

Reproductive and Hormonal Risk Profile According to Language Acculturation and Country of Residence in the *Ella* Binational Breast Cancer Study

Jesse N. Nodora, DrPH,¹ Linda Gallo, PhD,² Renee Cooper, BA,² Betsy C. Wertheim, MS,³ Loki Natarajan, PhD,¹ Patricia A. Thompson, PhD,³ Ian K. Komenaka, MD,⁴ Abenaa Brewster, MD,⁵ Melissa Bondy, PhD,⁶ Adrian Daneri-Navarro, PhD,⁷ María Mercedes Meza-Montenegro, PhD,⁸ Luis Enrique Gutierrez-Millan, PhD,⁹ and María Elena Martínez, PhD¹

Abstract

Background: We compared the distribution of breast cancer reproductive and hormonal risk factors by level of acculturation and country of residence in women of Mexican descent.

Methods: To compare the distribution of breast cancer reproductive and hormonal risk factors by level of acculturation and country of residence in women of Mexican descent, taking into account level of education, we analyzed data on 581 Mexican and 620 Mexican American (MA) women with a history of invasive breast cancer from the *Ella* Binational Breast Cancer Study. An eight-item language-based acculturation measure was used to classify MA women. Multivariate logistic regression was used to test associations between language acculturation, country of residence, and reproductive and hormonal risk factors.

Results: After adjustment for age and education, compared to women residing in Mexico, English-dominant MAs were significantly more likely to have an earlier age at menarche (< 12 years; odds ratio [OR] = 2.08; 95% confidence interval [CI], 1.30–3.34), less likely to have a late age at first birth (≥ 30 years; OR = 0.49; 95% CI, 0.25–0.97), and less likely to ever breastfeed (OR = 0.13; 95% CI, 0.08–0.21).

Conclusions: Differences in reproductive and hormonal risk profile according to language acculturation and country of residence are evident; some of these were explained by education. Results support continued efforts to educate Mexican and MA women on screening and early detection of breast cancer along with promotion of modifiable factors, such as breastfeeding.

Introduction

GLOBALLY, BREAST CANCER is the most common cancer in women causing approximately 460,000 deaths in 2008.¹ Since 1980, incidence and mortality rates for breast cancer have been increasing.² The most notable increases have occurred in low- and middle-income countries, which accounted for 45% of all newly diagnosed cases of breast cancer in 2009. Although more developed countries have a higher incidence rate (67.8/100,000) than less developed

countries (23.8/100,000),³ there are disproportionately more deaths reported in low-income countries (58% of the total deaths in 2008) than in high-income countries (15% of the total deaths in 2008) among younger women (< 50 years of age).² Breast cancer mortality rates in women living in Mexico, a middle-income country, increased from 7.67 to 9.20 per 100,000 from 1985–1999 and then leveled off to 9.04 per 100,000 in 2005–2007.⁴ Detailed regional mortality analyses show that these increases are expanding beyond developed regions and into underdeveloped parts of Mexico.⁵

¹Moore's University of California San Diego Cancer Center, University of California, San Diego, La Jolla, California.

²San Diego State University, San Diego, California.

³Arizona Cancer Center, University of Arizona, Tucson, Arizona.

⁴Department of Surgery, Maricopa Medical Center, Phoenix, Arizona.

⁵University of Texas M.D. Anderson Cancer Center, Houston, Texas.

⁶Department of Pediatrics, Dan L. Duncan Cancer Center, Baylor College of Medicine, Houston, Texas.

⁷University of Guadalajara, Guadalajara, México.

⁸Instituto Tecnológico de Sonora, Ciudad Obregón, México.

⁹University of Sonora, Hermosillo, México.

