P4-02-25



Analysis of prognosis in different subtypes of invasive lobular carcinoma using a National Cancer Database Breast Cancer Registry of Japan

Background

- Invasive lobular carcinoma (ILC) has more likely to be hormone receptor (HR) positive, and several studies reported that the prognosis of ILC was better than invasive ductal carcinoma (IDC) [1].
- However, ILC also has different prognosis according to the subtypes as IDC does[2], and better prognosis of ILC might depend on their high HR positivity.
- Additionally, there are many reports that chemotherapy (CT) does not improve the prognosis of ILC due to the high positivity of HR [3].
- ILC usually constitutes small population of invasive breast cancer [1], therefore, the data from multiple institutions is needed for more accurate analyses.
- The National Clinical Database (NCD) is a platform for nationwide cancer registry in Japan. It contains records of more than 300,000 breast cancer patients from more than 800 institutions in Japan.

Objectives

- To compare the prognosis of IDC and ILC in each subtype
- To assess the effect of CT on luminal ILC

Study design

Analysis of the prognosis in IDC and ILC

- Inclusion criteria
- IDC or ILC
- Do not have distant metastasis
- Received surgery for primary breast cancer
- Did not receive preoperative therapy
- Do not have bilateral breast cancer
- The cases with 10-year follow-up data
- ✓ To evaluate the prognosis of each subtype, we compared DFS and OS for IDC and ILC in each subtype.

Analysis of the effect of CT in luminal ILC

Inclusion criteria

www.PosterPresentation

- Satisfy above criteria
- Luminal ILC with pT2N0M0 or pT1-2N1M0
- Received endocrine therapy (ET)
- ✓ To evaluate the effect of CT in luminal ILC, we compared DFS and OS for ET+CT group and ET only group in luminal ILC.
- Because it was presumed that there are differences in pathological and clinical characteristics, we have planned to make the matched cohorts by using exact matching for comparing their prognosis.
- DFS was defined as the time from surgery to local or distant recurrence or death from any cause. OS was defined as the time between the surgery and the death from any cause.
- Peason's Chi squared test was used to identify the characteristics. Survival curves were constructed by Kaplan-Meier method and were compared by log-rank test. All analyses were conducted using SAS version 9.4 (SAS Institute, Cary, NC, USA).

Table1: Consort diagram of this study

| All records of breast cancer in Japanese Breast Cancer Registry, 2004-2012 n=318,338 |
|--|
| \downarrow |
| IDC or ILC n=250,736 |
| |
| IDC or ILC satisfying criteria n=250,736 |
| <u> </u> |
| Cohort for analysis IDC; n=136,654 ILC; n=5,705 |
| Ļ |
| Matched cohort for prognostic analysis IDC; n=5,633 ILC; n=5,633 |

(matched cohort)

| Total |
|----------------------------------|
| Age |
| <40 |
| <60 |
| <80 |
| 80 and above |
| Menopause status |
| Menopause |
| Pre menopause |
| Unknown |
| Tumor size |
| <2cm |
| 2cm-, <5cm |
| 5cm- |
| Unknown |
| ER |
| Positive |
| Negative |
| Not administered |
| No information |
| PgR |
| Positive |
| Negative |
| Not administered |
| No information |
| HER2 |
| Positive |
| Negative |
| Not administered / No informatio |
| Lymph node status |
| None |
| 1 to 3 |
| 4 to 9 |
| 10 and above |
| Not administered / No informatio |
| Initial surgical treatment |
| Mastectomy |
| Breast-conserving surgery |
| |
| |

San Antonio Breast Cancer Symposium - December 6-10, 2022

Yayoi Adachi¹⁾, Sota Asaga²⁾, Hiraku Kumamaru³⁾, Yutaka Yamamoto⁴⁾, Shigeru Imoto²⁾, Hiromitsu Jinno⁵⁾ 1)UT southwestern medical center, Dallas, Tx, USA 2) Kyorin University of Tokyo, Japan 3) University School of Medicine, Tokyo, Japan 3) University of Tokyo, Japan 3) University School of Medicine, Tokyo, Japan 3) University School of Medicine

