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Comparison of clinicopathologic features and survival in young American women aged 18–39 years in different ethnic groups with breast cancer

P Liu¹, X Li¹, E A Mittendorf², J Li¹, X L Du³, J He⁴, Y Ren⁴, J Yang⁵, K K Hunt² and M Yi^{1,2}

¹Department of Translational Medicine, The First Affiliated Hospital of Xian Jiaotong University, School of Medicine, 277 West Yanta Road, Xian, Shaanxi 710061, China; ²Department of Surgical Oncology, The University of Texas MD Anderson Cancer Center, Houston, TX, USA; ³Division of Epidemiology, Human Genetics, and Environmental Sciences, The University of Texas School of Public Health, Houston, TX, USA; ⁴Department of Surgical Oncology, The First Affiliated Hospital of Xian Jiaotong University, School of Medicine, Xian, China and ⁵Department of Medical Oncology, The First Affiliated Hospital of Xian Jiaotong University, School of Medicine, Xian, China

Background: Ethnic disparities in breast cancer diagnoses and disease-specific survival (DSS) rates in the United States are well known. However, few studies have assessed differences specifically between Asians American(s) and other ethnic groups, particularly among Asian American(s) subgroups, in women aged 18–39 years.

Methods: The Surveillance, Epidemiology, and End Results database was used to identify women aged 18–39 years diagnosed with breast cancer from 1973 to 2009. Incidence rates, clinicopathologic features, and survival among broad ethnic groups and among Asian subgroups.

Results: A total of 55 153 breast cancer women aged 18–39 years were identified: 63.6% non-Hispanic white (NHW), 14.9% black, 12.8% Hispanic-white (HW), and 8.7% Asian. The overall incidence rates were stable from 1992 to 2009. Asian patients had the least advanced disease at presentation and the lowest risk of death compared with the other groups. All the Asian subgroups except the Hawaiian/Pacific Islander subgroup had better DSS than NHW, black, and HW patients. Advanced tumour stage was associated with poorer DSS in all the ethnic groups. High tumour grade was associated with poorer DSS in the NHW, black, HW, and Chinese groups. Younger age at diagnosis was associated with poorer DSS in the NHW and black groups.

Conclusion: The presenting clinical and pathologic features of breast cancer differ by ethnicity in the United States, and these differences impact survival in women younger than 40 years.

Breast cancer is the most common cancer in women and the leading cause of cancer death among women in all ethnic groups worldwide (NHS, 2013). However, breast cancer is uncommon in young women: only $\sim 7\%$ of all breast cancers are diagnosed in women younger than 40 years, and fewer than 4% are diagnosed in women younger than 35 years (Chung *et al*, 1996; Brinton *et al*,

2008). Nonetheless, a recent study showed a small but significant increase in the incidence of breast cancer with distant disease in the United States in relatively young women aged 25–39 years from 1976 to 2009 (Johnson *et al*, 2013), causing some concern and prompting further studies to confirm the findings and to determine the potential reasons.

*Correspondence: Dr M Yi; E-mail: myi@mdanderson.org

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Numerous studies have demonstrated that breast tumours in younger women have a more aggressive biology, which correlates with poorer outcomes when compared with older women (Nixon et al, 1994; Chung et al, 1996; Dubsky et al, 2002; Han et al, 2004; Aebi and Castiglione, 2006). However, data regarding disparities among these young women are limited. The few studies that have been published have focused on comparisons among black, Hispanic, and white women (Newman et al, 2002; Shavers et al, 2003). In a study of breast cancer patients of all ages, our group previously found that patients in all Asian subgroups were younger at diagnosis than NHW patients except Japanese patients, and patients in most Asian subgroups had similar survival rates compared with NHW women except Japanese patients had a better survival rate and Hawaiian/Pacific Islander patients had a worse survival rate (Yi et al, 2012). The current study was performed to identify disparities specifically in young women. Using age 39 years as a cutoff, we sought to identify differences in disease presentation, clinicopathologic features, and survival among the broad ethnic groups and among the Asian subgroups.

