–5 X 3 cycles → CCRT	43.1 months	3 year PFS 73.9 vs. 57.2 P = 0.042	3 year OS 86.3 vs. 68.9 P = 0.059	3 year DMFS HR 0.53 P = 0.18
Hong et al. [27] (2018) N = 479	CCRT (cisplatin 80 mg/m ²)	MEPFL regimen (mitomycin 8 mg/m ² D1, Epirubicin 60 mg/m ² D1, Cisplatin 60 mg/m ² D1, 5FU 450 mg/m ² D8. caLV 30 mg/m ² D8) X3 cycles →CCRT	72 months	5 year DFS 61 vs. 50 P = 0.0264	5 year OS 72 vs. 68 P = 0.624	5 year DMFS 76 vs. 71 P = 0.28
Tan et al. [28] (2015) N = 180	CCRT (cisplatin (40 mg/m ² weekly)	Paclitaxel 70 mg/m ² , carboplatin AUC(2.5), Gemcitabine 1 g/m ² D1, D8 X 3 cycles → CCRT	3.2 years	3 year DFS 74.9 vs. 67.4 P = 0.362	3 year OS 94.3 vs. 92.3 P = 0.494	3 year DMFS 83.8 vs. 79.9 P = 0.547

Stage III-Stage IVa Ca Nasopharynx(N-480)

T3-T4N0 excluded
IMRT was mandatory

Concurrent Chemo RT
Cisplatin 100mg/m²d1q3weeks

Primary end point –FFS

**TPFx3cycles - CDDP 60Mg/m²D1,
Docetaxel 60mg/M²d1
5FU 600/M²CID1-D5q3weeks**

Concurrent Chemo RT

Lancet Oncol 2016; 17: 1509–20

Induction chemotherapy plus concurrent chemoradiotherapy versus concurrent chemoradiotherapy alone in locoregionally advanced nasopharyngeal carcinoma: a phase 3, multicentre, randomised controlled trial



Ying Sun*, Wen-Fei Li*, Nian-Yong Chen*, Ning Zhang*, Guo-Qing Hu*, Fang-Yun Xie*, Yan Sun*, Xiao-Zhong Chen, Jin-Gao Li, Xiao-Dong Zhu, Chao-Su Hu, Xiang-Ying Xu, Yuan-Yuan Chen, Wei-Han Hu, Ling Guo, Hao-Yuan Mo, Lei Chen, Yan-Ping Mao, Rui Sun, Ping Ai, Shao-Bo Liang, Guo-Xian Long, Bao-Min Zheng, Xing-Lai Feng, Xiao-Chang Gong, Ling Li, Chun-Ying Shen, Jian-Yu Xu, Ying Guo, Yu-Ming Chen, Fan Zhang, Li Lin, Ling-Long Tang, Meng-Zhong Liu, Jun Ma

IJC International Journal of Cancer



Cancer Therapy and Prevention

Concurrent chemoradiotherapy with/without induction chemotherapy in locoregionally advanced nasopharyngeal carcinoma: Long-term results of phase 3 randomized controlled trial

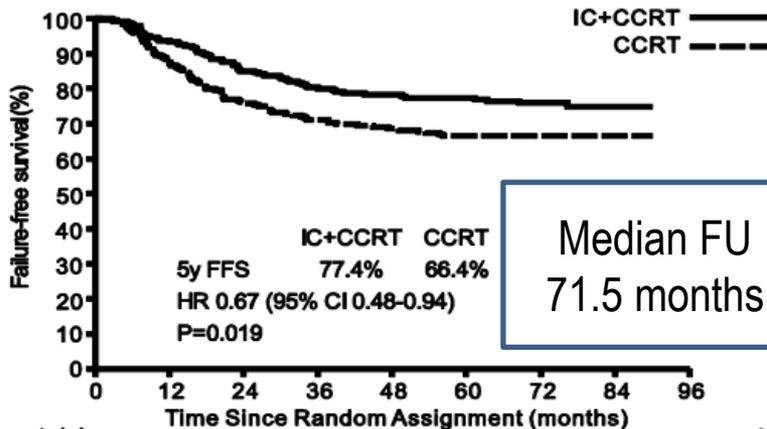
Wen-Fei Li, Nian-Yong Chen, Ning Zhang, Guo-Qing Hu, Fang-Yun Xie, Yan Sun, Xiao-Zhong Chen, Jin-Gao Li, Xiao-Dong Zhu, Chao-Su Hu, Xiang-Ying Xu, Yuan-Yuan Chen, Wei-Han Hu, Ling Guo, Hao-Yuan Mo, Lei Chen, Yan-Ping Mao, Rui Sun, Ping Ai, Shao-Bo Liang, Guo-Xian Long, Bao-Min Zheng, Xing-Lai Feng, Xiao-Chang Gong, Ling Li, Chun-Ying Shen, Jian-Yu Xu, Ying Guo, Yu-Ming Chen, Fan Zhang, Li Lin, Ling-Long Tang, Meng-Zhong Liu, Jun Ma ✉, Ying Sun ✉ ... See fewer authors ^

First published: 06 January 2019 | <https://doi.org/10.1002/ijc.32099> | Cited by: 5

Conflict of interest: The authors declare that they have no competing interests.

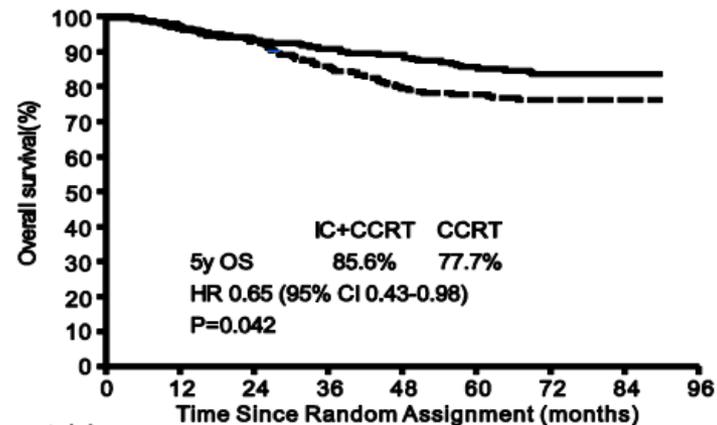
Funding information: Shenzhen Main Luck Pharmaceuticals Inc., Sun Yat-sen University Clinical Research 5010 Program, Grant/Award Number: 2007037; National Natural Science Foundation of China, Grant/Award Number: 81702682; Special support program of Sun Yat-sen University Cancer Center, Grant/Award Number: 16zxtzlc06; Natural Science Foundation of Guangdong Province, Grant/Award Number: 2017A030312003; Health & Medical Collaborative Innovation Project of Guangzhou City, Grant/Award Number: 201803040003; Innovation Team Development Plan of the Ministry of Education, Grant/Award Number: IRT_17R110; Overseas Expertise Introduction Project for Discipline Innovation, Grant/Award Number: 111 Project, B14035

<https://doi.org/10.1002/ijc.32099>



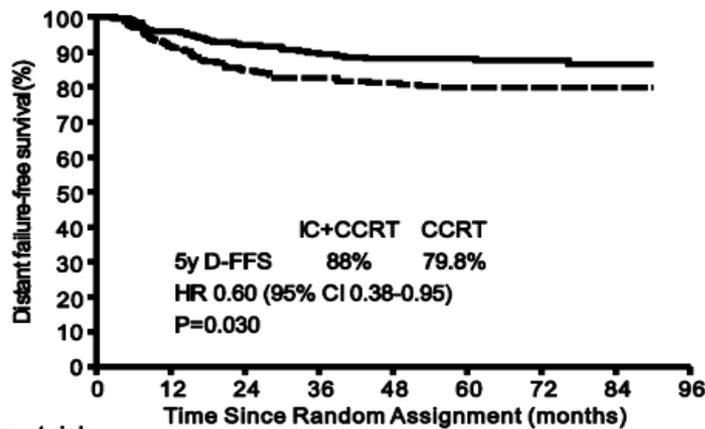
Number at risk

	0	12	24	36	48	60	72	84	96
CCRT 239	210	181	170	161	152	96	17	0	
IC+CCRT 241	225	203	191	181	169	113	20	0	



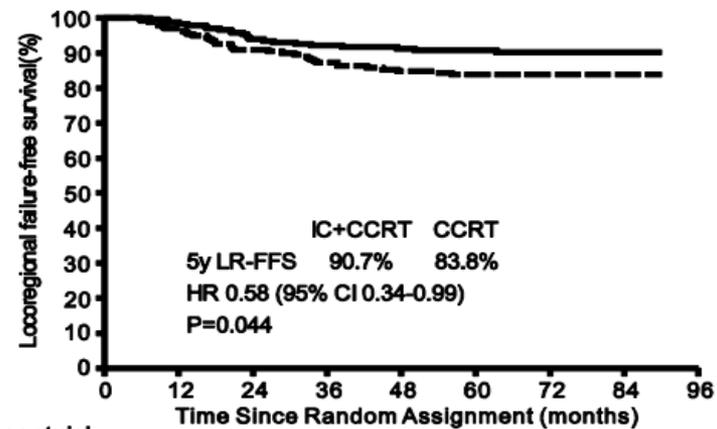
Number at risk

	0	12	24	36	48	60	72	84	96
CCRT 239	231	221	205	187	178	108	20	0	
IC+CCRT 241	234	223	216	207	186	124	21	0	



Number at risk

	0	12	24	36	48	60	72	84	96
CCRT 239	215	197	185	176	168	103	19	0	
IC+CCRT 241	228	214	204	194	179	118	21	0	



Number at risk

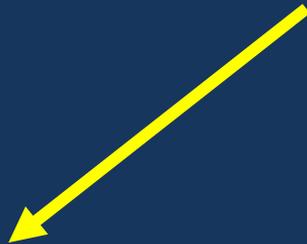
	0	12	24	36	48	60	72	84	96
CCRT 239	225	201	188	171	162	101	18	0	
IC+CCRT 241	231	212	203	193	175	118	20	0	

Toxicity profile

	Induction chemotherapy plus concurrent chemoradiotherapy group (n=239)		Concurrent chemoradiotherapy group (n=238)		p value*	
	Grade 3	Grade 4	Grade 3	Grade 4	Grade 3	Grade 4
Any†	132 (55%)	42 (18%)	125 (53%)	3 (1%)	0.55	<0.0001
Haematological						
Neutropenia	64 (27%)	37 (15%)	16 (7%)	1 (<1%)	<0.0001	<0.0001
Febrile neutropenia	5 (2%)	2 (1%)	0	0	0.061	0.50
Neutropenic infection	1 (<1%)	0	0	0	1.00	..
Leucopenia	86 (36%)	12 (5%)	40 (17%)	1 (<1%)	<0.0001	0.0020
Anaemia	4 (2%)	0	5 (2%)	0	0.75	..
Thrombocytopenia	5 (2%)	1 (<1%)	2 (1%)	0	0.45	1.00
Non-haematological						
Stomatitis (mucositis)	96 (40%)	2 (1%)	82 (34%)	2 (1%)	0.20	1.00
Vomiting	52 (22%)	4 (2%)	45 (19%)	0	0.44	0.12
Nausea	46 (19%)	4 (2%)	40 (17%)	0	0.49	0.12
Dry mouth	13 (5%)	..‡	13 (5%)	..‡	0.99	..
Dermatitis	8 (3%)	1 (<1%)	10 (4%)	0	0.62	1.00
Oesophagitis, dysphagia, or odynophagia	5 (2%)	0	9 (4%)	0	0.27	..
Hepatotoxicity	7 (3%)	0	2 (1%)	0	0.18	..
Allergic reaction	2 (1%)	0	0	0	0.50	..