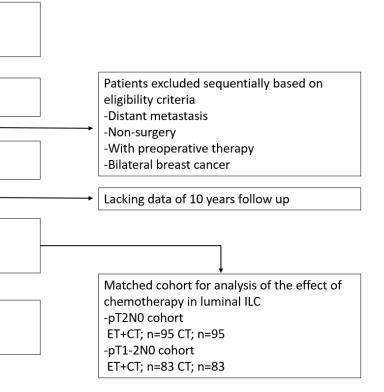


Figure1: Prognosis of overall population

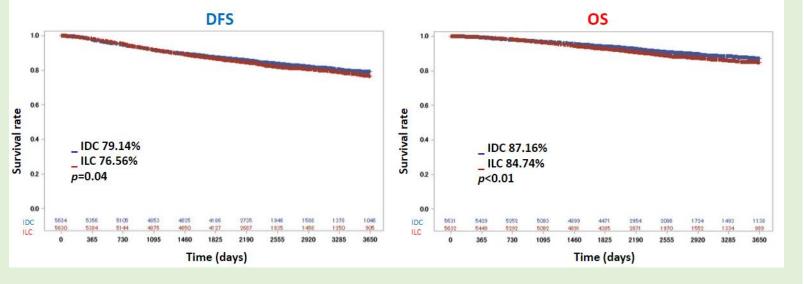
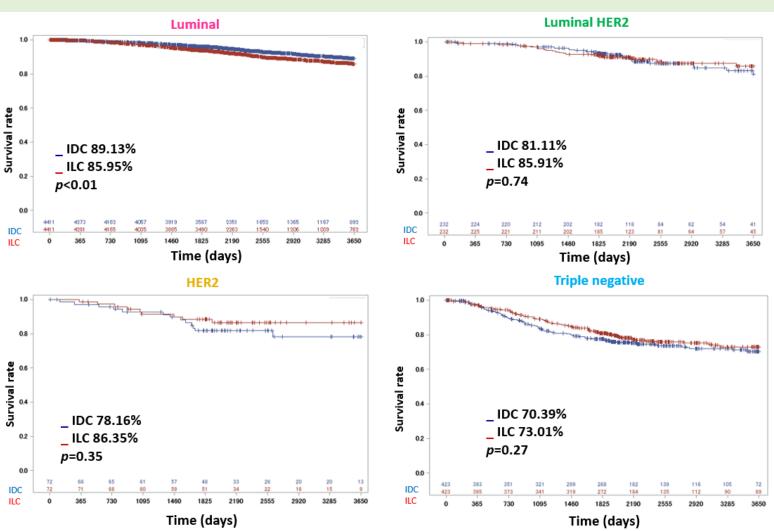
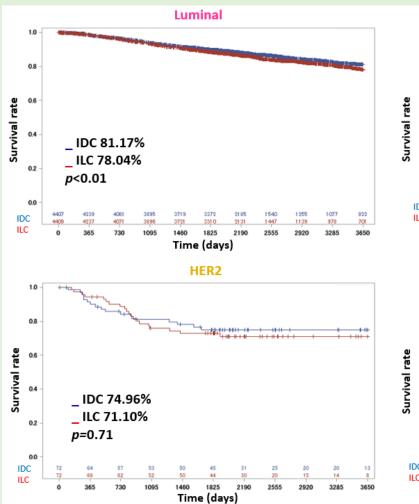