PATIENTS AND METHODS

Patient selection and data collection. The Surveillance Epidemiology and End Results (SEER) database of the National Cancer Institute was used to identify patients with primary tumour sites coded as C50.0–C50.9 (breast) between 1973 and 2009. Data were obtained from all 18 US cancer registries participating in the SEER program using SEER*Stat software version 8.0.2 (http://seer.cancer.gov/seerstat). Male patients and patients whose race was coded as 'American Indian/Alaska Native', 'Other unspecified (1991 +)', or 'Unknown' were excluded. Ethnicity was categorised into four broad groups: non-Hispanic-white (NHW), black, Hispanic white (HW), and Asian. Eight subgroups of Asian patients were identified: Filipino, Chinese, Japanese, Asian Indian/Pakistani, Korean, Vietnamese, Hawaiian/Pacific Islander, and others.

Statistical analysis. Age-adjusted breast cancer SEER incidence rates in women aged 18–39 years and older than 39 years were calculated separately by using SEER*Stat software version 8.0.2 and data from incidence – SEER 13 (1992–2009). A χ^2 test was used to assess differences in categorical variables (disease stage, surgery type, tumour grade, hormone receptor status and radiation treatment status) and a Kruskal–Wallis equality-of-populations rank test was used to assess differences in continue variables (age at diagnosis) among the broad ethnic groups and among the eight Asian subgroups.

Overall survival (OS) was calculated from the date of diagnosis to the date of death, date last known to be alive, or 30 November 2009, whichever occurred first. Disease-specific survival (DSS) was calculated from the date of diagnosis to the date of breast cancerrelated death, date last known to be alive, or 30 November 2009. Overall survival and DSS curves were calculated using the Kaplan–Meier method. Patients who were lost to follow-up or who survived beyond 30 November 2009, were censored.

Multivariate Cox proportional hazards models were used to determine the influence of patient, tumour, and treatment factors of known or potential prognostic value on DSS in each broad ethnic group and in each Asian subgroup with a sample of >500 patients. Stata SE version 12.0 statistical software (StataCorp LP, College Station, TX, USA) was used for statistical analyses. All tests were two-tailed, and statistical significance was set at P < 0.05.

RESULTS

Patient and tumour characteristics. The SEER database revealed 971 565 patients diagnosed with breast cancer from 1973 to 2009, including 55 908 (5.8%) aged 18–39 years. Hispanic white had the highest percentage of patients whose age at diagnosis was between 18 and 39 years old (10.8%) while NHW had the lowest percentage (4.7%, black patients (9.5%), and Asian patients (8.6%)). Of these, 55 153 patients belonged to 1 of the 4 broad ethnic groups being evaluated in this study: 35 101 (63.6%) NHW, 8215 (14.9%) black, 7067 (12.8%) HW, and 4770 (8.7%) Asian.

The clinicopathologic characteristics of patients in the broad ethnic groups are shown in Table 1. Our analyses revealed differences among the groups with respect to disease stage, surgery type, tumour grade, oestrogen receptor (ER) status, progesterone receptor (PR) status, and radiation treatment status. The incidence of localised tumours was higher in Asian patients (52.9%) than in NHW (51.6%, P = 0.1), black (44.3%, P < 0.0001), and HW (44.3%, P < 0.0001) patients. The incidence of distant disease was higher in black patients (8.3%) than in patients in any of the other three groups (NHW 4.6%, HW 6.8%, and Asian 4.8%). At least 55% of patients in each broad ethnic group had undergone total mastectomy. The incidence of grade III and IV tumours was lowest in the Asian patients (56.3%) and highest in the black patients (70.5%). At least 75% of patients in each broad ethnic group had invasive ductal carcinoma. The incidence of ER-positive tumours was highest in the Asian patients (67.8%) and lowest in the black patients (49.9%). In all, 70% Asian patients received radiation therapy after segmental mastectomy while only 62.2% black patients received radiation theory after segmental mastectomy (P < 0.0001).