Compared with women living in Mexico, Hispanic/Latina women in the United States (US) have a higher mortality rate (15 per 100,000), which is lower than that of non-Hispanic whites (NHWs) (23 per 100,000).⁶ It has also been reported that non-US-born post-menopausal Hispanic women have a lower risk of breast cancer than US-born Hispanic women.⁷

The variation in breast cancer burden among women living in Mexico, Hispanic women in the US, and NHW women in the US might be explained, in part, by differences in each population's breast cancer risk profile¹; genetic admixture has also been shown to alter breast cancer risk both in Mexican women⁸ and US Hispanic/Latinas.^{9–11} Reproductive and hormonal factors, including early age at menarche, late age at first birth, nulliparity, and lack of breastfeeding, among others, are known predictors of breast cancer risk;¹² however, growing evidence suggests that specific risk factors correlate with distinct breast cancer tumor subtypes.¹³ Understanding differences in patterns of reproductive and hormonal characteristics among different subgroups of Hispanic women (e.g., by culture) might help explain the differences in corresponding breast cancer incidence rates or the tumor subtype distribution.

Acculturation is generally defined as the cultural exchange that occurs when two distinct cultures come in contact for an extended period of time, as occurs in immigration.¹⁴ During extended exposures among different cultures, attitudes, values, behaviors, and the language of each culture are influenced by one another.¹⁵ Acculturation has been used as an explanatory factor for both adverse and beneficial health effects in immigrant populations, including Hispanic/Latinos.^{16–19} Many acculturation measures use language as a proxy.^{20,21} This approach has been criticized, suggesting that it does not accurately explain the dynamic process of acculturation or identify the contextual factors associated with the outcome of interest.^{14,16,22} For example, it could be argued that this proxy measure is primarily an indicator of socioeconomic status (SES). Lacking in the literature, however, are studies considering whether or not measures of SES account for observed associations of acculturation with health outcomes of interest. Furthermore, to our knowledge, a comparison that includes a population residing in the country of origin of the immigrant population has not been reported in the literature.

In this report, we compare the distribution of breast cancer reproductive and hormonal risk factors by level of acculturation and country of residence in women of Mexican descent, taking into account level of education. Education is considered a central component of SES given its relevance to occupational opportunities and earning potential.²³ Compared with income inquiries, the response rate to education questions is high and education is applicable to all adults regardless of age or employment status.²⁴ This is particularly important in women who may be outside of the labor force due to caregiving responsibilities. An important and unique aspect of this study is the inclusion of a population of women residing in Mexico, the country of origin and cultural nexus for US participants.

Methods

Participants

Details regarding the study design and recruitment for the *Ella* Binational Breast Cancer Study have been previously

published.²⁵ Briefly, the *Ella* Study is a case series of women of self-reported Mexican descent who were age ≥ 18 years and diagnosed with invasive breast cancer in the previous 24 months prior to enrollment. Participants were recruited from two study sites in the US (University of Arizona in Tucson, Arizona and MD Anderson Cancer Center in Houston, Texas) and three sites in Mexico (Universidad de Sonora in Hermosillo, Sonora; Instituto Tecnológico de Sonora in Ciudad Obregon, Sonora; and Universidad de Guadalajara in Guadalajara, Jalisco). Recruitment took place from March 2007 through June 2011. All recruitment sites used a predominantly clinic-based recruitment strategy; response rates for all aspects of the study (i.e., risk factor questionnaire, medical record access, and tissue collection) ranged from 95%–99%.²⁵ The dataset for the present analyses includes women who provided risk factor questionnaire and medical record data and who have known data on acculturation, education, and age at diagnosis ($n=1201$). The Institutional Review Board from each institution approved the study, and all participants provided written informed consent.

Data collection

Risk factor data were collected via an in-person interview at the recruitment site or participant's home, depending on her preference. At the time of the interview, participants also provided consent to abstract their medical records to obtain clinical data. The risk factor questionnaire included information on sociodemographic data, reproductive factors, and other breast cancer risk factors; it also included questions on acculturation, which are described in the subsequent section. Education was based on the highest level achieved by the participant.

