

# Pattern of care and outcome of non-small cell lung cancer patients receiving gefitinib beyond radiologic progression.

Hideo Kunitoh, Yukio Hosomi, Chiharu Tanai, Kiyotaka Yoh, Yasushi Goto, and Yasuo Ohashi

Department of Medical Oncology, Japanese Red Cross Medical Center, Tokyo, Japan; Department of Thoracic Oncology and Respiratory Medicine, Tokyo Metropolitan Cancer and Infectious Diseases Center Komagome Hospital, Tokyo, Japan; Division of Respiratory Medicine, NTT Medical Center, Tokyo, Japan; Division of Thoracic Oncology, National Cancer Center Hospital East, Kashiwa, Japan; Department of Thoracic Oncology, National Cancer Center Hospital, Tokyo, Japan; Chuo University, Tokyo, Japan

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## Background

- We previously showed (Yamane et al, ESMO 2014) that 1/3 of non-small cell lung cancer patients (NSCLC pts) with activating epidermal growth factor receptor mutation (EGFRm+) continued to receive an EGFR-tyrosine kinase inhibitor after radiological “progressive disease” (R-PD) was seen, if they remained clinically stable, until clinical deterioration (C-PD).

## Objectives

- To explore for patients subgroups who would be likely to receive benefit from “continuation of TKI beyond PD” strategy, in gefitinib-receiving cohort.
- Focusing on elderly (≥70 y.o.) and female patients, as compared to non-elderly and male patients, respectively.

# Study subjects (Patients)

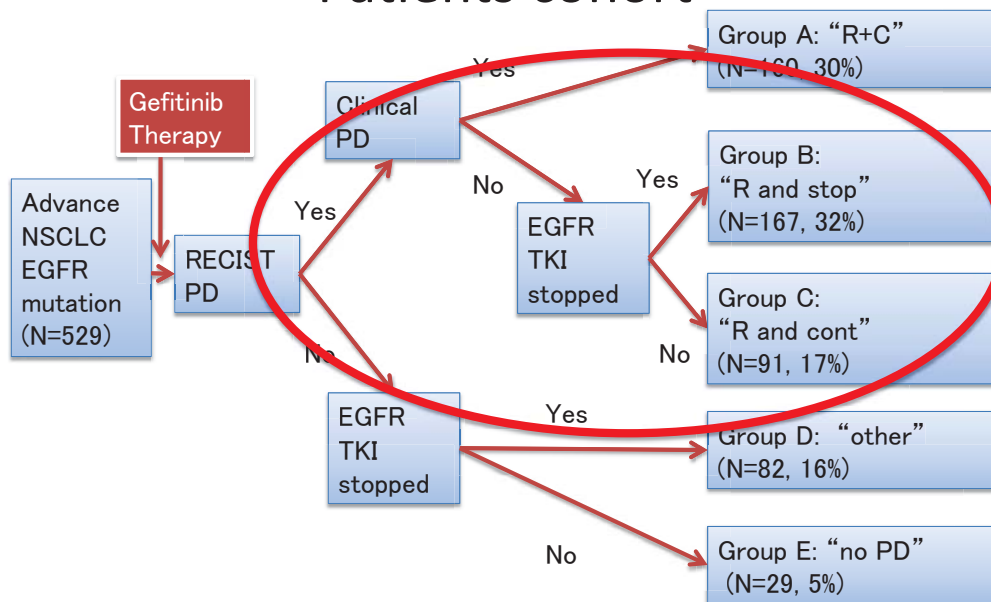
- Inclusion criteria
  - Advanced or post-operational recurrent non-small-cell lung cancer
  - Diagnosed as having tumor harboring EGFR mutation
    - Definition of EGFR gene mutation positive (mutation of sensitive gene)
      - (A) Deletion of Exon19 (irrespective of the subtype)
      - (B) Exon 21 L858R
      - (C) Other rare mutations (Exon 18 G791X, etc.)
    - EGFR gene mutation excluded from this study:
      - (A) Exon 20 insertion mutation
      - (B) T790M
  - Treatment with Gefitinib or was started from January 1, 2009 until December 31, 2011 as the initial anti-cancer therapy
- Exclusion criteria
  - Prior treatment with cytotoxic chemotherapy
  - Concomitant malignancy

# Definition of

## “Clinical PD (disease progression)”

- Symptomatic progression
- Declining of PS due to progression
- Threat to major organ(s)
- Unequivocal multi-organ progression