The Asian patients were categorised into eight subgroups as follows: Filipino (21.8%), Chinese (20.9%), Japanese (12.6%), Hawaiian/Pacific Islander (11.3%), Korean (7.1%), Asian Indian/Pakistani (8.6%), Vietnamese (6.4%), and other (11.2%). Table 2 lists the clinicopathologic characteristics of patients in the eight subgroups. Disease stage at presentation, tumour histology and grade, and ER and PR status were different among the subgroup. Japanese patients generally had more favourable clinicopathologic features: they had the highest incidence of localised disease (59.3%) and grade I tumours (12.3%). Asian Indian/Pakistani women generally had less favourable clinicopathologic features: they had the highest incidence of regional/distant disease (56.3%) and grade III/IV tumours (64.2%) and the lowest incidence of ER-positive tumours (61.0%). As in the broad ethnic group comparisons, the majority of patients in all the subgroups had undergone surgery. The smallest proportion of patients who did not undergo surgery was in the Japanese subgroup (2.0%), whereas the largest proportion of patients who did not undergo surgery was in the Asian Indian/Pakistani subgroup (5.9%). There were no difference in received radiation therapy after segmental mastectomy among Asian subgroups (P = 0.8). In all, 45% Asian Indian/Pakistani received radiation therapy after total mastectomy while only 26.5% Vietnamese received radiation therapy (P < 0.0001).

Breast cancer incidence rates. Figure 1 shows age-adjusted breast cancer incidence rates in women aged 18–39 years (A) and older than 39 years (B) from the SEER 13 registries (1992–2009). Although the incidence rates for patients older than 39 years decreased during this time period, the incidence rates for women aged 18–39 years were stable. In this younger age group, black patients had the highest incidence rates, and two dramatic increases were noted in 2000 and 2004. In young NHW women, there was a slight increase from 1992 to 2009, whereas the

Table 1. Comparison of patient, tumour, and treatment characteristics among the four broad ethnic groups of women aged 18–39 years with breast cancer

Characteristic	Non-Hispanic white $(n = 35101)$	Black (n = 8215)	Hispanic white $(n = 7067)$	Asian (<i>n</i> = 4770)	P -value
Disease stage					<0.0001a
Localised	17 602 (51.6)	3538 (44.3)	3035 (44.3)	2477 (52.9)	
Regional	14 936 (43.8)	3785 (47.4)	3356 (48.9)	1978 (42.3)	
Distant	1581 (4.6)	659 (8.3)	465 (6.8)	225 (4.8)	
Unknown	982	233	211	90	
Age at diagnosis (years)					0.0001 ^b
Mean (median)	35 (36)	34.4 (35)	34.5 (35)	35 (36)	
<35 years	12 421 (35.4)	3499 (42.6)	2930 (41.5)	1718 (36.0)	< 0.0001
≥35 years	22 680 (64.6)	4716 (57.4)	4137 (58.5)	3052 (64.0)	
Surgery					<0.0001a
No primary surgery	1030 (3.0)	583 (7.2)	477 (6.8)	175 (3.7)	
SM	11 835 (34.2)	2929 (36.2)	2584 (36.9)	1775 (37.5)	
TM	21 731 (62.8)	4584 (56.6)	3948 (56.3)	2780 (58.8)	
Unknown	505	119	58	40	
Tumour grade					< 0.0001ª
I	1816 (7.3)	278 (4.6)	336 (5.7)	336 (8.8)	
II	7905 (31.8)	1521(24.9)	1744 (29.5)	1338 (35.0)	
III	14 045 (56.4)	4079 (66.8)	3553 (60.1)	2018 (52.7)	
IV	1119 (4.5)	225 (3.7)	276 (4.7)	135 (3.5)	
Unknown	10216	2112	1158	943	
Histology					< 0.0001
IDC	26 973 (76.8)	6227 (75.8)	5441 (77.0)	3793 (79.5)	
ILC	1170 (3.3)	194 (2.4)	186 (2.6)	97 (2.0)	
IDC and ILC	1522 (4.3)	231 (2.8)	337 (4.8)	194 (4.1)	
Other	5436 (15.5)	1563 (19.0)	1103 (15.6)	686 (14.4)	
Oestrogen receptor status					<0.0001 ^a
Positive	13 362 (62.0)	2667 (49.9)	3035 (57.8)	2379 (67.8)	
Negative	8010 (37.2)	2636 (49.3)	2185 (41.7)	1115 (31.7)	
Borderline	172 (0.8)	42 (0.8)	26 (0.5)	17 (0.5)	
Unknown	13 557	2870	1821	1259	
Progesterone receptor status					<0.0001 ^a
Positive	11 995 (56.3)	2280 (43.2)	2606 (50.5)	2180 (63.1)	
Negative	9117 (42.8)	2941 (55.8)	2514 (48.7)	1241 (35.9)	
Borderline	198 (0.9)	51 (1.0)	41 (0.8)	35 (1.0)	
Unknown	13791	2943	1906	1314	
Adjuvant radiation therapy					< 0.0001
No	21 357 (60.8)	5105 (62.1)	4206 (59.5)	2722 (57.1)	
Yes	13 744 (39.2)	3110 (37.9)	2861 (40.5)	2048 (42.9)	
Adjuvant radiation therapy after SM					< 0.0001
No	3576 (30.2)	1107 (37.8)	971 (37.6)	532 (30.0)	
Yes	8259 (69.8)	1822 (62.2)	1613 (62.4)	1243 (70.0)	
Adjuvant radiation therapy after TM					< 0.0001
No	16 318 (75.1)	3331 (72.7)	2721 (68.9)	1984 (71.4)	
Yes	5413 (24.9)	1253 (27.3)	1227 (31.1)	796 (28.6)	