Acculturation measure

Methodology regarding the measure used to assess level of acculturation, including reliability of the measure, has been previously published.²⁶ The cultural orientation scales were language-based and primarily based on Marin and Gamba's (1996) Bidimensional Acculturation Scale,²⁷ which has been previously validated in Hispanic populations.^{20,28,29} The questions included two orthogonal, four-item measures of cultural orientation; one scale assessed degree of English language use/exposure and one scale assessed degree of Spanish language use/exposure. Mexican American (MA) women who completed both scales were placed into one of three acculturation groups using the recommended 2.99 average cutoff for both scales: (1) bilingual (≥ 3.0 average in both the English and Spanish scales); (2) Spanish-dominant (≥ 3.0 average in the Spanish scale only); and (3) English-dominant (≥ 3.0 average in English scale only). The questionnaire was only administered to women residing in the US, since all participating women in Mexico were Spanish speakers. Mexican women were used as a single comparison group to the three groups in the US and were considered to have the lowest level of acculturation.

Statistical methods

Associations between level of acculturation and each breast cancer risk factor were tested using logistic regression, with Mexican women as the reference group, adjusted for age

(continuous) and level of education (<high school, high school, and >high school, treated as an ordinal variable). Risk factor categorizations were dichotomized and conducted *a priori* based on existing literature or on the distribution of the variables in the total population; these include: age at menarche (<12 years),³⁰ age at first full-term pregnancy (>30 years),³⁰ time since last pregnancy (<10 years),³¹ parity (≥ 3 pregnancies), ever breastfeeding, breastfeeding duration (≥ 6 months), ever use of menopausal hormone therapy (HT), and ever use of hormone contraception. All analyses were conducted using Stata 12.1 (Stata-Corp).

Results

Table 1 shows the distribution of breast cancer reproductive and hormonal risk factors for Mexican women and MA women by level of language acculturation. Mexican women had the highest mean age at breast cancer diagnosis, which decreased as language acculturation increased among MA women. Mexican and Spanish-dominant MA women had lower education levels than bilingual or English-dominant

MA women. Across levels of increasing acculturation (from Mexican women to English-dominant MA women), there were trends of decreasing age at menarche and number of full-term pregnancies and decreasing proportions of women with a history of breastfeeding and postmenopausal status.

Level of education was associated with several risk factors (Table 2). Women with less education had older age at menarche, younger mean age at first full-term pregnancy, and a higher mean number of full-term pregnancies than those with higher education. Compared to women with greater than a high school education, women with less than a high school education were much more likely to have breastfed and to breastfeed for longer duration. The percentage of women reporting use of HT increased with education.

After adjustment for age and education, bilingual (odds ratio [OR]=1.63; 95% confidence interval [CI], 1.12–2.38) and English-dominant (OR=2.08; 95% CI, 1.30–3.34) women were significantly more likely to have a younger age at menarche than Mexican women (Table 3). Bilingual (OR=0.50; 95% CI, 0.30–0.83) and English-dominant (OR=0.49; 95% CI, 0.25–0.97) women had significantly lower odds of having a first full-term pregnancy at age ≥ 30

TABLE 1. BREAST CANCER REPRODUCTIVE AND HORMONAL RISK FACTORS BY LEVEL OF ACCULTURATION IN *ELLA* PARTICIPANTS WITH KNOWN AGE AT DIAGNOSIS AND LEVEL OF EDUCATION (N=1201)

