

Original Research Article

Triple-Negative Breast Cancer Survival in Kurdish Patients

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Abstract: Triple negative breast cancers (TNBCs) are defined as the lack of expression of estrogen receptor (ER), progesterone receptor (PR), and human epidermal growth factor receptor 2 (HER-2). In this study we evaluated survival and clinicopathologic characteristics in Kurdish patients in Iran with TNBC. Between of 2009 and 2015, 58 patients with TNBC referred to Department of Radiation Oncology in Imam Reza Hospital, Kermanshah, Iran. We analyzed sex, age, type of pathology, grade tumor, laterality, stage, tumor size, overall survival and vascular, perineural, margin and skin invasion. The OS was plotted with GraphPad Prism 5 software and IBM SPSS version 19. The mean age for the patients at diagnosis was 44.29(±9.29), 100% female. Mean tumor size was 3.05 cm (range, 0.6-6 cm). Grade III was the most grades in TNBC (50%). Fifty four patients (93.1%) had invasive ductal carcinoma, one (1.7%) had invasive lobular carcinoma and 3 (5.2%) had invasive medullary. Sex, nineteen, twenty one and twelve patients were in stage I, II, III and IV, respectively. Twenty six patients did not have vascular invasion (n=46). Twenty six patients did not have perineuralinvasion (n=42). Forty two patients did not have margin invasion (n=50). Forty five patients did not have skin invasion (n=50).Overall survival for all of patients that the overall survival rate was 62%. Information about TNBC is low in Iran, especially in west of Iran. We need more research in this area because TNBC had a poorer overall survival than those without.

Keywords: Kurdish, Overall Survival, TNBC.

INTRODUCTION:

Breast cancer is a heterogeneous disease with a different characteristic such as age, tumor stage, lymph node involvement and pathologic grade [1]. Triple negative breast cancers (TNBCs) are defined as the lack of expression of estrogen receptor (ER), progesterone receptor (PR), and human epidermal growth factor receptor 2 (HER-2) [2]. These cancers occur in approximately 10% to 25% of all patients with breast cancer [4]. Around 6.6% of all BC cases are diagnosed in women less than 40 of age, 2.4% in women less than 35, and 0.65% in women less than 30 [3]. Among all the breast cancer subtypes, TNBC is associated with a worse prognosis. It has a characteristic recurrence pattern with the peak risk of recurrence and the majority of deaths occurring in the first 3 and 5 years after the initial treatment, respectively. Comparing to endocrine sensitive tumours, the risk for the late recurrence (beyond 5 years after the diagnosis) decreases by 50% [5]. In this study we evaluated survival and clinicopathologic characteristics in Kurdish patients in Iran with TNBC.

PATIENTS AND METHODS:

Between of 2009 and 2015, 58 patients with TNBC referred to Department of Radiation Oncology in Imam Reza Hospital, Kermanshah, Iran. We analyzed sex, age, type of pathology, grade tumor, laterality, stage, tumor size, overall survival and vascular, perineural, margin and skin invasion. HER2 positivity was defined as either HER2 gene amplification by fluorescent in situ hybridization or scored as 3+ by IHC. In case of HER2 2(+), fluorescent in situ hybridization was performed to determine HER2 positivity. TNBC was defined as ER (-), PR (-), and HER2 (-). The OS was plotted with GraphPad Prism 5 software and IBM SPSS version 19.

RESULTS:

The mean age for the patients at diagnosis was 44.29(±9.29), 100% female. Mean tumor size was 3.05 cm (range, 0.6-6 cm). Grade III was the most grades in TNBC (50%), following by grade I (10.3%) and II (39.7%). Fifty four patients (93.1%) had invasive ductal carcinoma, one (1.7%) had invasive lobular carcinoma and 3 (5.2%) had invasive medullary. Sex,

nineteen, twenty one and twelve patients were in stage I, II, III and IV, respectively. Location of tumor in thirty four patients was in left breast and twenty four patients were in right breast. Tumor size of nineteen patients was between 0.1 cm to 2 cm and tumor size of thirty three patients was between 2.1 cm to 5 cm and tumor size of six patients was greater than 5 cm. Twenty six patients did not have vascular invasion and twenty patients had vascular invasion (n=46). Twenty six

patients did not have perineuralinvasion and sixteen patients had perineuralinvasion (n=42). Forty two patients did not have margin invasion and eight patients had margin invasion (n=50). Forty-five patients did not have skin invasion and five patients had skin invasion (n=50) (Table 1).