Data are n or n (%). *p values were calculated with the χ^2 test (or Fisher's exact test). †No grade 3-4 nephrotoxicity, ototoxicity, or neurotoxicity was recorded. ‡According to the Common Terminology Criteria for Adverse Events (version 3.0) dry mouth has only grade 1-3.

Stage III-Stage IVa
Ca Nasopharynx(n=480)



Concurrent Chemo RT CCRT
CCRT- Cisplatin 100mg/m² d1q3weeks

IC-3 cycles-CDDP 80 mg/m² D1 plus
Gemcitbaine 1000mg/m² D1& D8 q3

Primary end point –RFS



Concurrent Chemo RT

N Engl J Med 2019;381:1124-35.

ORIGINAL ARTICLE

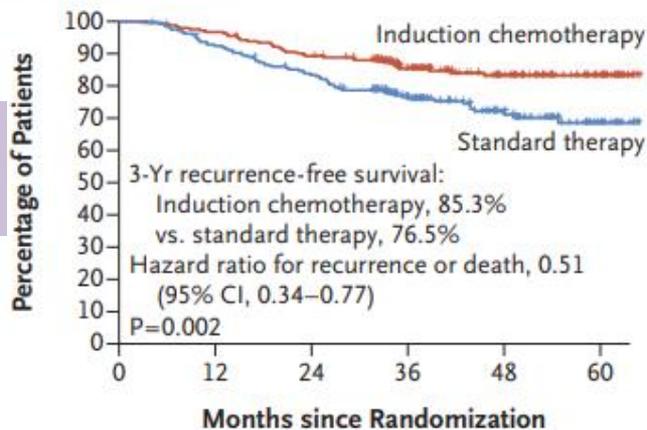
Gemcitabine and Cisplatin Induction Chemotherapy in Nasopharyngeal Carcinoma

Y. Zhang, L. Chen, G.-Q. Hu, N. Zhang, X.-D. Zhu, K.-Y. Yang, F. Jin, M. Shi, Y.-P. Chen, W.-H. Hu, Z.-B. Cheng, S.-Y. Wang, Y. Tian, X.-C. Wang, Yan Sun, J.-G. Li, W.-F. Li, Y.-H. Li, L.-L. Tang, Y.-P. Mao, G.-Q. Zhou, R. Sun, X. Liu, R. Guo, G.-X. Long, S.-Q. Liang, L. Li, J. Huang, J.-H. Long, J. Zang, Q.-D. Liu, L. Zou, Q.-F. Su, B.-M. Zheng, Y. Xiao, Y. Guo, F. Han, H.-Y. Mo, J.-W. Lv, X.-J. Du, C. Xu, N. Liu, Y.-Q. Li, M.L.K. Chua, F.-Y. Xie, Ying Sun, and J. Ma

Event	Induction Chemotherapy (N = 239)		Standard Therapy (N = 237)	
	Grade 1 or 2	Grade 3 or 4	Grade 1 or 2	Grade 3 or 4
	<i>number of patients with event (percent)</i>			
Any acute adverse event	58 (24.3)	181 (75.7)	105 (44.3)	132 (55.7)
Leukopenia	168 (70.3)	63 (26.4)	178 (75.1)	48 (20.3)
Neutropenia	135 (56.5)	67 (28.0)	147 (62.0)	25 (10.5)
Febrile neutropenia	0	1 (0.4)	0	0
Neutropenic infection	0	0	0	0
Anemia	178 (74.5)	23 (9.6)	157 (66.2)	2 (0.8)
Thrombocytopenia	91 (38.1)	27 (11.3)	54 (22.8)	3 (1.3)
Mucositis	139 (58.2)	69 (28.9)	154 (65.0)	76 (32.1)
Vomiting	85 (35.6)	54 (22.6)	52 (21.9)	33 (13.9)
Nausea	176 (73.6)	55 (23.0)	188 (79.3)	33 (13.9)

Primary end point

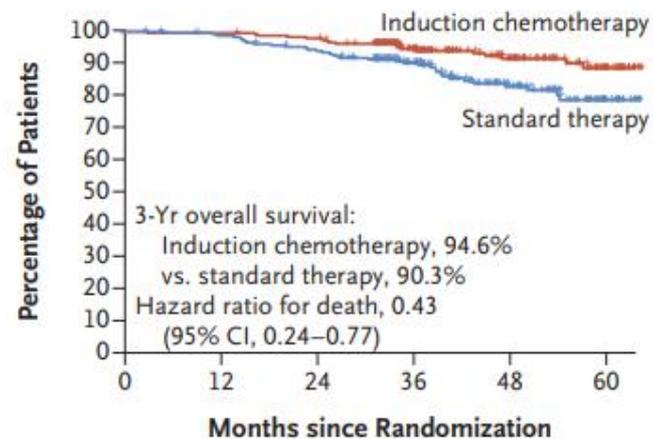
Recurrence-free Survival



No. at Risk

Induction chemotherapy	242	234	215	146	93	35
Standard therapy	238	217	194	130	73	26

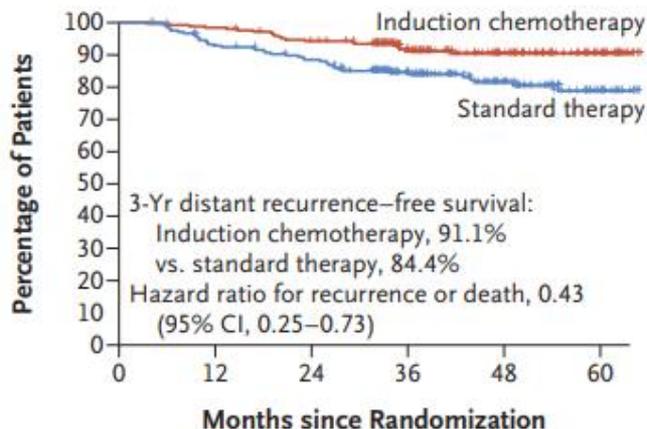
Overall Survival



No. at Risk

Induction chemotherapy	242	241	236	162	100	36
Standard therapy	238	232	219	152	87	29

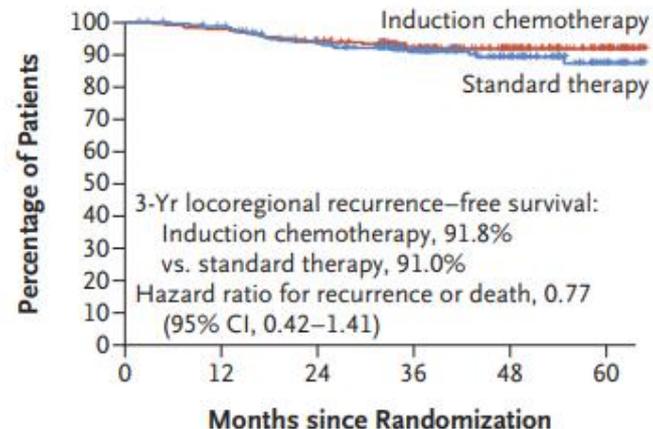
Distant Recurrence-free Survival



No. at Risk

Induction chemotherapy	242	238	226	154	96	35
Standard therapy	238	217	204	140	80	28

Locoregional Recurrence-free Survival



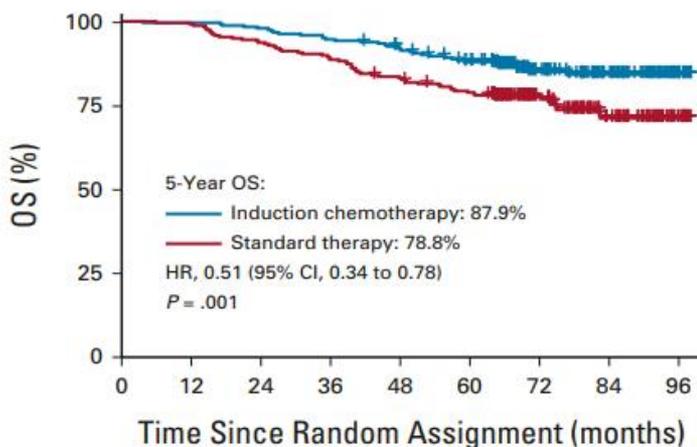
No. at Risk

Induction chemotherapy	242	237	225	152	97	36
Standard therapy	238	230	206	141	81	27

Final Overall Survival Analysis of Gemcitabine and Cisplatin Induction Chemotherapy in Nasopharyngeal Carcinoma: A Multicenter, Randomized Phase III Trial