Participant characteristics	Mexican American			
	Mexican (n=581)	Spanish dominant (n=202)	Bilingual (n=295)	English dominant (n=123)
Age at diagnosis (years), mean \pm standard deviation (SD)	54.5 \pm 12.5	51.2 \pm 12.0	49.5 \pm 11.7	49.2 \pm 12.2
Highest level of education, n (%)				
Less than high school	389 (67.0)	139 (68.8)	57 (19.3)	17 (13.8)
High school or equivalent	130 (22.4)	51 (25.3)	90 (30.5)	42 (34.2)
Post high school	62 (10.7)	12 (5.94)	148 (50.2)	64 (52.0)
Age at menarche (years), mean \pm SD ^a	13.0 \pm 1.5	12.9 \pm 1.5	12.4 \pm 1.7	12.2 \pm 1.6
Age at menarche <12 years, n (%) ^a	93 (16.0)	32 (15.9)	84 (28.5)	42 (34.2)
Hormone contraceptive use (ever), n (%) ^b	304 (52.3)	99 (49.0)	203 (68.8)	75 (62.0)
Nulliparity, n (%)	55 (9.5)	12 (5.9)	32 (10.8)	20 (16.3)
Age at first full-term pregnancy, mean \pm SD ^c	22.9 \pm 5.5	21.9 \pm 5.0	22.9 \pm 5.5	23.0 \pm 5.8
Age at first full-term pregnancy ≥ 30 years, n (%) ^c	73 (13.9)	15 (7.89)	34 (12.9)	13 (12.6)
Number of full-term pregnancies, mean \pm SD	3.6 \pm 2.6	3.4 \pm 2.0	2.7 \pm 1.8	2.1 \pm 1.5
Number of full-term pregnancies ≥ 3 , n (%)	302 (72.6)	131 (69.0)	160 (60.8)	51 (49.5)
Ever breastfed (if parous), n (%) ^d	466 (89.1)	144 (76.2)	165 (63.2)	53 (51.5)
Lifetime breastfeeding (if parous, months), mean \pm SD ^d	30.5 \pm 39.9	18.9 \pm 23.5	8.4 \pm 12.7	4.7 \pm 10.7
Lifetime breastfeeding ≥ 6 months (if parous), n (%) ^d	391 (74.8)	117 (61.9)	102 (39.1)	23 (22.3)
Time since last pregnancy (if premenopausal, years), mean \pm SD ^e	13.5 \pm 7.3	12.5 \pm 7.7	12.9 \pm 8.3	13.2 \pm 7.9
Time since last pregnancy ≤ 10 years (if premenopausal), n (%) ^e	77 (35.3)	46 (44.2)	60 (40.3)	22 (34.9)
Postmenopausal, n (%) ^f	329 (57.3)	88 (44.9)	123 (41.8)	44 (36.4)
Age at natural menopause (years), mean \pm SD ^g	49.0 \pm 5.2	48.8 \pm 5.1	50.1 \pm 4.8	50.4 \pm 3.4
Age at natural menopause ≥ 50 years, n (%) ^g	136 (51.1)	31 (46.3)	39 (55.7)	13 (68.4)
Menopausal hormone therapy use, n (%) ^h	63 (11.0)	19 (9.45)	57 (19.4)	20 (16.5)

^aMissing data for age at menarche for 1 participant.

^bMissing data for hormone contraceptive use for 2 participants.

^cMissing data for age at first full-term pregnancy for 1 participant, plus 119 women were nulliparous.

^dMissing data for breastfeeding for 6 participants, plus 119 women were nulliparous.

^eMissing data for time since last pregnancy for 67 participants, plus 584 women were postmenopausal, and 16 women had unknown menopausal status.

^fMissing data for menopausal status for 16 participants.

^gMissing data for age at menopause for 162 participants who had nonnatural (or unknown cause of) menopause, plus 601 women were premenopausal, and 16 women had unknown menopausal status.

^hMissing data for menopausal hormone therapy use for 12 participants.

TABLE 2. BREAST CANCER REPRODUCTIVE AND HORMONAL RISK FACTORS BY LEVEL OF EDUCATION IN *ELLA* STUDY PARTICIPANTS WITH KNOWN AGE AT DIAGNOSIS AND LEVEL OF EDUCATION (N=1201)

