Chest CT scan+CXR versus CXR For follow up

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IFCT 0302



Concept

- Need for follow up?
- Diagnose-
 - -Recurrence at earlier stage
 - -Second primary
- Methods-

-History and physical examination/CXR/ CT scan/ Bronchoscopy

Recurrence pattern



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Local recurrence following lung cancer surgery: Incidence, risk factors, and outcomes

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Author	Year	Number of patients	Percent of patients recurring (n)	Percent of pati recurrence	ents with initial purely local (n)	Percent of patients with initial recurrence mixed (<i>n</i>)	Percent of patients with any initial local recurrence (n)
Taylor ⁵	2012	1143	33 (378)		8 (94)	None listed	8 (94)
Saynak ¹¹	2010	335	33 (111)		12 (41)	12 (41)	24 (82)
Kelsey ⁹	2009	975	26 (250)		7 (63)	8 (78)	15 (141)
Hung ¹³	2009	933	31 (289)		8 (74)	5 (49)	13 (123)
Nakagawa ^a , ¹²	2008	397	22 (87)		7 (30)	None listed	7 (30)
Sugimura ¹⁰	2007	1073	36 (390)		7 (79)	6 (62)	13 (141)
Martini ^a ,7	2005	598	27 (159)		5 (32)	None listed	5 (32)

Post-excision local recurrence rate (%) by lung cancer stag							
	Stage						
Study	IA	IB	IIA	IIB	IIIA		
Choi 2011 ^a , ¹⁶	4	7	11	12	24		
Saynak 2010 ^b , ¹¹	19	19	27	27	38–40		
Pisters 2005 ¹⁵	10	10	N/A	12	15		



Recurrence Pattern

Differences in Patterns of Recurrence in Early-Stage Versus Locally Advanced Non-Small Cell Lung Cancer

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Median follow up 35 months

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In stage IIIA-
52% developed recurrence, 85% were distant, and by CT surveillance-49%
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In early stage-20% developed recurrence and 74% were distant, by CT surveillance- 61%

Survival was better in CT detected recurrences compared to symptomatic





Follow up strategy

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Postoperative follow-up strategy based on recurrence dynamics for non-small-cell lung cancer[†]

Katsuya Watanabe^{a,*}, Masahiro Tsuboi^b, Kentaro Sakamaki^c, Teppei Nishii^a, Taketsugu Yamamoto^a, Takuya Nagashima^a, Kohei Ando^a, Yoshihiro Ishikawa^a, Tekkan Woo^a, Hiroyuki Adachi^a, Yutaka Kumakiri^a, Takamitsu Maehara^a, Haruhiko Nakayama^a and Munetaka Masuda^a

To clarify postoperative follow up strategy based on time of recurrence



Follow up strategy



Follow up strategy



Bimodal recurrence pattern

Hospital visitation programmesfocus on 6–8 and 22–24 months after surgery

Male- 6-8 months Female- 22 to 24 months

Newer evidences on recurrence

Recurrence location	Arm A: Lobectomy (N=554)	Arm B: Segmentectomy (N=552)
Total	44 (7.9%)	67 (12.1%)
Loco-regional	17 (3.1%)	38 (6.9%)
Distant	14 (2.5%)	7 (1.3%)
Loco-regional + distant	13 (2.3%)	20 (3.6%)
Unclassified	0	2
Proportion of local recurrence	30 (5.4%)	58 (10.5%)

	Lobar N=351	Sublobar N=336	Total N=687	P-Value ¹
Overall	103 (29.3%)	102 (30.4%)	205 (29.8%)	0.8364
Locoregional only	35 (10%)	45 (13.4%)	80 (11.6%)	0.2011
Regional only	9 (2.6%)	6 (1.8%)	15 (2.2%)	0.6623
Any Distant	59 (16.8%)	51 (15.2%)	110 (16.0%)	0.6323





Second primary

SEER database review- 2004 to 2014



Recurrence and Second Primary

Rates of Recurrence and Metachronous Cancer After Surgery



Recommendations

Organization	Summary of recommendations
ACCP [18]	Surveillance by clinical examination and chest radiograp and then yearly for patients with good performance st
ESMO [<mark>19</mark>]	A follow-up visit every 3-6 months is recommended dur For follow-up, history and physical examination, chest Cl
NCCN [<mark>20</mark>]	History and physical examination with contrast-enhance Then history and physical examination and non-contrast
ASCO [21]	For patients treated with curative intent, in the absence of performed every 3 months during the first 2 years; ever For patients treated with curative intent, there is no clear patients in whom no interventions are planned
NICE [22]	Offer all patients an initial specialist follow-up appointment ongoing care. Offer regular appointments thereafter, rethey experience symptoms Offer protocol-driven follow-up led by a lung cancer cline expectancy of more than 3 months Ensure that patients know how to contact the lung cancer scheduled hospital visits

	Classification of recommendation
hy or CT should be performed every 6 months for 2 years tatus and pulmonary function	Grade 1C
ing 2–3 years, less often–e.g. annually–thereafter	III, B
T and, to a lesser extent, chest X-ray are appropriate tools	III, B
d CT scan every 4-6 months for 2 years	2B
enhanced CT scan annually	2B
of symptoms, a history and physical examination should be ery 6 months thereafter through year 5; and yearly thereafter r role for routine studies in asymptomatic patients and	None
ent within 6 weeks of completing treatment to discuss rather than relying on patients requesting appointments when	None
nical nurse specialist as an option for patients with a life	
er clinical nurse specialist involved in their care between their	



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- •Phase 3
- •Randomised
- •Open label
- •122 French Hospitals
- •Approved by Ethical committee, Data Monitoring +
- •Study period: Jan 3, 2005 Nov 30, 2012