 $Abbreviations: IDC = invasive \ ductal \ cancer; \ ILC = invasive \ lobular \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy. \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy. \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy \ ductal \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy \ ductal \ duc$

 $\mathbf{b}_{\mathsf{Kruskal-Wallis}} \ \mathsf{equality-of-populations} \ \mathsf{rank} \ \mathsf{test}.$

incidence rates in HW women were slightly decreased. Incidence rates in Asian women were decreased from 1992 to 1997, increased from 1997 to 2001, and aside from an increase in 2008 in the 18–39 age group, then decreased again. Incidence rates in the Asian subgroups were not available.

Survival. The median follow-up for the study cohort was 6 years. The 5- and 10-year DSS and OS rates are shown in Table 3. Korean and Japanese patients had the best 5-year DSS rates (89.3% and 86.0%, respectively), and black and Hawaiian/Pacific Islander patients had the poorest 5-year DSS rates (71.8% and 76.1%,

^aP-value calculated after excluded unknown category.

Table 2. Comparison of patient, tumour, and treatment characteristics among subgroups of Asian women (n = 4770) aged 18–39 years with breast cancer

Characteristic	Filipino (n = 1040)	Japanese $(n = 602)$	Chinese $(n = 995)$	Hawaiian/Pacific Islander (n = 540)	Korean (n = 339)	Asian Indian/ Pakistani (n = 410)	Vietnamese (<i>n</i> = 307)	Other (n = 537)	P- value
Disease stage						,			< 0.000
Localised	529 (51.9)	352 (59.3)	536 (55.1)	257 (49.0)	172 (51.7)	176 (43.7)	174 (57.2)	281 (53.2)	
Regional	432 (42.3)	223 (37.5)	397 (40.8)	230 (43.8)	150 (45.0)	206 (51.1)	117 (38.5)	223 (42.2)	
Distant	59 (5.8)	19 (3.2)	40 (4.1)	38 (7.2)	11 (3.3)	21 (5.2)	13 (3.3)	24 (4.6)	
Unknown	20	8	22	15	6	7	3	9	
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Age at diagnosis									0.06ª
(years)									
Mean (median)	35.1 (36)	35.2 (36)	35.1 (36)	34.7 (36)	35.3 (36)	34.4 (35)	34.9 (36)	35.2 (36)	
< 35	368 (35.4)	203 (33.7)	349 (35.1)	209 (38.7)	114 (33.6)	174 (42.4)	115 (37.5)	184 (34.6)	0.08
≥35	672 (64.6)	399 (66.3)	646 (64.9)	331 (61.3)	225 (66.4)	236 (57.6)	192 (62.5)	351 (65.4)	
Surgery									0.001
No primary surgery	37 (3.6)	12 (2.0)	33 (3.3)	25 (4.7)	11 (3.2)	24 (5.9)	7 (2.3)	26 (4.8)	
SM	357 (34.5)	227 (38.2)	370 (37.5)	183 (34.3)	151 (44.7)	173 (42.4)	119 (38.8)	195 (36.8)	