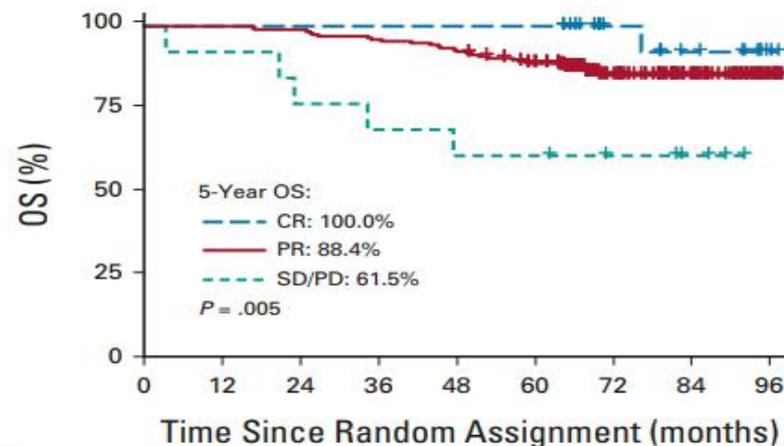
Yuan Zhang, MD, PhD¹; Lei Chen, MD, PhD¹; Guo-Qing Hu, MD²; Ning Zhang, MD³; Xiao-Dong Zhu, MD, PhD⁴; Kun-Yu Yang, MD⁵; Feng Jin, MD⁶; Mei Shi, MD, PhD⁷; Yu-Pei Chen, MD¹; Wei-Han Hu, MD¹; Zhi-Bin Cheng, MD⁸; Si-Yang Wang, MD⁹; Ye Tian, MD¹⁰; Xi-Cheng Wang, MD¹¹; Yan Sun, MD, PhD¹²; Jin-Gao Li, MD¹³; Wen-Fei Li, MD, PhD¹; Yu-Hong Li, MD¹⁴; Yan-Ping Mao, MD, PhD¹; Guan-Qun Zhou, MD, PhD¹; Rui Sun, MD¹; Xu Liu, MD, PhD¹; Rui Guo, MD, PhD¹; Guo-Xian Long, MD, PhD²; Shao-Qiang Liang, MD³; Ling Li, MD, PhD⁴; Jing Huang, MD, PhD⁵; Jin-Hua Long, MD⁶; Jian Zang, MD⁷; Qiao-Dan Liu, MD, PhD⁹; Li Zou, MD, PhD¹⁰; Qiong-Fei Su, MD¹¹; Bao-Min Zheng, MD, PhD¹²; Yun Xiao, MD¹³; Ying Guo, PhD¹⁵; Fei Han, MD, PhD¹; Hao-Yuan Mo, MD¹⁶; Jia-Wei Lv, MD¹; Xiao-Jing Du, MD, PhD¹; Cheng Xu, MD, PhD¹; Na Liu, MD, PhD¹; Ying-Qin Li, MD, PhD¹; Fang-Yun Xie, MD¹; Ying Sun, MD, PhD¹; Jun Ma, MD¹; and Ling-Long Tang, MD, PhD¹

With a median follow-up of 69.8 months



No. at risk:

Induction chemotherapy	242	241	236	228	217	202	114	69	10
Standard therapy	238	234	221	209	195	183	105	53	10

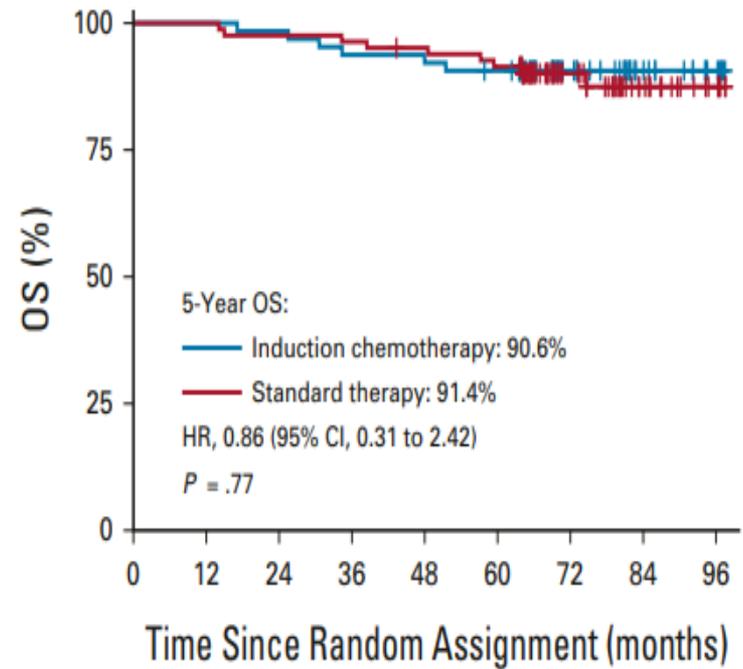
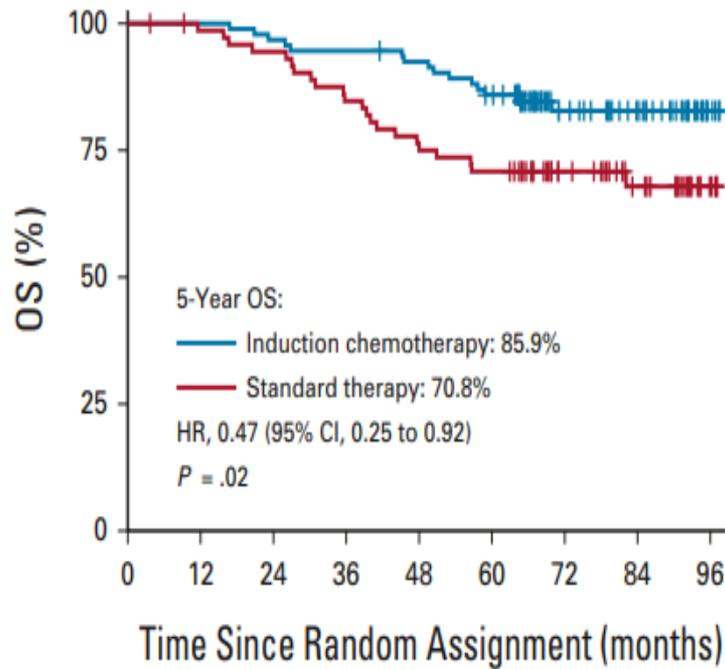


No. at risk:

CR	24	24	24	24	24	2	13	9	2
PR	202	202	199	192	182	168	93	55	8
SD/PD	13	12	10	9	8	8	6	3	0

EBV DNA >4000

EBV DNA <4000



No. at risk:

Induction chemotherapy	93	93	90	88	85	78	43	32	4
Standard therapy	74	71	68	61	55	51	33	22	4

No. at risk:

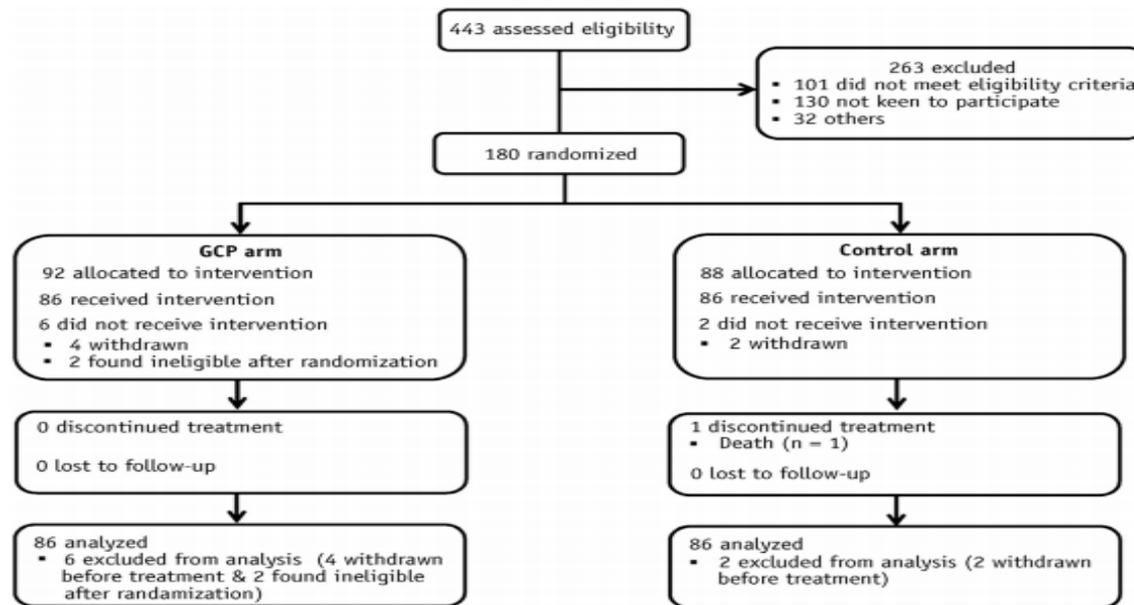
Induction chemotherapy	64	64	63	60	59	57	31	16	5
Standard therapy	82	82	80	79	77	74	37	17	6

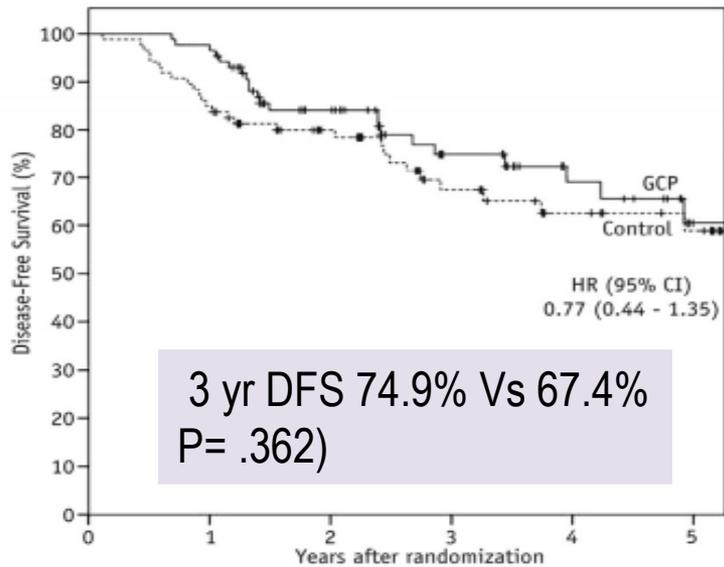
Clinical Investigation

Concurrent Chemo-Radiation With or Without Induction Gemcitabine, Carboplatin, and Paclitaxel: A Randomized, Phase 2/3 Trial in Locally Advanced Nasopharyngeal Carcinoma