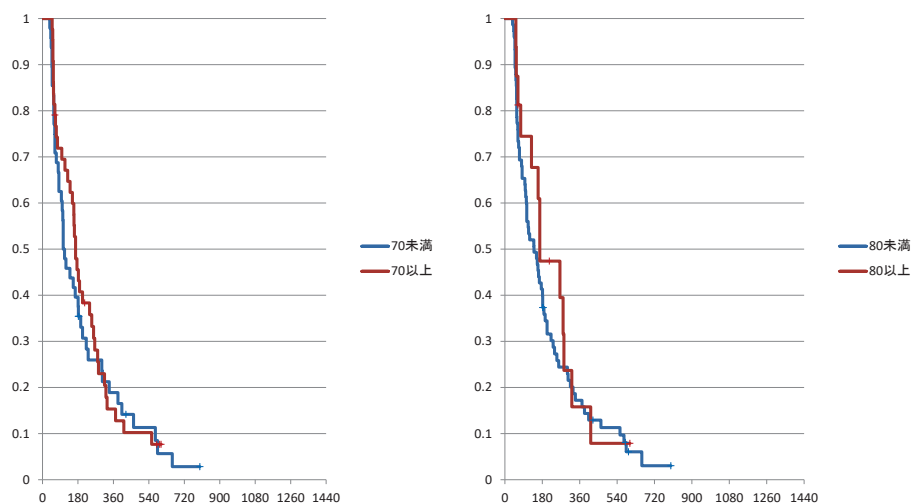
### Patients cohort



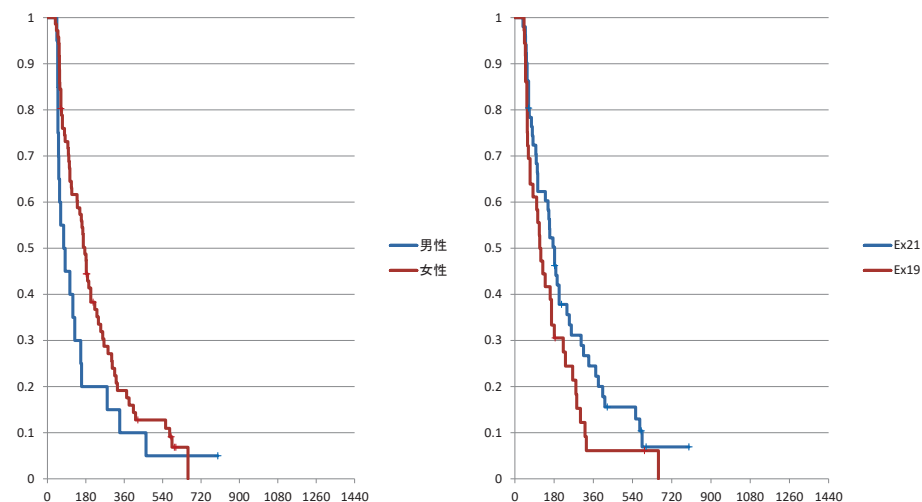
Patient characteristics of those who remained clinically stable on radiological PD (groups B and C)

| Group              | B (N=167)  | C (N=91)   | P value |
|--------------------|------------|------------|---------|
| Gender M           | 58 (35%)   | 20 (22%)   | 0.03    |
| Gender F           | 109 (65%)  | 71 (78%)   |         |
| Age median (Range) | 69 (31~88) | 69 (27~93) | >0.2    |
| < 70 y.o.          | 95 (61%)   | 48 (53%)   | >0.2    |
| > 70 y.o.          | 72 (39%)   | 43 (47%)   |         |
| < 80 y.o.          | 147 (88%)  | 75 (82%)   | >0.2    |
| > 80 y.o.          | 20 (12%)   | 16 (18%)   |         |
| PS 0-1             | 131 (78%)  | 69 (76%)   | >0.2    |
| PS 2-4             | 36 (22%)   | 22 (24%)   |         |
| Never smoker       | 106 (63%)  | 71 (78%)   | 0.05    |
| Ever smoker        | 61 (37%)   | 20 (22%)   |         |
| No prior Tx        | 123 (74%)  | 59 (65%)   | 0.15    |
| Prior Tx           | 44 (26%)   | 32 (35%)   |         |
| Mutation ex19      | 81 (49%)   | 36 (40%)   | >0.2    |
| Mutation ex21      | 77 (46%)   | 51 (56%)   |         |
| Mutation others    | 9 (5%)     | 4 (4%)     |         |
| CR/PR              | 123 (74%)  | 72 (79%)   | >0.2    |
| NC/PD/NE           | 44 (26%)   | 19 (21%)   |         |