TM	639 (61.9)	355 (59.8)	584 (59.2)	325 (61.0)	176 (52.1)	211 (51.7)	181 (58.9)	309 (58.3)	
Unknown	7	8	8	7	1	2	0	1	
Tumour grade		-	-				-		0.002 ^k
I grade	67 (8.1)	53 (12.3)	68 (8.7)	37 (8.9)	32 (11.1)	21 (5.6)	18 (6.5)	40 (9.0)	0.002
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II	286 (34.7)	156 (36.2)	288 (37.0)	138 (33.3)	86 (30.0)	112 (30.2)	102 (36.8)	170 (38.3)	
III	442 (53.6)	200 (46.4)	406 (52.2)	225 (54.2)	152 (53.0)	227 (61.2)	145 (52.4)	221 (49.8)	
IV	29 (3.5)	22 (5.1)	16 (2.1)	15 (3.6)	17 (5.9)	11 (3.0)	12 (4.3)	13 (2.9)	
Unknown	216	171	217	125	52	21	18	40	
Histology									0.03
IDC	817 (78.6)	498 (82.7)	785 (78.9)	427 (79.1)	273 (80.5)	324 (79.0)	242 (78.8)	427 (79.5)	
ILC	21 (2.0)	10 (1.7)	19 (1.9)	9 (1.7)	6 (1.8)	14 (3.4)	1 (0.3)	17 (3.2)	
IDC and ILC	41 (3.9)	17 (2.8)	43 (4.3)	20 (3.7)	20 (5.9)	24 (5.9)	17 (5.5)	12 (2.2)	
Other	161 (15.5)	77 (12.8)	148 (14.9)	84 (15.5)	40 (11.8)	48 (11.7)	47 (15.3)	81 (15.1)	
Oestrogen									0.08 ^b
receptor status									
Positive	513 (68.4)	247 (68.0)	510 (70.7)	266 (68.6)	183 (66.1)	216 (61.0)	175 (68.4)	286 (71.1)	
Negative	237 (31.6)	116 (32.0)	211 (29.3)	122 (31.4)	94 (33.9)	138 (39.0)	81 (31.6)	116 (28.9)	
Unknown	290	239	274	152	62	56	51	135	
Progesterone									0.1 ^b
receptor status									0.1
Positive	465 (63.7)	232 (64.4)	478 (67.0)	251 (65.9)	171 (63.1)	199 (56.9)	160 (62.7)	259 (65.6)	
Negative	265 (36.3)	128 (35.6)	236 (33.0)	130 (34.1)	100 (36.9)		95 (37.3)	136 (34.4)	
Unknown	310	242	281	159	68	60	52	142	
Adjuvant radiation therapy									< 0.0001
No	605 (58.2)	337 (56.0)	567 (57.0)	323 (59.8)	186 (54.9)	190 (46.3)	176 (57.3)	338 (62.9)	
Yes	435 (41.8)	265 (44.0)	428 (43.0)	217 (40.2)	153 (45.1)	220 (53.7)	131 (42.7)	199 (37.1)	
Adjuvant radiation			· · ·			1 1			0.8
therapy after SM									0.0
No	106 (29.7)	65 (28.6)	101 (27.3)	60 (32.8)	47 (31.1)	50 (28.9)	37 (31.1)	66 (33.8)	
Yes	251 (70.3)	162 (71.4)	269 (72.7)	123 (67.2)	104 (68.9)	123 (71.1)	82 (68.9)	129 (66.2)	
Adjuvant radiation therapy after TM									< 0.0001
No	457 (71.5)	252 (71.0)	426 (73.0)	233 (71.7)	128 (72.7)	116 (55.0)	133 (73.5)	239 (77.3)	
Yes	182 (28.5)	103 (29.0)	158 (27.0)	92 (28.3)	48 (27.3)		48 (26.5)	70 (22.7)	

 $Abbreviations: IDC = invasive \ ductal \ cancer; \ ILC = invasive \ lobular \ cancer; \ SM = segmental \ mastectomy; \ TM = total \ mastectomy.$

^aKruskal-Wallis equality-of-populations rank test.

bExclude unknown category.