Figure2: DFS in each subgroup







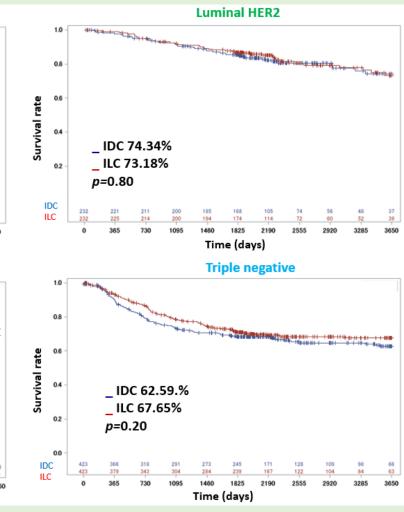


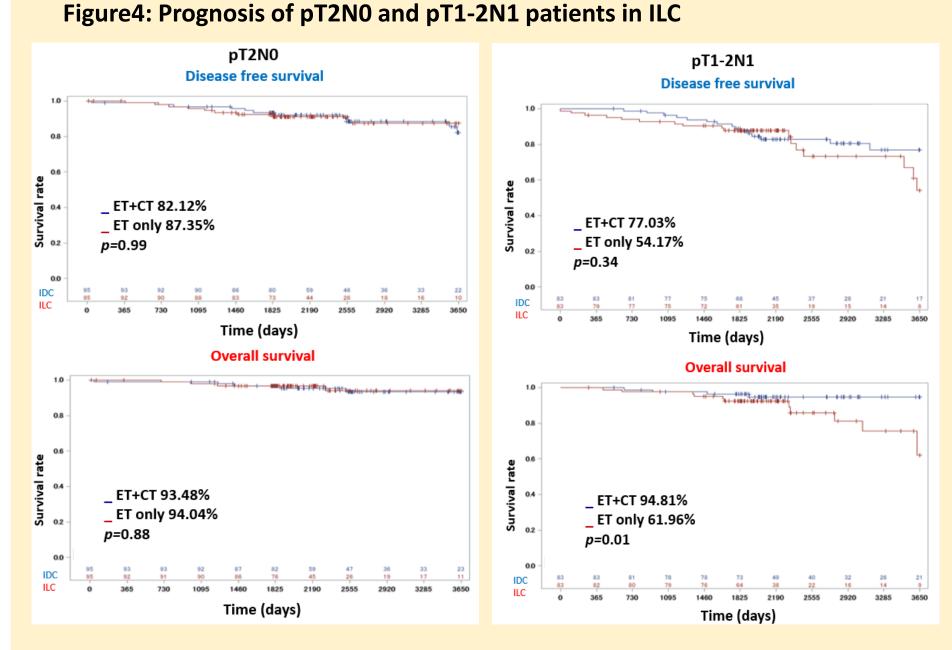
Table2: Patient characteristics of IDC and ILC

| ID | C | ILC | | | |
|------|-------|------|-------|--|--|
| n | % | n | % | | |
| 5633 | | 5633 | | | |
| | | | | | |
| 152 | 2.70 | 152 | 2.70 | | |
| 2673 | 47.45 | 2673 | 47.45 | | |
| 2407 | 42.73 | 2407 | 42.73 | | |
| 401 | 7.12 | 401 | 7.12 | | |
| | | | | | |
| 3702 | 65.72 | 3702 | 65.72 | | |
| 1750 | 31.07 | 1750 | 31.07 | | |
| 181 | 3.21 | 181 | 3.21 | | |
| | | | | | |
| 2280 | 40.48 | 2280 | 40.48 | | |
| 2612 | 46.37 | 2612 | 46.37 | | |
| 534 | 9.48 | 534 | 9.48 | | |
| 207 | 3.67 | 207 | 3.67 | | |
| | | | | | |
| 4910 | 87.16 | 4910 | 87.16 | | |
| 566 | 10.05 | 566 | 10.05 | | |
| 138 | 2.45 | 149 | 2.65 | | |
| 19 | 0.34 | 8 | 0.14 | | |
| | | | | | |
| 3793 | 67.34 | 3793 | 67.34 | | |
| 1678 | 29.79 | 1678 | 29.79 | | |
| 142 | 2.52 | 154 | 2.73 | | |
| 20 | 0.36 | 8 | 0.14 | | |
| | | | | | |
| 304 | 5.40 | 304 | 5.40 | | |
| 4845 | 86.01 | 4845 | 86.01 | | |
| 484 | 8.59 | 484 | 8.59 | | |
| | | | | | |
| 3561 | 63.22 | 3561 | 63.22 | | |
| 1152 | 20.45 | 1152 | 20.45 | | |
| 373 | 6.62 | 373 | 6.62 | | |
| 340 | 6.04 | 340 | 6.04 | | |
| 207 | 3.67 | 207 | 3.67 | | |
| | | | | | |
| 3053 | 54.20 | 3053 | 54.20 | | |
| 2580 | 45.80 | 2580 | 45.80 | | |

Results

Table3: Patient characteristics of pT2N0 and pT1-2N1 patients in ILC (matched cohort)