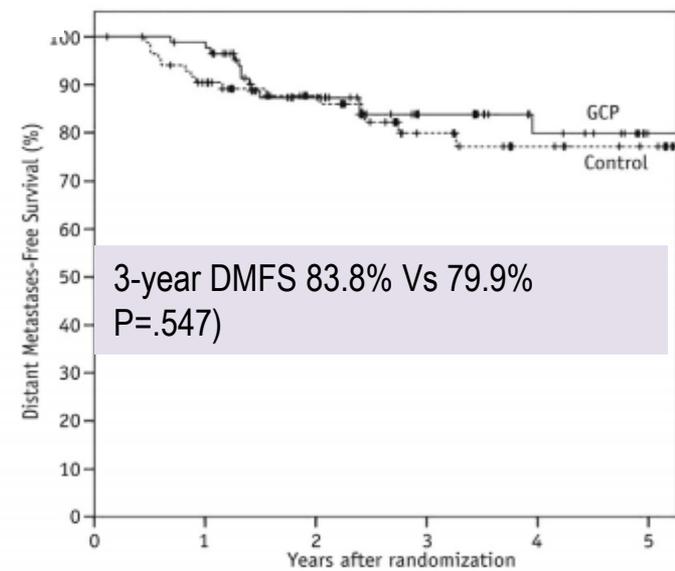
Terence Tan, FRCR,* Wan-Teck Lim, MRCP,[†] Kam-Weng Fong, FRCR,* Shie-Lee Cheah, FRCR,* Yoke-Lim Soong, FRCR,* Mei-Kim Ang, MRCP,[†] Quan-Sing Ng, MRCP,[†] Daniel Tan, MRCP,[†] Whee-Sze Ong, MAppStats,[‡] Sze-Huey Tan, MSc,[‡] Connie Yip, FRCR,* Daniel Quah, FRCR,* Khee-Chee Soo, MD,[§] and Joseph Wee, FRCR*





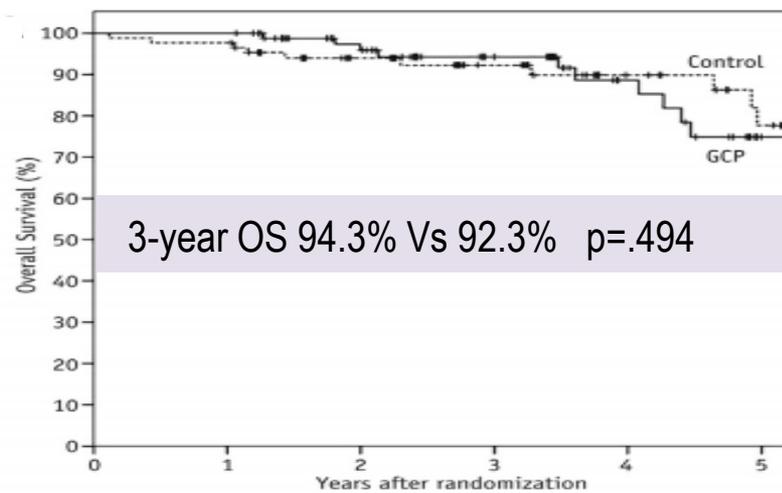
Number at risk

Control	86	73	52	32	21	16
GCP	86	84	57	32	20	7



Number at risk

Control	86	73	52	32	21	16
GCP	86	84	57	32	20	7



Number at risk

Control	86	84	61	43	28	18
GCP	86	86	66	41	26	10

Criteria	Cao	Ying Sun	Terence Tan	GORTEC	J.Ma	Hong
Inclusion criteria	Stage III & IV except (T3N0-N1)	Stage III & IV except (T3, T4N0)	Stage III & IV	Stage III & IV	Stage III & IV	Stage IVa & IV b
Age	18-60	Up to 59 yrs	Above 18 yrs	-	18-64 yrs	Less than 70 yrs
NACT	2 cycles of PF	3 cycles - Modified TPF	3 cycles Carbo+gem+pacli	3 cycles of TPF	3 cycles of Cis Gem	MEPFL X3 cycles
OS Benefit	Yes	Yes	No	No	Yes	No
Toxicity	Grade 3/4 toxicity 63.4% Vs 48.3% P=0.001	Grade IV acute toxicity more p=0.0001	More	Same	More	More
FFS	Yes	Yes	No	Yes	Yes	Yes

Best IC regime?

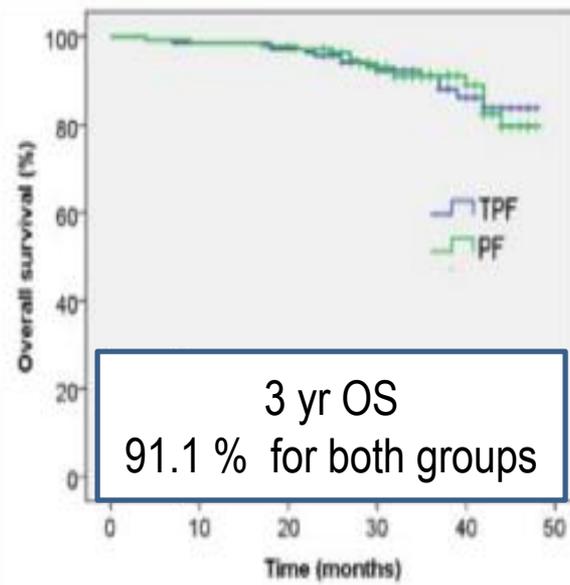
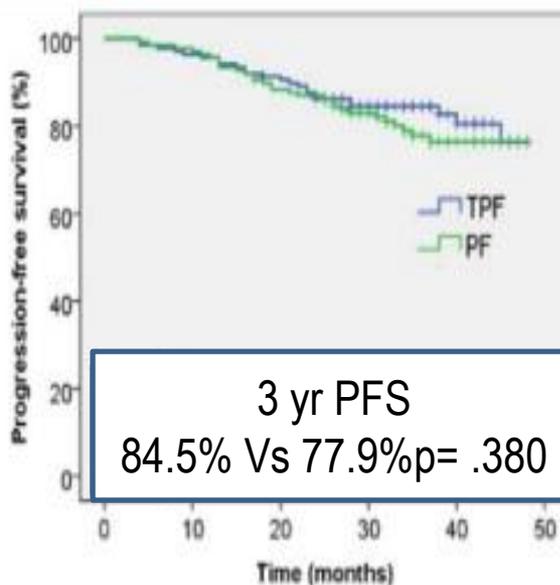
Cisplatin and Fluorouracil Induction Chemotherapy With or Without Docetaxel in Locoregionally Advanced Nasopharyngeal Carcinoma^{1,2,3,4}



Ting Jin^{*,†,5}, Wei-feng Qin^{†,‡,5}, Feng Jiang^{†,‡}, Qi-feng Jin^{†,‡}, Qi-chun Wei⁵, Yong-shi Jia⁵, Xiao-nan Sun⁶, Wen-feng Li^{**} and Xiao-zhong Chen^{†,‡}

*Key Laboratory of Head & Neck Cancer Translational Research of Zhejiang Province, Hangzhou, Zhejiang 310022, People's Republic of China; [†]Department of Radiation Oncology, Zhejiang Cancer Hospital, Hangzhou, Zhejiang 310022, People's Republic of China; [‡]Key Laboratory of Radiation Oncology in Zhejiang Province, Hangzhou, Zhejiang 310022, People's Republic of China; ⁵Department of Radiation Oncology, Key Laboratory of Cancer Prevention and Intervention, China National Ministry of Education, The Second Affiliated Hospital, School of Medicine, Zhejiang University, Hangzhou, Zhejiang 310009, People's Republic of China; ⁶Department of Radiation Oncology, Zhejiang Provincial People's Hospital, Hangzhou, Zhejiang 310014, People's Republic of China; ⁶Department of Radiation Oncology, Sir Run Run Shaw Hospital, College of Medicine, Zhejiang University, Hangzhou, Zhejiang 310000, People's Republic of China; ^{**}Department of Chemoradiation Oncology, The First Affiliated Hospital of Wenzhou Medical University, Wenzhou, Zhejiang 325000, People's Republic of China

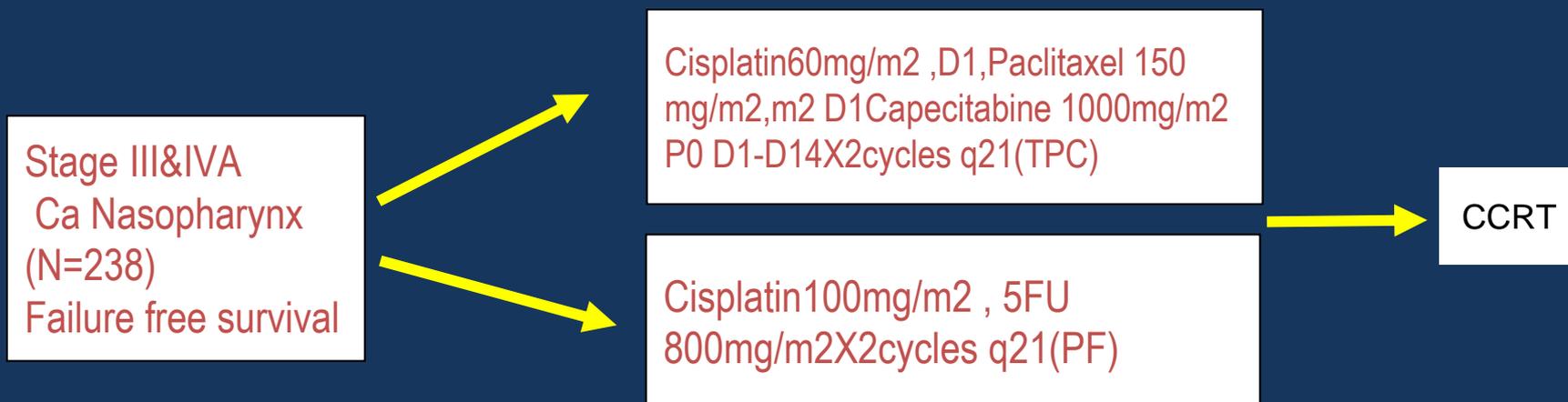
3Cycles of TPF Vs 3cycles PF
 Non inferiority trial
 TPF- CDDP 75 mg/M2 D1
 Docetaxel 75/m2 D1
 5FU 600/M2 D1-5 CI
 PF
 CDDP 100MG/M2 D1
 5FU 800/M2 IV D1-5 CI
 N= 276
 All patients received IMRT
 18- 70 yrs
 Primary end point PFS