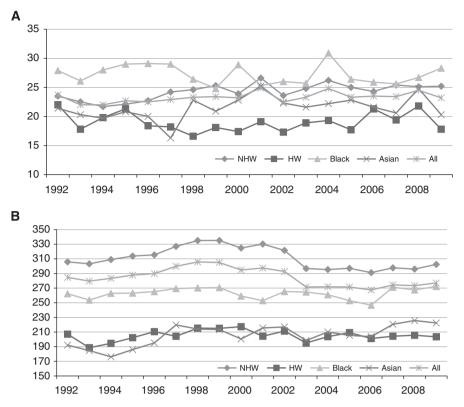


Figure 1. Age-adjusted breast cancer incidence rates in women aged 18–39 years (**A**) and older than 39 years (**B**) from the SEER 13 registries (1992–2009). Rates are per 100 000 and age adjusted to the 2000 US standard population (19 age groups; Census P25-1130) standard.

respectively). All Asian subgroups except Hawaiian patients (75.9%) had a better 5-year OS rate than did NHW, black, and HW patients (80.0%, 69.4%, and 77.0%, respectively).

Table 4 shows the association between clinicopathologic factors and DSS in the four broad ethnicity groups. Advanced tumour stage, high tumour grade, and negative ER status were highly associated with poorer DSS in each ethnic group. Age younger than 35 years was associated with poorer DSS in NHW and black patients but not in HW or Asian patients. Table 5 shows the association between clinicopathologic factors and DSS in four Asian ethnicity subgroups that had a sample of >500 patients (Filipino, Japanese, Chinese, and Hawaiian/Pacific Islander). Advanced tumour stage was highly associated with poorer DSS in all four subgroups. Higher tumour grade was associated with poorer DSS in only in the Chinese subgroup. The ER-negative status was associated with DSS in the Hawaiian subgroup only. Age at diagnosis was not associated with DSS in these four Asian subgroups.

DISCUSSION

The current study represents one of the most comprehensive population-based analyses of breast cancer patients aged 18–39 years evaluated by ethnicity. Our study included >55 000 (5.8%) breast cancer women aged 18–39 years, which is comparable to the 7% of breast cancer patients aged 18–39 years reported in other studies (Chung et al, 1996; Brinton et al, 2008). Consistent with other studies (Gray et al, 1980; Pathak et al, 2000; Joslyn et al, 2005; Brinton et al, 2008) our study showed that black women had higher incidence rates of breast cancer than NHW women before age 40, whereas NHW women had higher incidence rates than black women after age 39 (crossover pattern). The HW and Asian

women had lower breast cancer incidence rates than NHW women in both age groups and did not exhibit the crossover pattern observed among black women. Interestingly, for women aged 18-39 years, the incidence rates were increased only in NHW women from 1992 to 2009. This observation must be put into context, however, as in NHW patients, only 4.7% of breast cancers were diagnosed in patients aged 18-39 years, compared with 10.8% in the HW patients, 9.5% in the black patients, and 8.6% in the Asian patients, numbers that are consistent with previously published reports (Chung et al, 1996; Brinton et al, 2008; Telli et al, 2011; Yi et al, 2012). It is possible the overall age distribution of Asian women is younger compared with the other ethnic groups in the United States, resulting in a disproportionate breakdown by age. This could be due to immigration patterns, as the Asian population reflects the rapid increases in younger age in the latter half of the twentieth century.