## Group C: R-PD to C-PD



## Group C: R-PD to C-PD

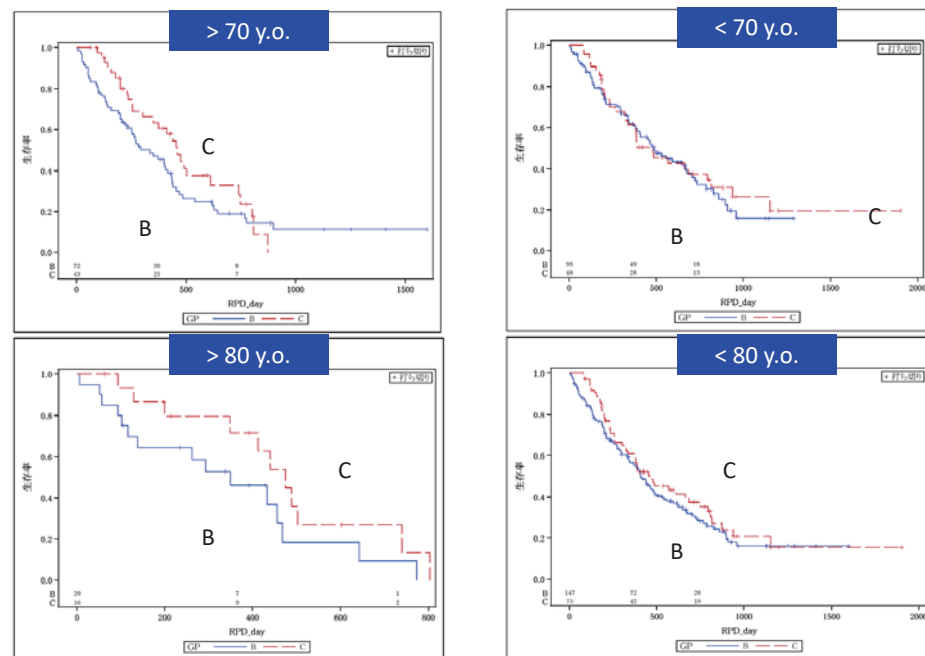


## Reason of C-PD in Group C

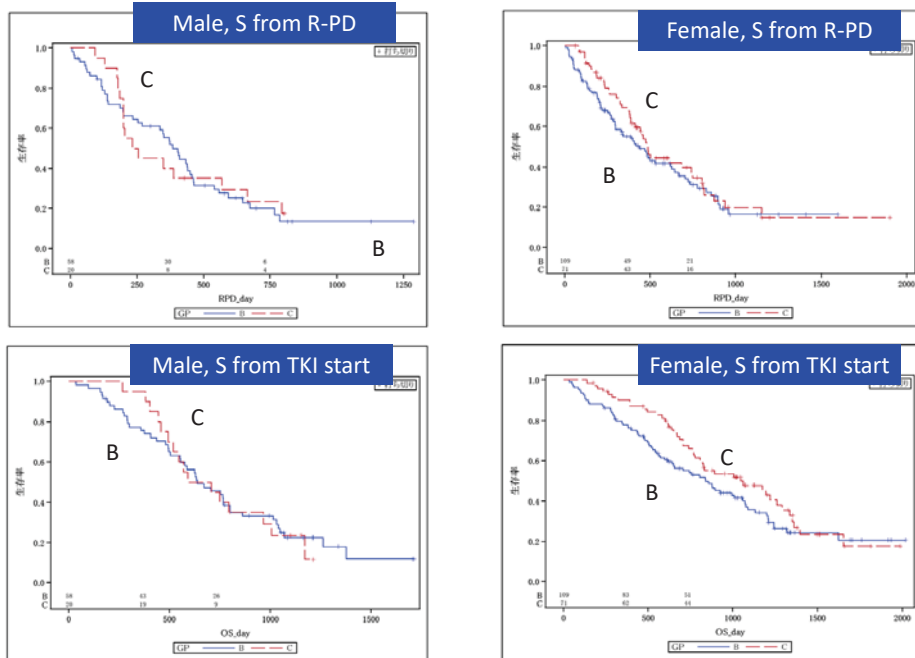
|        | All      | <70      | >70      | <80      | >80     | Male    | Female   |
|--------|----------|----------|----------|----------|---------|---------|----------|
| Reason | N=91     | N=48     | N=43     | N=85     | N=16    | N=20    | N=71     |
| 1      | 36 (40%) | 18 (38%) | 18 (42%) | 31 (36%) | 5 (31%) | 8 (40%) | 28 (39%) |
| 2      | 26 (29%) | 12 (25%) | 14 (33%) | 20 (24%) | 6 (38%) | 6 (30%) | 20 (28%) |
| 3      | 11 (12%) | 6 (13%)  | 5 (12%)  | 10 (12%) | 1 (6%)  | 4 (20%) | 7 (10%)  |
| 4      | 41 (45%) | 27 (56%) | 14 (33%) | 37 (44%) | 4 (25%) | 7 (35%) | 34 (48%) |

1. Symptomatic progression
2. Declining of PS due to progression
3. Threat to major organ(s)
4. Unequivocal multi-organ progression

## Survival from R-PD and Age



## Survival and gender



## Summary (cont'd)

- Period from R-PD to clinical PD tended to be longer in female patients who received gefitinib “beyond PD”.
- Survival after R-PD tended to be longer in elderly (but not non-elderly) patients who received gefitinib “beyond PD” than who did not.

## Summary

- Female patients were more likely to receive gefitinib continuation on clinically stable, radiological PD (R-PD).
- Age was not significantly associated with gefitinib continuation on R-PD.

## Conclusions and Acknowledgement

- Female and elderly patients could be candidates for the subpopulations who are likely to receive benefit from the strategy of gefitinib “beyond PD” continuation.
- More research should be warranted.
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