The Figure 1 shows the OS for all of patients that the overall survival rate was 62%.

Table 1: Characteristics for the patients with triple negative breast cancer

| Characteristics | n(%) | Mean±SD | Range |
|---------------------------|----------|------------|-------|
| Age | | 44.29±9.29 | 25-68 |
| Sex | | | |
| Female | 58(100) | | |
| Male | 00(0) | | |
| Kind of Pathology | | | |
| Invasive Ductal Carcinoma | 54(93.1) | | |
| Lobular Carcinoma | 01(1.7) | | |
| Medullary Carcinoma | 03(5.2) | | |
| Laterality | | | |
| Left | 34(59) | | |
| Right | 24(41) | | |
| Stage | | | |
| I | 06(10) | | |
| II | 19(33) | | |
| III | 21(36) | | |
| IV | 12(21) | | |
| Grade | | | |
| I | 06(10.3) | | |
| II | 23(39.7) | | |
| III | 29(50) | | |
| Tumor Size | | | |
| 0.1-2 | 19(33) | | |
| 2.1-5 | 33(57) | | |
| >5 | 06(10) | | |
| Vascular | | | |
| Negative | 26(57) | | |
| Positive | 20(43) | | |
| Unknown | 12 | | |
| Perineural | | | |
| Negative | 26(62) | | |
| Positive | 16(38) | | |
| Unknown | 16 | | |
| Margin | | | |
| Negative | 42(84) | | |
| Positive | 08(16) | | |
| Unknown | 08 | | |
| Skin | | | |
| Negative | 45(90) | | |
| Positive | 05(10) | | |
| Unknown | 08 | | |

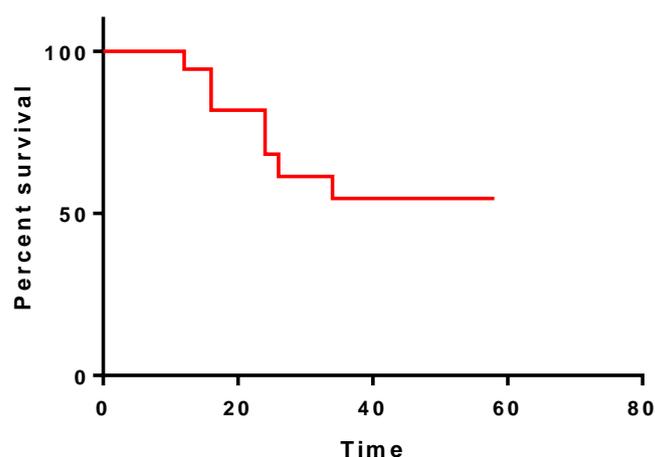


Fig 1: The 5-year overall survival for the patients with triple negative breast cancer

DISCUSSION:

Patients with TNBC had a poorer overall survival than those without. The reason for the poor prognosis of TNBC may be special biological characteristics such as younger age, higher rate of breast cancer family history, bigger tumor size, more advanced clinical stage upon diagnose, higher rate of lymph node metastasis, higher histological grade [16]. A study reported that Mean age of diagnosis of TNBC was found to be 46.26 years and other study [6]. This rate was found to be 47.64 and 49.99 by Liedtke et al [2]. Other study mean age of presentation for the patients was 44.5 years [7]. In our study, the mean age was 44.29 years and range of 25-68 years. Our study showed that the most percentage of TNBC were invasive ductal carcinoma (93.1%) and grade III (50%) that two studies [12, 13] had results similar to our study. Most patients in this study are located in stage III. Surgical margin status after breast conserving surgery is also an established risk factor for local recurrence, specifically in-breast recurrence [11]. Twenty six patients did not have vascular invasion (n=46). Twenty six patients did not have perineural invasion (n=42). Forty two patients did not have margin invasion (n=50). Forty five patients did not have skin invasion (n=50). The size of the tumor is also a well-known risk factor for recurrence [15]. These patients had larger tumors so that more than two-thirds of them were greater than 2 cm [14] like our study and mean tumor size was 3.05 cm. Two publications which have reported an overall 5-year survival of between 59-77% in patients with triple negative disease [8, 9]. Other study reported that the 5-year overall survival was 77% for African-American women and 72% for Caucasian women [10] in our patients in wSest of Iran is 62%.