Interestingly, the Asian patients in our study presented with less advanced stages of disease and had a lower risk of death compared with patients in the other ethnic groups. This finding conflicts with the findings of previous studies that included patients of all ages, which showed that Asian patients were more likely to have advanced disease than NHW patients but less likely to have advanced disease than black and HW patients (Li et al, 2003; Martinez et al, 2007; Berz et al, 2009; Ooi et al, 2011; Yi et al, 2012). Among the Asian subgroups, Asian Indian/Pakistani patients had the highest proportion of patients with advanced disease, high tumour grades, and ER- and PR-negative tumours, but their mortality risk was similar to that in the other subgroups, which is consistent with previous studies evaluating Asian women of all ages (Moran et al, 2011; Yi et al, 2012). Hawaiian/Pacific Islander patients had the poorest DSS of all the Asian subgroups even though they were comparable to the other subgroups in terms of disease stage and tumour grade, which is consistent with the findings we previously published from a study evaluating patients

Table 3. Survival (5-year and 10-year) rates for each ethnic group of women aged 18-39 years with breast cancer Hawaiian/ Asian Pacific Indian/ NHW Black HW Asian Filipino Japanese Chinese Islander Korean Pakistani Vietnamese Other (n = 7067)(n = 4770)(n = 995)(n = 540) (n = 1040)(n = 339)(n = 410)(n = 307)(n = 35101)(n = 8215)(n = 602)(n = 537)(%) (%) (%) (%) (%) (%) (%) (%) (%) (%) (%) (%) Disease-specific survival 81.4 71.8 79.1 84.6 82.9 86.0 85.6 89.3 84.9 85.6 90.3 5-Year 76.1 10-Year 73.0 70.3 77.2 73.0 80.2 78.3 67.9 83.8 73.8 80.6 63.9 86.5 Overall survival 5-Year 80.0 69 4 77.0 83.5 811 86.0 84 8 746 88.3 82.5 86.1 89 5 10-Yea 70.6 60.2 67.2 74.7 69 9 79.1 76.5 65.2 80.7 68.7 80.8 82.4 Abbreviations: HW = Hispanic white; NHW = non-Hispanic white

Table 4. Multivariate analysis of clinicopathologic variables associated with disease-specific survival in women with breast cancer aged 18–39 years in four broad ethnic groups using a Cox proportional hazards model

	NHW (n = 35 101)		Black (<i>n</i> = 8215)		HW (n=7067)		Asian (n = 4770)	
Variable	HR	P -value	HR	P -value	HR	P -value	HR	P -value
Disease stage								•
Localised	Referent							
Regional	2.43	< 0.001	2.89	< 0.001	2.53	< 0.001	3.25	< 0.001
Distant	8.19	< 0.001	8.29	< 0.001	7.59	< 0.001	12.15	< 0.001
Tumour grade								
I	Referent							
II	2.37	< 0.001	1.21	0.4	1.65	0.05	2.58	0.006
III	3.40	< 0.001	1.66	0.02	2.06	0.005	3.55	< 0.001
IV	3.13	< 0.001	1.70	0.05	2.31	0.005	3.52	0.002
Histology								
IDC	Referent							
ILC	1.40	0.005	1.87	0.02		NS		NS
IDC and ILC	1.08	0.4	1.10	0.6		NS		NS
Other	0.93	0.3	0.88	0.2		NS		NS
Negative ER status	1.14	0.02	1.46	< 0.001	1.33	0.008	1.44	0.01
Negative PR status	1.12	0.02		NS		NS		NS
Age <35 years	1.09	0.02	1.14	0.03	1.1	0.3	1.12	0.3
Adjuvant radiation therapy	1.07	0.07	0.76	< 0.001		NS		NS
Primary surgery								
SM	Referent							
TM	1.38	< 0.001	1.20	0.01	1.63	< 0.001	1.37	0.006
None	2.63	< 0.001	1.38	0.04	2.70	< 0.001	2.51	0.002

Abbreviations: ER = oestrogen receptor; HR = hazard ratio; HW = Hispanic white; IDC = invasive ductal cancer; ILC = invasive lobular cancer; NHW = non-Hispanic white; NS = not significant; PR = progesterone receptor; SM = segmental mastectomy; TM = total mastectomy.

in all age groups (Yi *et al*, 2012). These data suggest that this subgroup may experience issues related to access to care (i.e., screening and follow-up) rather than differences in tumour biology (Ooi *et al*, 2011).