Participant characteristics	<High school (n=602)	High school (n=313)	>High school (n=286)
Age at diagnosis (years), mean \pm SD	55.5 \pm 12.8	49.9 \pm 10.6	47.6 \pm 11.4
Age at menarche (years), mean \pm SD ^a	13.0 \pm 1.6	12.6 \pm 1.6	12.3 \pm 1.6
Age at menarche < 12 years, n (%) ^a	92 (15.3)	69 (22.0)	90 (31.5)
Hormone contraceptive use (ever), n (%) ^b	298 (49.5)	195 (62.5)	188 (66.0)
Nulliparity, n (%)	44 (7.3)	29 (9.3)	46 (16.1)
Age at first full-term pregnancy, mean \pm SD ^c	21.6 \pm 5.2	22.8 \pm 4.8	25.4 \pm 5.7
Age at first full-term pregnancy \geq 30 years, n (%) ^c	53 (9.52)	26 (9.15)	56 (23.3)
Number of full-term pregnancies, mean \pm SD	3.9 \pm 2.6	2.7 \pm 1.7	2.1 \pm 1.5
Number of full-term pregnancies \geq 3, n (%)	444 (79.6)	169 (59.5)	111 (46.3)
Ever breastfed (if parous), n (%) ^d	464 (83.8)	198 (70.0)	166 (69.5)
Lifetime breastfeeding (if parous, months), mean \pm SD ^d	30.5 \pm 39.6	11.6 \pm 18.2	8.4 \pm 12.2
Lifetime breastfeeding \geq 6 months (if parous), n (%) ^d	392 (70.8)	143 (50.5)	98 (41.0)
Time since last pregnancy (if premenopausal, years), mean \pm SD ^e	14.6 \pm 8.1	12.9 \pm 7.0	11.3 \pm 7.6
Time since last pregnancy \leq 10 years (if premenopausal), n (%) ^e	75 (34.6)	55 (33.7)	75 (48.7)
Postmenopausal, n (%) ^f	356 (59.9)	132 (43.0)	96 (33.8)
Age at natural menopause (years), mean \pm SD ^g	48.8 \pm 5.4	50.2 \pm 4.3	49.5 \pm 4.5
Age at natural menopause \geq 50 years, n (%) ^g	142 (50.2)	50 (56.8)	27 (52.9)
Menopausal hormone therapy use, n (%) ^h	62 (10.4)	42 (13.6)	55 (19.4)

^aMissing data for age at menarche for 1 participant.

^bMissing data for hormone contraceptive use for 2 participants.

^cMissing data for age at first full-term pregnancy for 1 participant, plus 119 women were nulliparous.

^dMissing data for breastfeeding for 6 participants, plus 119 women were nulliparous.

^eMissing data for time since last pregnancy for 67 participants, plus 584 women were postmenopausal, and 16 women had unknown menopausal status.

^fMissing data for menopausal status for 16 participants.

^gMissing data for age at menopause for 162 participants who had non-natural (or unknown cause of) menopause, plus 601 women were premenopausal, and 16 women had unknown menopausal status.

^hMissing data for menopausal hormone therapy use for 12 participants.

years after adjustment for age and education. All three groups of MA women were significantly less likely than Mexican women to report ever breastfeeding (ORs=0.37, 0.21, and 0.13 for Spanish-dominant, bilingual, and English-dominant, respectively); results for breastfeeding \geq 6 months were nearly identical, and these did not change after adjustment for age and education. Although the associations were attenuated after adjustment for age and education, bilingual MA women had significantly higher odds of having used HT (OR=1.84; 95% CI, 1.16–2.93) as well as hormone contraception (OR=1.59; 95% CI, 1.14–2.22) than Mexican women. English-dominant women had higher odds of being nulliparous in the unadjusted model; however, the ORs were attenuated and nonsignificant after adjustment for age and education.

Discussion

In this binational study of women of Mexican descent, we observed differences in the reproductive risk profile according to language acculturation and country of residence. Across levels of increasing acculturation (from Mexican women to English-dominant MA women), we observed increasing trends for lower age at menarche and decreasing trends for proportion of women with a history of breastfeeding as well as breastfeeding duration, after adjustment for age and education. HT and hormone contraception use was highest in bilingual MA women. In a novel finding, the present study shows that English-dominant women had lower odds of being diagnosed within 10 years of their last birth, a transient high-risk period for developing breast cancer;³²

however, due to the small number in this group, the OR lacked precision.