| | pT2N0 | | | | pT1-2N1 | | | |
|------------------|-------|--------|---------|--------|---------|--------|---------|--------|
| | ET+CT | | ET only | | ET+CT | | ET only | |
| | n | % | n | % | n | % | n | % |
| Total | 95 | | 95 | | 83 | | 83 | |
| Age(years) | | | | | | | | |
| <20 | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 |
| 20-29 | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 |
| 30-39 | 2 | 2.11 | 2 | 2.11 | 2 | 2.41 | 2 | 2.41 |
| 40-49 | 23 | 24.21 | 23 | 24.21 | 28 | 33.73 | 28 | 33.73 |
| 50-59 | 23 | 24.21 | 23 | 24.21 | 24 | 28.92 | 24 | 28.92 |
| 60-69 | 29 | 30.53 | 29 | 30.53 | 21 | 25.30 | 21 | 25.30 |
| 70-79 | 16 | 16.84 | 16 | 16.84 | 8 | 9.64 | 8 | 9.64 |
| 80- | 2 | 2.11 | 2 | 2.11 | 0 | 0.00 | 0 | 0.00 |
| Menopause status | | | | | | | | |
| Menopause | 63 | 66.32 | 63 | 66.32 | 46 | 55.42 | 46 | 55.42 |
| Premenopause | 32 | 33.68 | 32 | 33.68 | 37 | 44.58 | 37 | 44.58 |
| Tumor size | | | | | | | | |
| <2cm | 0 | 0.00 | 0 | 0.00 | 20 | 24.10 | 20 | 24.10 |
| 2cm- | 95 | 100.00 | 95 | 100.00 | 63 | 75.90 | 63 | 75.90 |
| ER | | | | | | | | |
| Positive | 95 | 100.00 | 95 | 100.00 | 83 | 100.00 | 83 | 100.00 |
| Negative | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 | 0 | 0.00 |
| PgR | | | | | | | | |
| Positive | 68 | 71.58 | 68 | 71.58 | 72 | 86.75 | 72 | 86.75 |
| Negative | 27 | 28.42 | 27 | 28.42 | 11 | 13.25 | 11 | 13.25 |
| Radiotherapy | | | | | | | | |
| Yes | 1 | 1.05 | 1 | 1.05 | 4 | 4.82 | 4 | 4.82 |
| No | 94 | 98.95 | 94 | 98.95 | 79 | 95.18 | 79 | 95.18 |



In overall subtypes, the 10-year DFS of ILC was poor than those of IDC (76.56% vs 79.14%, p=0.04). (Figure 1)

- In the analysis by each subtype, there was no statistical difference in DFS for luminal HER2, HER2, and TN cohorts, however luminal ILC had statistically significant poor DFS than luminal IDC (78.04% vs 81.17%, p<0.01). (Figure 2)
- The analysis of 10-year OS showed similar results, and there were no differences in the OS of luminal HER2, HER2 and TN cohorts between ILC and IDC. However, ILC had worse OS than IDC in luminal cohort (85.95% vs 89.13%, p<0.01). (Figure 3)
- In pT2N0 cohort, there was no statistical differences in the 10-year DFS and OS between the ET+CT and ET only group (DFS: ET+CT 82.12% vs ET only 87.35% (*p*=0.99), OS: ET+CT 93.48% vs ET only 94.04% (*p*=0.88)). (Figure 4)
- In pT1-2N1 cohort, the ET only group tended to have poor DFS (ET+CT 77.03% vs ET only 54.17% (*p*=0.34)). The ET only group had poor OS compared to the ET+CT group (ET+CT 94.81% vs ET only 61.96%(*p*=0.01)). (Figure 4)

Conclusions

Although luminal HER2, HER2 and TN cohorts had no differences in prognosis between ILC and IDC, luminal ILC had a poor prognosis than luminal IDC. Therefore, luminal ILC needs stronger approach to improve their prognosis. And it was suggested that chemotherapy is effective for recurrent high-risk luminal ILC such as those with positive lymph node metastasis.

References

[1] Pestalozzi BC et al, Distinct Clinical and Prognostic Features of Infiltrating Lobular Carcinoma of the Breast: Combined Results of 15 International Breast Cancer Study Group Clinical Trials, J Clin Oncol. 2008; 26(18):3006–14

[2] Iorfida M et al. Invasive lobular breast cancer: subtypes and outcome. Breast Cancer Res Treat. 2012;133(2):713-23 [3] Carey et al. The triple negative paradox: primary tumor chemosensitivity of breast cancer subtypes. Clin Cancer Res, 2007. 13(8): 2329-34 [4] Hayashi, N., et al., Annual report of the Japanese Breast Cancer Registry for 2017. Breast Cancer, 2020. 27(5): 803-9

Acknowledgments

We would like to thank the Japanese Breast Cancer Society staff and National Clinical Database office staff for their support in this study. This presentation is the intellectual property of the author/presenter. Contract them at <u>yayoi.adachi@utsouthwestern.edu</u> for permission to reprint and/or distribute.