Our study also showed that age as a risk factor for DSS varied between ethnic groups. Although being younger than 35 at diagnosis was associated with poorer DSS in NHW and black patients, it was not associated with DSS in HW patients or Asian patients. Our study also showed that there was no survival difference in Asian subgroups when compared patients younger than 35 with 35–39, which is consistent with the studies conducted in Japan (Yoshida *et al*, 2011) and Korean (Kim *et al*, 2007). This may suggest that, in HW and Asian groups, tumour biology and age are independent prognostic factors for survival. An interesting observation in our study is the differences in biologic factors between Asians and other ethnic groups, specifically in tumour grade, ER/PR status, and % presenting with localised disease. Previous studies have shown that certain breast cancer subtypes to

Table 5. Multivariate analysis of clinicopathologic variables associated with disease-specific survival in women with breast cancer aged 18–39 years in Asian subgroups with a sample size of >500 using a Cox proportional hazards model

	Filipino (<i>n</i> = 1040)		Chinese (n = 995)		Japanese (<i>n</i> = 602)		Hawaiian/Pacific Islander (n = 540)	
Variable	HR	P -value	HR	P -value	HR	P -value	HR-value	P -value
Disease stage								
Localised	Referent							
Regional	2.8	< 0.001	3.25	< 0.001	4.11	< 0.001	3.92	< 0.001
Distant	10.3	< 0.001	16.9	< 0.001	9.44	< 0.001	4.37	0.01
Tumour grade								
1	Referent							
II		NS	1.96	0.3		NS		NS
III		NS	2.77	0.09		NS		NS
IV		NS	4.35	0.046		NS		NS
Histology								
IDC	Referent							
ILC		NS	7.32	0.001		NS		NS
IDC and ILC		NS	0.57	0.2		NS		NS
Others		NS	1.09	0.7		NS		NS
Negative ER status		NS		NS		NS	2.02	0.003
Negative PR status		NS		NS		NS		NS
Age <35 years		NS		NS		NS		NS
Adjuvant radiation therapy		NS		NS		NS		NS
Primary surgery	•	•	•		•			
SM	Referent							
TM	1.70	0.002		NS		NS	1.92	0.02
None	4.84	< 0.001		NS		NS	7.46	0.004

Abbreviations: ER=oestrogen receptor; HR=hazard ratio; IDC=invasive ductal cancer; ILC=invasive lobular cancer; NS=not significant; PR=progesterone receptor; SM=segmental mastectomy; TM=total mastectomy;

include triple-negative breast cancer (Bauer *et al*, 2007; Millikan *et al*, 2008) occur more often in young patients. Although population-based studies have identified a higher proportion of triple negative breast cancers in premenopausal Black women (Carey *et al*, 2006; Bauer *et al*, 2007; Morris *et al*, 2007), the relationship between race and tumour biology has not been completely elucidated. Ongoing work by our group is further investigating the relationship among age, race, and tumour biology in Asian patients.

Because this study was a retrospective investigation of a population-based database, data regarding socioeconomic status, family history of breast cancer, lifestyle factors, Her2 status, and the administration of neoadjuvant or adjuvant systemic therapies were limited. This prevented us from evaluating these factors as potential confounders or effect modifiers of the relationships observed. In addition, SEER age-adjusted incidence rates were limited to the four broad ethnic groups only because no data were available for Asian subgroups. Incidence rates were also limited to the time period from 1992 to 2009, when data were available. The long duration of our study period (1973–2009) may affect the results due to missing data and change of management. Instead of using TNM stage, we only used localised, regional, distant as disease stage. We cannot provide information about non-invasive disease or DCIS in our study.

In conclusion, our study demonstrates that presenting clinical and pathologic features of breast cancer differ by ethnicity in the United States and that these differences impact survival in women younger than age 40 years.

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