Effect of Induction Chemotherapy With Paclitaxel, Cisplatin, and Capecitabine vs Cisplatin and Fluorouracil on Failure-Free Survival for Patients With Stage IVA to IVB Nasopharyngeal Carcinoma

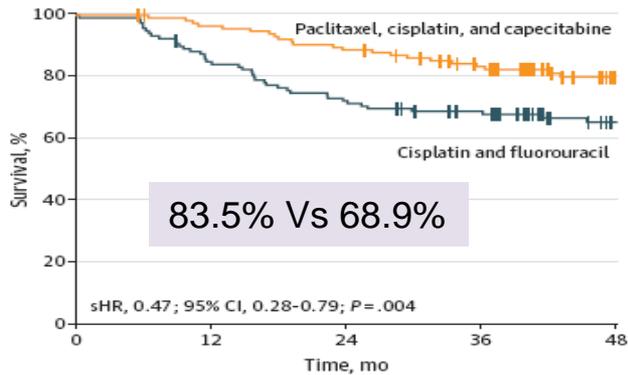
A Multicenter Phase 3 Randomized Clinical Trial

Wang-Zhong Li, MD; Xing Lv, MD; Dan Hu, MS; Shu-Hui Lv, MS; Guo-Ying Liu, MD; Hu Liang, MD; Yan-Fang Ye, MS; Wen Yang, MD; Han-Xiong Zhang, MD; Tai-Ze Yuan, MD; De-Shen Wang, MD; Nian Lu, MD; Liang-Ru Ke, MD; Wu-Bing Tang, MD; Li-Hua Tong, MS; Zhi-Jie Chen, MS; Ting Liu, MS; Ka-Jia Cao, MD; Hao-Yuan Mo, MD; Ling Guo, MD; Chong Zhao, MD; Ming-Yuan Chen, MD; Qiu-Yan Chen, MD; Pei-Yu Huang, MD; Rui Sun, MD; Fang Qiu, MD; Dong-Hua Luo, MD; Lin Wang, MD; Yi-Jun Hua, MD; Lin-Quan Tang, MD; Chao-Nan Qian, MD; Hai-Qiang Mai, MD; Xiang Guo, MD; Yan-Qun Xiang, MD; Wei-Xiong Xia, MD



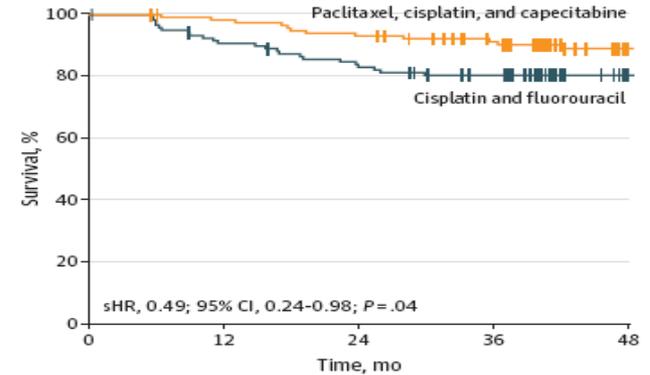
Results- median FU 48.4 months

Failure-free survival



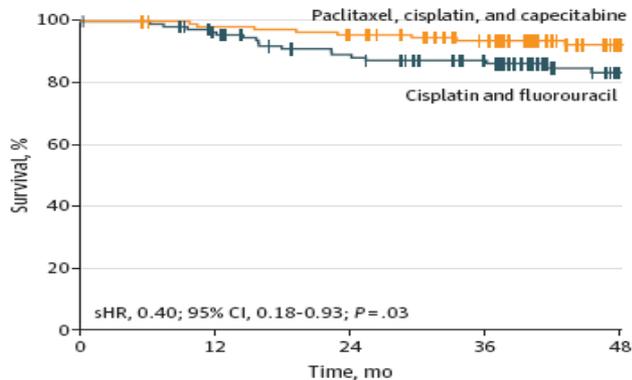
No. at risk	0	12	24	36	48
Cisplatin and fluorouracil	120	101	86	76	44
Paclitaxel, cisplatin, and capecitabine	118	112	103	89	51

Distant metastasis-free survival



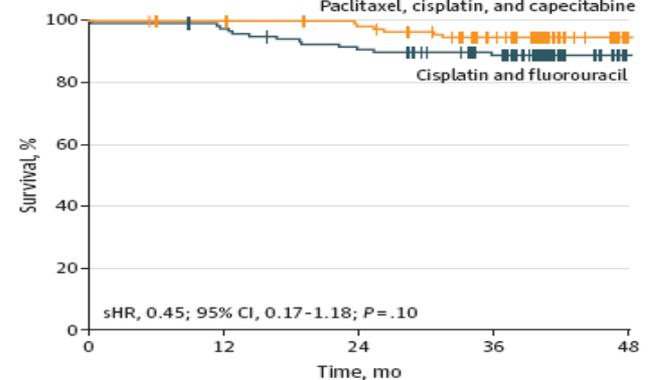
No. at risk	0	12	24	36	48
Cisplatin and fluorouracil	120	107	97	88	52
Paclitaxel, cisplatin, and capecitabine	118	114	108	96	55

Locoregional relapse-free survival



No. at risk	0	12	24	36	48
Cisplatin and fluorouracil	120	109	96	85	49
Paclitaxel, cisplatin, and capecitabine	118	114	109	95	54

Overall survival



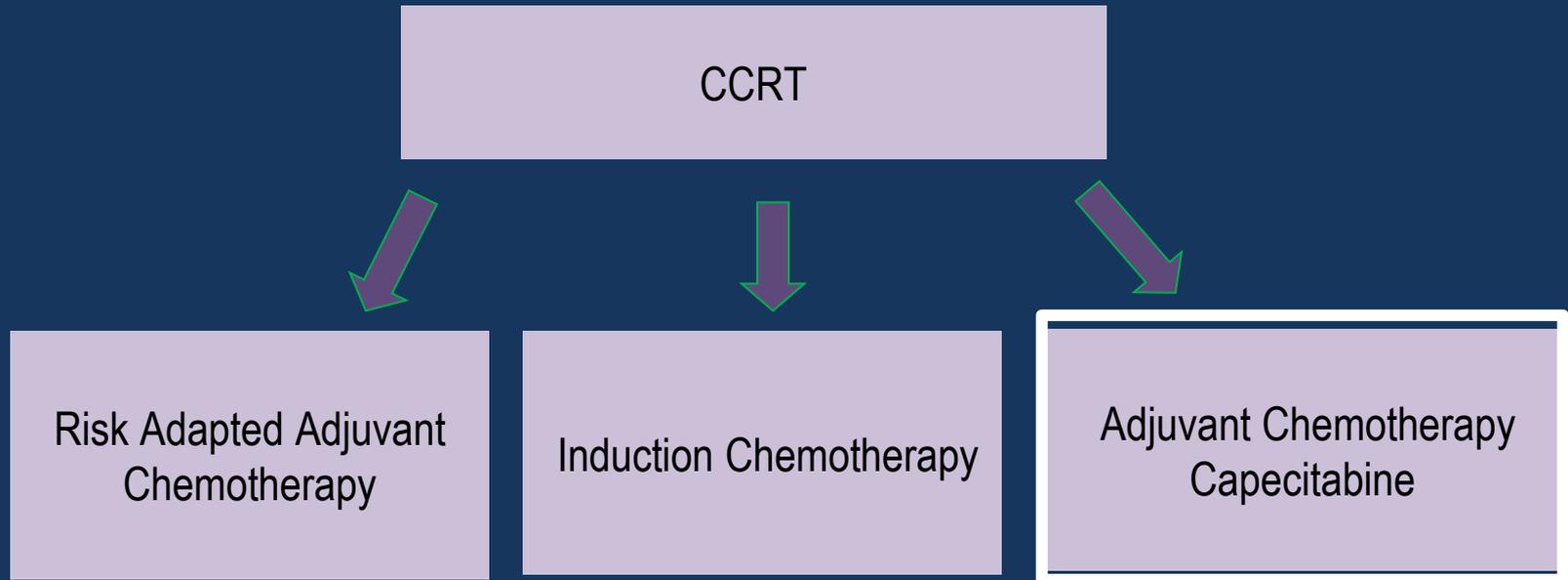
No. at risk	0	12	24	36	48
Cisplatin and fluorouracil	120	116	107	94	57
Paclitaxel, cisplatin, and capecitabine	118	116	112	97	57

Toxicity – Similar

Adverse event	Patients, No. (%)					
	Paclitaxel, cisplatin, and capecitabine (n = 118)			Cisplatin and fluorouracil (n = 120)		
	Grade 0-1	Grade 2	Grade 3-4	Grade 0-1	Grade 2	Grade 3-4
Acute hematologic toxicity						
Leukopenia	60 (50.8)	40 (33.9)	18 (15.3)	65 (54.2)	28 (31.7)	17 (14.2)
Neutropenia	73 (61.9)	30 (25.4)	15 (12.7)	70 (58.3)	28 (23.3)	22 (18.3)
Anemia	68 (57.6)	37 (31.4)	13 (11.0)	70 (58.3)	39 (32.5)	11 (9.2)
Thrombocytopenia	108 (91.5)	8 (6.8)	2 (1.7)	105 (87.5)	8 (6.8)	2 (1.7)
Acute nonhematologic toxicity						
Dry mouth	80 (67.8)	31 (26.3)	7 (5.9)	73 (60.8)	35 (29.2)	12 (10.0)
Mucositis	53 (44.9)	32 (27.1)	33 (28.0)	48 (40.0)	38 (31.7)	34 (28.3)
Dermatitis	94 (79.7)	22 (18.6)	2 (1.7)	87 (72.5)	29 (24.2)	4 (3.3)
Diarrhea	111 (94.1)	5 (4.2)	2 (1.7)	112 (93.3)	6 (5.0)	2 (1.7)
Nausea	62 (52.5)	38 (32.2)	18 (15.3)	63 (52.5)	32 (26.7)	25 (20.8)
Vomiting	71 (60.2)	25 (21.2)	22 (18.6)	66 (55.0)	35 (29.2)	19 (15.8)
Hepatotoxicity	96 (81.4)	19 (16.1)	3 (2.5)	101 (84.2)	16 (13.3)	3 (2.5)
Nephrotoxicity ^a	112 (94.9)	6 (5.1)	0	112 (93.3)	7 (5.8)	1 (0.8)
Hand-foot syndrome	116 (98.3)	2 (1.7)	0	119 (99.2)	1 (0.8)	0
Allergic reaction	112 (94.9)	5 (4.2)	1 (0.8)	117 (97.5)	3 (2.5)	0
Weight loss	91 (77.1)	25 (21.2)	2 (1.7)	90 (75.0)	27 (22.5)	3 (2.5)