CONCLUSION:

Information about TNBC is low in Iran, especially in west of Iran. We need more research in

this area because TNBC had a poorer overall survival than those without.

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REFERENCES:

1. Brenton JD, Carey LA, Ahmed AA, Caldas C; Molecular classification and molecular forecasting of breast cancer: ready for clinical application? *J Clin Oncol.*, 2005; 23(29):7350-60.
2. Liedtke C, Mazouni C, Hess KR, André F, Tordai A, Mejia JA; Response to neoadjuvant therapy and long-term survival in patient with triple-negative breast cancer. *J Clin Oncol.*, 2008; 26(8):1275-81.
3. Amirifard N, Choubsaz M, Sadeghi M, Sadeghi E; Epidemiology of breast cancer in young women in the West of Iran. *Sch. J. App. Med. Sci.*, 2015; 3(8D):2980-2984.
4. Baur KR, Brown M, Cress RD, Parise CA, Caggiano V; Descriptive analysis of estrogen receptor (ER)-negative, Progesterone receptor (PR)-negative, and HER2- negative invasive breast cancer ,the so called triple-negative phenotype: a population-based study from the California cancer registry. *Cancer*, 2007; 109(9):1721-8.
5. Ovcaricek T, Frkovic S, Matos E, Mozina B, Borstnar S; Triple negative breast cancer – prognostic factors and survival. *Radiol Oncol.*, 2011; 45(1): 46–52.
6. Sajid MT, Ahmed M, Azhar M, Mustafa QU, Shukr I, Ahmed M, et al.; Age-related frequency of triple negative breast cancer in women, *J Coll Physicians Surg Pak.*, 2014; 24 (6): 400-3.
7. Lakshmaiah KC, Das U, Suresh TM, Lokanatha D, Babu GK, Jacob LA, et al.; A study of triple negative breast cancer at a tertiary cancer care

- center in southern India, *Ann Med Health Sci Res.*, 2014;4 (6): 933-7.
8. Schwentner L, Wolters R, Koretz K, Wischnewsky MB, Kreienberg R, Rottscholl R, et al.; A: Triple-negative breast cancer: the impact of guideline-adherent adjuvant treatment on survival – a retrospective multi-centre cohort study. *Breast Cancer Research and Treatment*, 2012; 132: 1073-1080.
 9. Dent R, Trudeau M, Pritchard KI, Hanna WM, Kahn HK, Sawka CA, et al.; Triple-negative breast cancer: clinical features and patterns of recurrence. *Clinical Cancer Research*, 2007; 13: 4429-4434.
 10. Chu QD, Henderson AE, Ampil F, Li BD; Outcome for patients with triple-negative breast cancer is not dependent on race/ethnicity. *International Journal of Breast Cancer*, 2012; 2012: 764570.
 11. Bartelink H, Maingon P, Poortmans P, Weltens C, Fourquet A, Jager J et al.; Whole-breast irradiation with or without a boost for patients treated with breast-conserving surgery for early breast cancer: 20-year follow-up of a randomised phase 3 trial. *Lancet Oncol*, 2015; 16(1):47-56.
 12. Basu S, Chen W, Tchou J, Mavi A, Cermik T, Czerniecki B, et al.; Comparison of triple-negative and estrogen receptor-positive/progestrone receptorpositive/HER2-negative breast carcinoma using quantitative fluorine-18 fluorodeoxyglucose/positron emission tomography imaging parameters. *American Cancer Society (ACS)*, 2007; 112(5):995-1000.
 13. Rakha EA, El-sayed ME, Green AR, Lee AH, Robertson JF, Ellis IO; Prognostic markers in triple-negative breast cancer. *Cancer*, 2007; 109(1):25-32.
 14. Ann J, Kang S, Kwun K; Clinicopathologic characteristics of triple-negative breast cancer in early stages. *Eur J Cancer Supp.*, 2008; 6(7):183.
 15. Kunkler IH, Williams LJ, Jack WJ, Cameron DA, Dixon JM; Breast-conserving surgery with or without irradiation in women aged 65 years or older with early breast cancer (PRIME II): a randomised controlled trial. *Lancet Oncol*., 2015;16:266-73.
 16. Guan ZY, Wang SS, Gao Y, Su ZY, Luo WB, Guan ZZ; Clinical characteristics and prognosis of triple-negative breast cancer: a report of 305 cases. *Chin J Cancer*, 2008; 27(6):561–565.