Randomisation

- •Two groups
 - Minimal follow up group (CXR)
 - •CT based follow up group(CECT chest+ CXR+ FOB)
- •Randomised- 8 weeks after surgery

- •Stratified Randomisation
 - •Centre
 - •Stage
 - Periop- chemo/radio
 - •Computer generated randomisation

Outcome

• Primary end point- OS

Secondary end point –DFS

Survival from recurrence or second primary
Genetic risk factor for lung cancer,
Health related QoL
Cost effectiveness



Awaited

Statistical analysis

- •Difference of 7.5% in 3 year OS
- •Estimated 3 yr OS of 40% in the minimal follow up group (changed to 68%)
- •Power- 90% and alpha level of 5%
- •Intention to treat analysis





Result

887 assigned to the CT-based follow-up group

887 included in the intention-to-treat population 709 included in the per-protocol population*

	Minimal follow-up group (n=888)	CT-based follow-up group (n=887)
Sex		
Male	678 (76-4%)	677 (76-3%)
Female	210 (23-6%)	210 (23.7%)
Median age, years (IQR)	63-0 (57-1-70-4)	62.9 (56.2-70.5)
Histological subtype		
Squamous	307 (34-6%)	307 (34-6%)
Adenocarcinoma	520 (58-5%)	522 (58-9%)
Large cell	38 (4-3%)	41 (4.6%)
Others	23 (2-6%)	17 (1-9%)
Smoking status		
Former or current smoker	818 (92-1%)	805 (90-7)
Never smoker	68 (7.7%)	80 (9.0%)
Unknown	2 (0.2%)	2 (0.3%)
Clinical stage		
E Construction of the second	606 (68-2%)	599 (67.5%)
II. Contraction of the second s	119 (13-4%)	125 (14.1%)
III	161 (18-1%)	161 (18-1%)
Unknown	2 (0.2%)	2 (0.3%)

Result

	Minimal follow-up	p CT-based follow-up	Clinical stage		
	group (n=888)	group (n=887)	I. State of the second s	606 (68.2%)	599 (67·5%
			II	119 (13-4%)	125 (14·1%
	678 (76-4%)	677 (76.3%)	III	161 (18-1%)	161 (18-1%
	210 (23-6%)	210 (23.7%)	Unknown	2 (0.2%)	2 (0.3%)
e, years (IQR)	63-0 (57-1-70-4)	62-9 (56-2-70-5)	Surgery		
l subtype			Lobectomy or bilobectomy	758 (85·4%)	775 (87.4%
US	307 (34.6%)	307 (34.6%)	Pneumonectomy	111 (12.5%)	95 (10-7%
rcinoma	520 (58·5%)	522 (58·9%)	Segmentectomy	16 (1-8%)	15 (1·7%)
I de la construcción de la constru	38 (4-3%)	41 (4-6%)	Unknown	3 (0.3%)	2 (0.2%)
	23 (2.6%)	17 (1.9%)	Pathological stage		
tatus			E Construction of the second	559 (62-9%)	561 (63-2%
r current smoker	818 (92.1%)	805 (90-7)	II. Contraction of the second s	158 (17.8%)	165 (18·6%
noker	68 (7.7%)	80 (9-0%)	III III III III III III III III III II	163 (18-4%)	152 (17.1%
n	2 (0.2%)	2 (0.3%)	Unknown	8 (0.9%)	9 (1.0%)
ge	606 (68-2%)	599 (67.5%)	Preoperative chemotherapy, or preoperative radiotherapy, or both	110 (12.4%)	116 (13-1%)
	119 (13.4%)	125 (14.1%)	Postoperative chemotherapy, radiotherapy, or both	342 (38.5%)	350 (39-5%
	161 (18.1%)	161 (18.1%)	Preoperative or postoperative radiotherapy, or both	80 (9.0%)	74 (8-3%)
n	2 (0.2%)	2 (0.3%)	Preoperative or postoperative chemotherapy, or both	397 (44-7%)	403 (45·4%



Figure 2: Kaplan-Meier overall survival curves by follow-up group (intention-to-treat population) HR=hazard ratio.

ITT- OS, DFS



• Most frequent site of metastases

- Ipsilateral lung
- Contralateral lung
- Brain

• 250 unjustified CT scans done in the minimal follow up group

Result

CXR

Recurrences- 27.7% Symptomatic- 82.5%

CT scan

Recurrences- 32.6% Symptomatic- 56%



Interesting findings

CXR

Surgery-5.7% Radical therapy for second primary- 19%

- CXR
- Only 3.3% were detected on FOB- which could not be picked up on a CT scan

CT scan

Surgery-10.3% Radical therapy for second primary- 29%

• Amongst all the recurrences detected in the CT scan group; 42% were not detected on a concomitant

Discussion

- CT scan group
 - higher proportion of asymptomatic recurrences
 - higher proportion of second primaries
 - earlier stage

Not powered enough to show OS benefit in recurrences and second primaries

STRENGTHS

- Randomised design
- 90% compliance
- Median follow up of 7.2 years
- Robust data on recurrences and second primaries

WEAKNESS

- Higher proportion of stage I and II cancers
- Started in 2005- almost 17 years ago- treatment strategies have changed significantly
- No central radiology review- not mentioned who interpreted the x ray or CT findings
- criteria where used at different time points in the study
- Significant cross over b/w groups- around 8 %

• Criteria for following up pulmonary nodules also have changed over time- not sure what

CONCLUSIONS

- CT scan-
 - More sensitive
 - Significant advancement in treatment-be performance status)
- Problems
 - Cost effective?
 - False positive
 - More robust evidence needed

• Significant advancement in treatment- better tolerated in asymptomatic patients (better



Thank You