3 drug NACT is not superior to 2 drug

Further progress



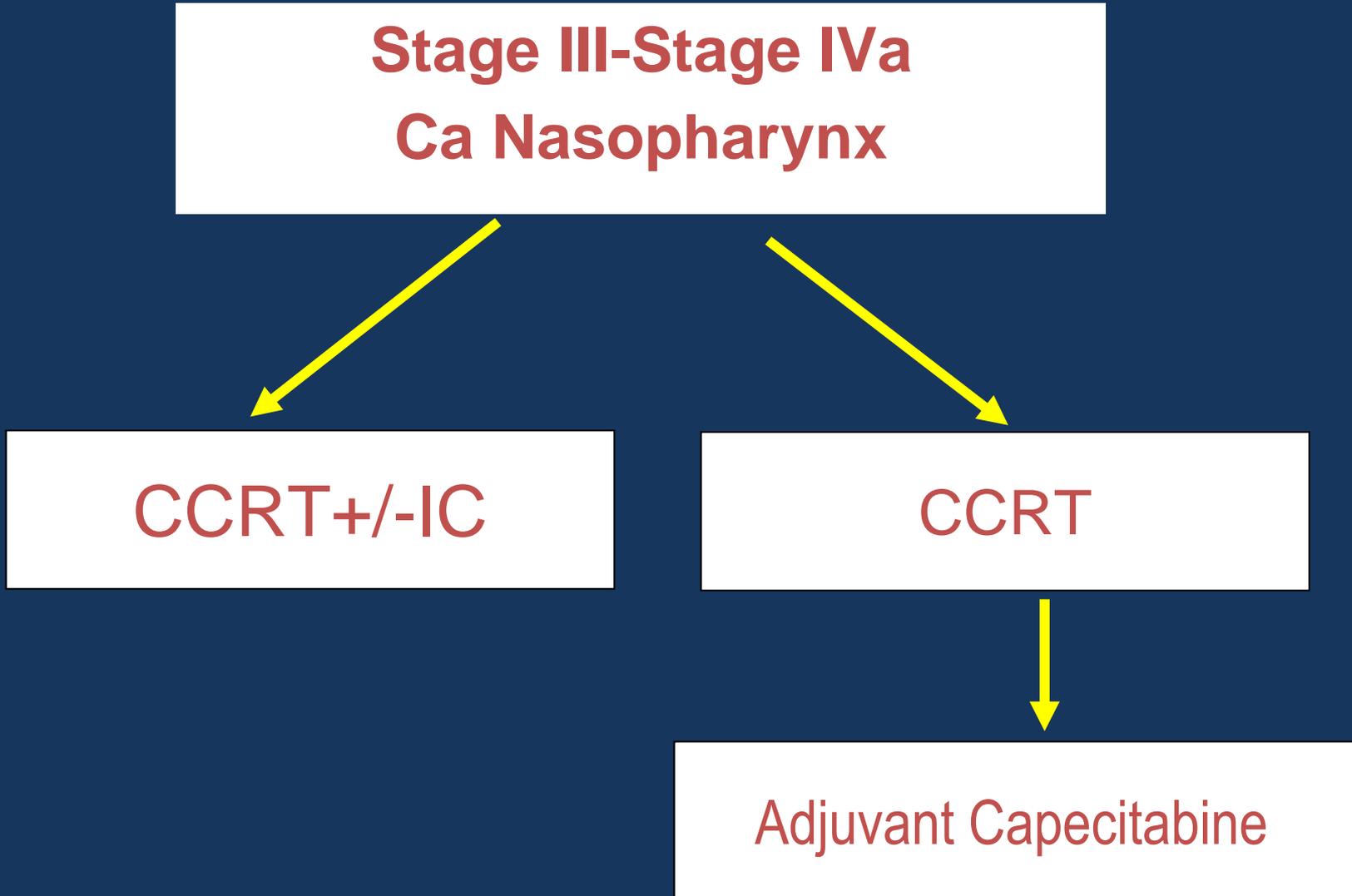
Adjuvant Chemotherapy

**Stage III-Stage IVa
Ca Nasopharynx**

CCRT+/-IC

CCRT

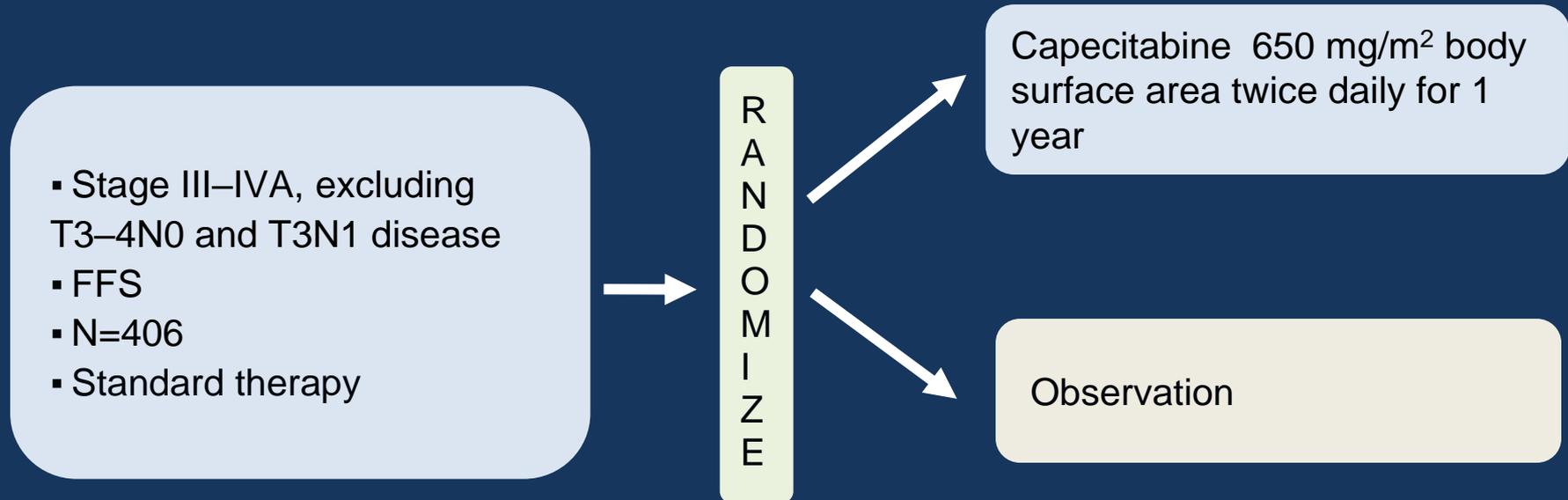
Adjuvant Capecitabine





Metronomic capecitabine as adjuvant therapy in locoregionally advanced nasopharyngeal carcinoma: a multicentre, open-label, parallel-group, randomised, controlled, phase 3 trial

Yu-Pei Chen*, Xu Liu*, Qin Zhou*, Kun-Yu Yang*, Feng Jin*, Xiao-Dong Zhu*, Mei Shi*, Guo-Qing Hu*, Wei-Han Hu*, Yan Sun, Hong-Fen Wu, Hui Wu, Qin Lin, Hui Wang, Ye Tian, Ning Zhang, Xi-Cheng Wang, Liang-Fang Shen, Zheng-Zheng Liu, Jing Huang, Xiu-Ling Luo, Ling Li, Jian Zang, Qi Mei, Bao-Min Zheng, Dan Yue, Jing Xu, San-Gang Wu, Yan-Xia Shi, Yan-Ping Mao, Lei Chen, Wen-Fei Li, Guan-Qun Zhou, Rui Sun, Rui Guo, Yuan Zhang, Cheng Xu, Jia-Wei Lv, Ying Guo, Hui-Xia Feng, Ling-Long Tang†, Fang-Yun Xie†, Ying Sun†, Jun Ma†



HEAD AND NECK CANCER

Adjuvant capecitabine in locoregionally advanced nasopharyngeal carcinoma: A multicenter randomized controlled phase III trial.



Check for updates

[Jingjing Miao](#), [Lin Wang](#), [Sze Huey Tan](#), [Jin-Gao Li](#), [Junlin Yi](#), [Ye Zhang](#), [Xiaochang Gong](#), [Xiang Yanqun](#), [Qiu-Yan Chen](#), [Mingyuan Chen](#), [Xing Lv](#), [Weixiong Xia](#), [Lin-Quan Tang](#), [Xiao-Wu Deng](#), [Xiang Guo](#), [Hai-Qiang Mai](#), [Fei Han](#), [Melvin Lee Kiang Chua](#), [Chong Zhao](#)

- T3-4N2 or T1-4N3
- Pre-treatment plasma EBV DNA >20,000 copy/ml
- GTVnx of >30 cm³
- SUVmax of >10.0 by ¹⁸FDG PET-CT in primary or node, with any larger than 4 cm
- FFS
- N=180



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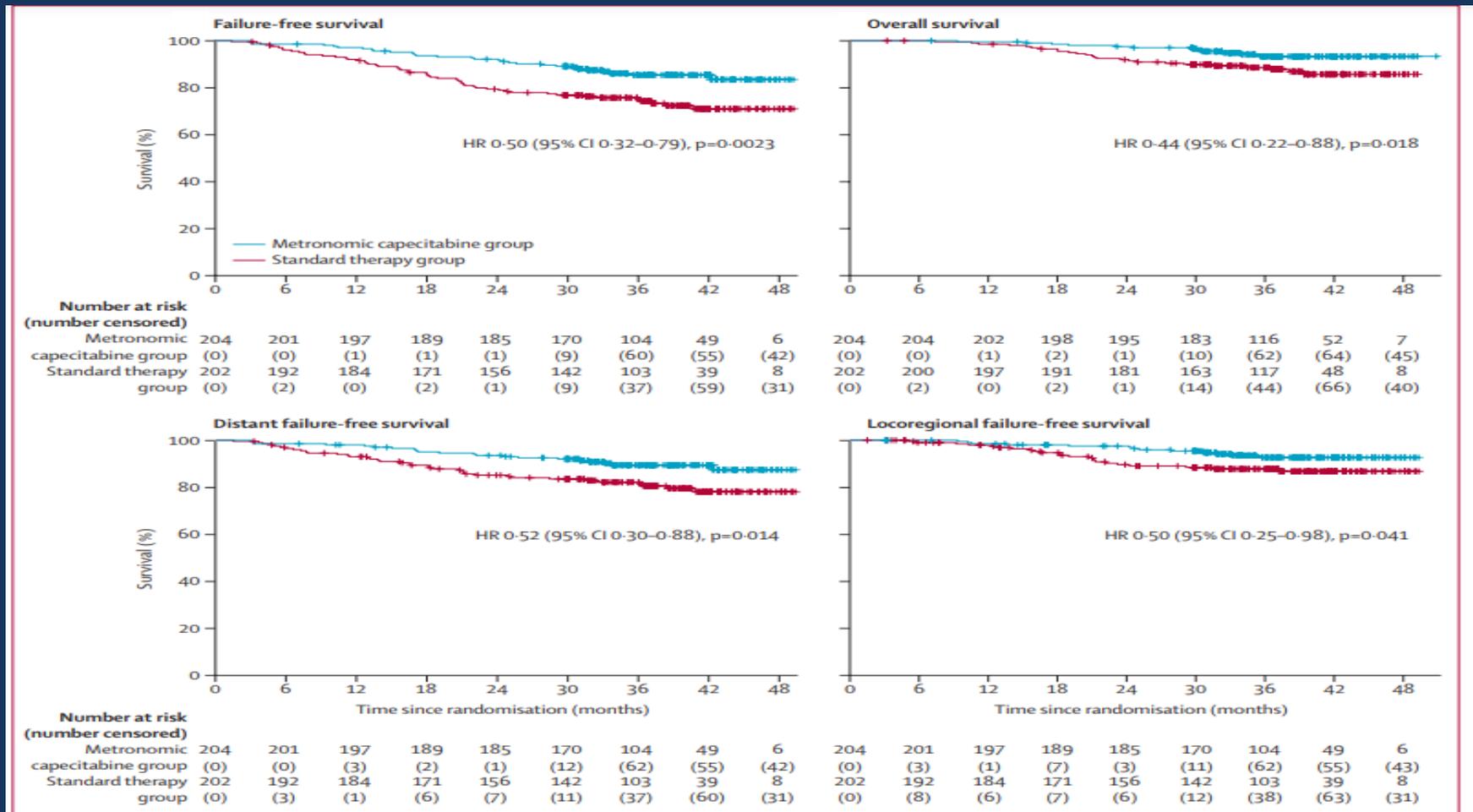


Capecitabine 1 gm/m² BID D1-D14 q3weeks X8 cycles



Observation

Results- Median follow up-38 months



Toxicity

	Metronomic capecitabine group (n=201)			Standard therapy group (n=200)		
	Any grade	Grade 1 or 2	Grade 3 or 4	Any grade	Grade 1 or 2	Grade 3 or 4
Any adverse event	182 (91%)	147 (73%)	35 (17%)	112 (56%)	101 (51%)	11 (6%)
Haematological adverse event						
Leukopenia	54 (27%)	48 (24%)	6 (3%)	39 (20%)	33 (17%)	6 (3%)
Neutropenia	37 (18%)	30 (15%)	7 (3%)*	25 (13%)	20 (10%)	5 (3%)
Anaemia	71 (35%)	70 (35%)	1 (<1%)	51 (26%)	49 (25%)	2 (1%)
Thrombocytopenia	24 (12%)	23 (11%)	1 (<1%)	19 (10%)	19 (10%)	0
Non-haematological adverse event						
Hand-foot syndrome	117 (58%)	<u>99 (49%)</u>	<u>18 (9%)</u>	0	0	0
Fatigue	55 (27%)	54 (27%)	1 (<1%)	36 (18%)	36 (18%)	0
Nausea	44 (22%)	42 (21%)	2 (1%)	21 (11%)	21 (11%)	0
Sensory neuropathy	37 (18%)	34 (17%)	3 (1%)	16 (8%)	14 (7%)	2 (1%)
Anorexia	36 (18%)	36 (18%)	0	18 (9%)	18 (9%)	0
Weight loss	27 (13%)	30 (15%)	0	13 (7%)	13 (7%)	0
Vomiting	26 (13%)	25 (12%)	1 (<1%)	14 (7%)	14 (7%)	0
Elevated ALT or AST concentrations	23 (11%)	23 (11%)	0	15 (8%)	15 (8%)	0
Mucositis or stomatitis	21 (10%)	<u>20 (10%)</u>	1 (<1%)	9 (5%)	9 (5%)	0
Diarrhoea	19 (9%)	<u>18 (9%)</u>	1 (<1%)	4 (2%)	4 (2%)	0

Results

Median follow up - 44.8 months

Variable			ITT		PP	
			CCRT+AC	CCRT alone	CCRT+AC	CCRT alone
			(N= 90)	(N= 90)	(N= 71)	(N= 90)
FFS	Failure or death, N (%)		15 (16.7)	27 (30.0)	8 (11.3)	27 (30.0)
	3-y FFS, %	<i>P</i> = 0.037	87.7	73.3	92.9	73.3
OS	Death, N (%)		8 (8.9)	12 (13.3)	3 (4.2)	12 (13.3)
	3-y OS, %	HR=0.66	92.6	88.9	98.6	88.9
DMFS	Distant metastasis or death, N (%)		13 (14.4)	19 (21.1)	6 (8.5)	19 (21.1)
	3-y DMFS, %	GR=0.67	88.8	81.1	94.3	81.1

Acute Toxicity

- G3-4 acute toxicity (57.8% vs 51.1%)
- Hand foot syndrome (3.5% vs 0%)
- Xerostomia (11.1% vs 3.3%)
- Mucositis (23.3% vs 16.7%)
- Anemia (5.6% vs 2.2%)

Adjuvant Capecitabine after CCRT

- Difference in inclusion criteria
- Difference in dose and duration
- Number of patients were less in the second study
- NACT permitted in one study
- 2 studies – OS Survival benefit
- Increased toxicity
- One published and one in abstract form
- Awaiting long term follow up
- Adjuvant in patients received NACT?



Contents lists available at ScienceDirect

Clinical and Translational Radiation Oncology

journal homepage: www.sciencedirect.com/journal/clinical-and-translational-radiation-oncology

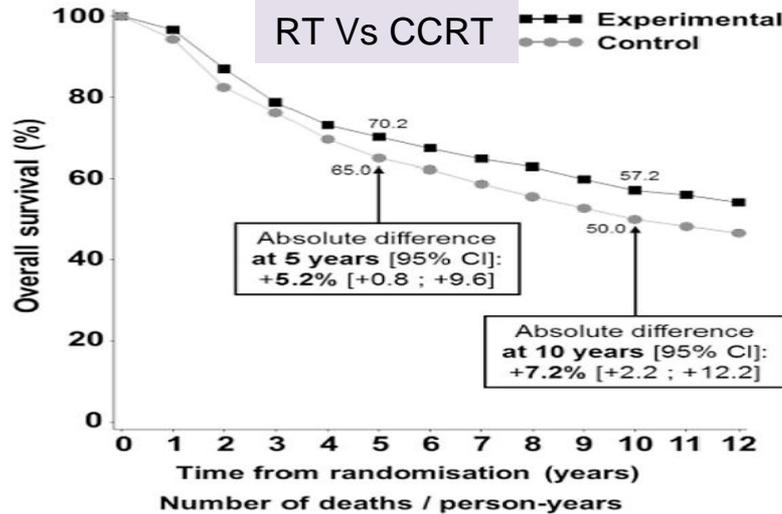


Original Research Article

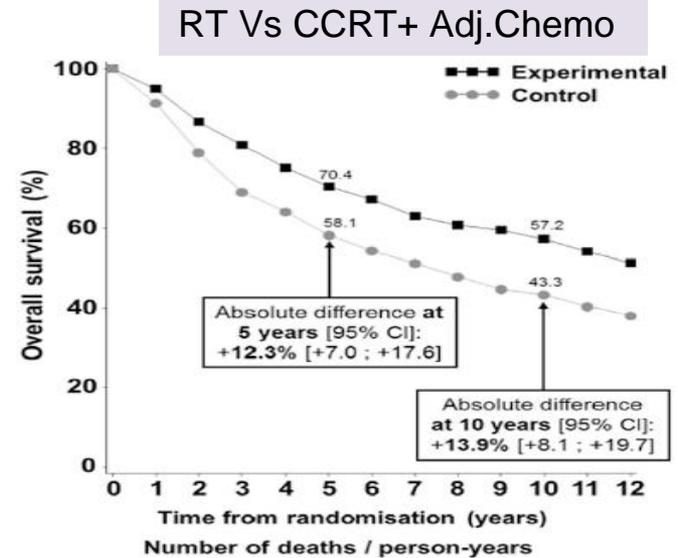


Meta-analysis of chemotherapy in nasopharynx carcinoma (MAC-NPC): An update on 26 trials and 7080 patients

Pierre Blanchard^{a,b,*}, Anne W.M. Lee^c, Alexandra Carmel^{b,d}, Ng Wai Tong^c, Jun Ma^e, Anthony T.C. Chan^f, Ruey Long Hong^g, Ming-Yuan Chen^h, Lei Chen^h, Wen-Fei Li^h, Pei-Yu Huang^h, Dora L.W. Kwongⁱ, Sharon S.X. Poh^j, Roger Ngan^c, Hai-Qiang Mai^h, Camille Ollivier^{b,d}, George Fountzilas^k, Li Zhang^h, Jean Bourhis^l, Anne Aupérin^{b,d}, Benjamin Lacas^{b,d}, Jean-Pierre Pignon^{b,d}, on behalf of the MAC-NPC collaborative Group



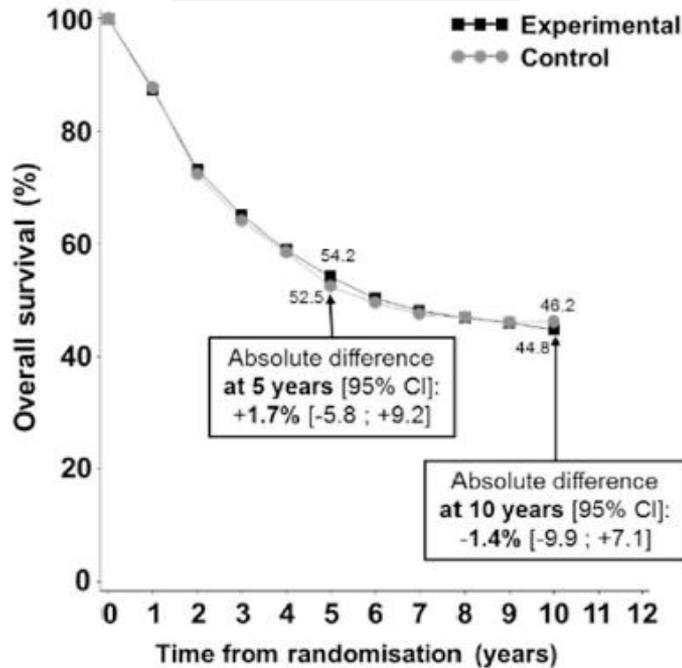
	Years [0;2[Years [2;5[Years [5;10[Years 10+
Experimental	116 / 1714	146 / 1974	90 / 2244	34 / 1147
Control	157 / 1655	146 / 1847	106 / 1979	33 / 934



	Years [0;2[Years [2;5[Years [5;10[Years 10+
Experimental	84 / 1188	102 / 1458	68 / 1537	70 / 1259
Control	135 / 1151	129 / 1231	74 / 1221	55 / 916

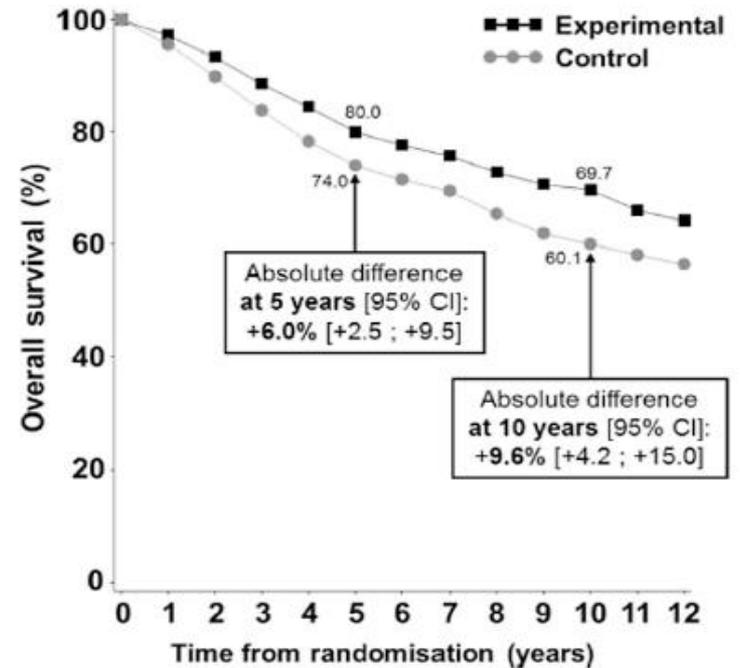
Efficacy

RT Vs NACT+RT



	Years [0;2[Years [2;5[Years [5;10[Years 10+
Experimental	102 / 664	61 / 599	17 / 383	0 / 36
Control	107 / 675	61 / 563	13 / 386	0 / 49

CCRT Vs NACT+CCRT



	Years [0;2[Years [2;5[Years [5;10[Years 10+
Experimental	80 / 2308	139 / 2718	48 / 1658	8 / 207
Control	119 / 2246	169 / 2558	63 / 1579	7 / 221

Conclusions

- Stage I- Radical RT
- UNI in N0- same efficacy with less late toxicity
- Stage II – No role for NACT
- Stage II- ? IMRT alone
- Stage III& IVa – IC followed by CCRT
- Superiority of 3 drug NACT is not proven
- Adjuvant Capecitabine - Promising results with excess toxicity
- Adjuvant CDDP+5FU in risk adapted approach – No benefit



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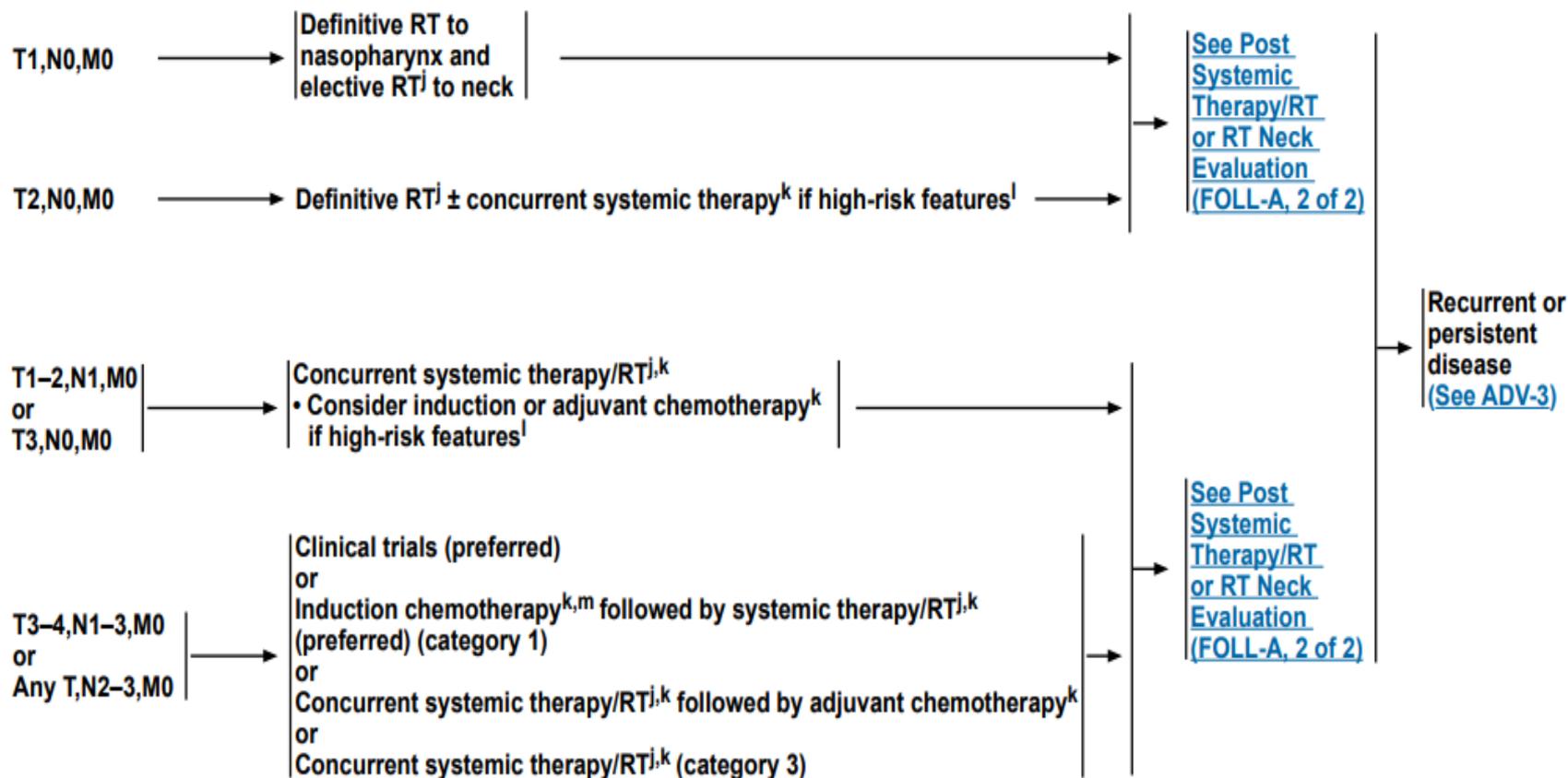
Cancer of the Nasopharynx

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CLINICAL STAGING

TREATMENT OF PRIMARY AND NECKⁱ

FOLLOW-UP



Research Article

Clinical Profile and Treatment Outcomes in Patients Treated with Intensity-Modulated Radiotherapy (IMRT) for Carcinoma Nasopharynx: A Retrospective Analysis

Farida Nazeer,¹ R. Rejnish Kumar,¹ Malu Rafi,¹ Tapesh Bhattacharya,¹ Aparna Mullangath Prakasan,¹ Kumar P. Naveen,¹ Preethi George,² Ramadas Kunnambath,¹ and Kainickal Cessal Thommachan ¹

¹Department of Radiation Oncology, Regional Cancer Centre, Thiruvananthapuram, Kerala, India

²Department of Cancer Epidemiology and Biostatistics, Regional Cancer Centre, Thiruvananthapuram, Kerala, India

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Chemotherapy sequencing	N=84	No. of patients (%)
Neoadjuvant chemotherapy alone		5 (6.3%)
Neoadjuvant + concurrent		34 (41.9%)
Concurrent + adjuvant		4 (4.9%)
Neoadjuvant + concurrent + adjuvant		3 (3.7%)
Concurrent alone		28 (34.5%)
No chemotherapy		7 (8.7%)

Stage (<i>n</i> = no. of patients)	Survival probability (%)
I (<i>n</i> = 2)	100
II (<i>n</i> = 19)	67.0
III (<i>n</i> = 31)	70.4
IV (<i>n</i> = 29)	68.1



Thank you
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