The *Ella* Binational Breast Cancer Study used a language-based acculturation measure, which was administered to participants living in the US. It has been argued that language-only measures of acculturation are partially indicative of SES, and therefore, studies that seek to examine the unique effects of acculturation should take SES into account.³³ Indeed, in the current study, our conceptualization of acculturation and educational attainment were strongly related. Only 10.7% of Mexican women reported education beyond high school, compared with 52% of MA women who were English dominant. To date, few studies have examined the effects of language-based acculturation while considering the influence of education, making it difficult to untangle the effects of these social-contextual factors. The current study assessed acculturation in relation to reproductive and hormonal breast cancer risk factors while controlling for the influence of education, a commonly used indicator of SES. The analyses identified clear associations between the reproductive and hormonal factors of interest and education for most variables; in some instances, associations between acculturation and reproductive factors were attenuated in the multivariate models that adjusted for education, which helps elucidate risk factors that are influenced by culture independent of education. For example, the association between acculturation and age at menarche was only partly attenuated by education. However, although we observed a significantly lower likelihood of having \geq 3 pregnancies for bilingual and English-dominant versus Mexican women in the unadjusted

TABLE 3. ASSOCIATIONS BETWEEN LEVEL OF ACCULTURATION (INDEPENDENT VARIABLE) AND REPRODUCTIVE AND HORMONAL RISK FACTORS (DEPENDENT VARIABLE)

Risk factor	Acculturation level	n (%) ^a	Crude	Odds ratio (95% confidence interval)			Age- and education-adjusted ^b
				Age-adjusted	Education-adjusted	Age- and education-adjusted ^b	
Age at menarche, < 12 years	Mexican	93 (16.0)	1.00	1.00	1.00	1.00	1.00
	Spanish-dominant Bilingual	32 (15.9)	0.99 (0.64–1.54)	0.99 (0.64–1.53)	1.02 (0.65–1.58)	1.02 (0.66–1.59)	1.02 (0.66–1.59)
	English-dominant	84 (28.5)	2.09 (1.49–2.92)	2.07 (1.47–2.91)	1.62 (1.11–2.37)	1.63 (1.12–2.38)	1.63 (1.12–2.38)
Nulliparous	Mexican	42 (34.2)	2.72 (1.76–4.18)	2.69 (1.74–4.17)	2.08 (1.30–3.33)	2.08 (1.30–3.34)	2.08 (1.30–3.34)
	Spanish-dominant Bilingual	55 (9.47)	1.00	1.00	1.00	1.00	1.00
	English-dominant	12 (5.94)	0.60 (0.32–1.15)	0.58 (0.30–1.11)	0.62 (0.33–1.19)	0.61 (0.32–1.17)	0.61 (0.32–1.17)
Age at first full-term pregnancy (if parous), ≥ 30 years	Mexican	32 (10.9)	1.16 (0.73–1.84)	1.10 (0.69–1.75)	0.80 (0.48–1.34)	0.79 (0.47–1.33)	0.79 (0.47–1.33)
	Spanish-dominant Bilingual	20 (16.3)	1.86 (1.07–3.23)	1.74 (1.00–3.05)	1.25 (0.68–2.29)	1.24 (0.67–2.27)	1.24 (0.67–2.27)
	English-dominant	73 (13.9)	1.00	1.00	1.00	1.00	1.00
Parity (if parous), ≥ 3 pregnancies	Mexican	15 (7.89)	0.53 (0.27–0.95)	0.55 (0.31–0.99)	0.55 (0.30–0.99)	0.59 (0.33–1.07)	0.59 (0.33–1.07)
	Spanish-dominant Bilingual	34 (12.9)	0.92 (0.59–1.42)	0.96 (0.62–1.50)	0.50 (0.30–0.82)	0.50 (0.30–0.83)	0.50 (0.30–0.83)
	English-dominant	13 (12.6)	0.89 (0.58–1.68)	0.95 (0.50–1.79)	0.48 (0.24–0.94)	0.49 (0.25–0.97)	0.49 (0.25–0.97)
Time since last pregnancy (among parous, premenopausal women), ≤ 10 years	Mexican	382 (72.6)	1.00	1.00	1.00	1.00	1.00
	Spanish-dominant Bilingual	131 (69.0)	0.84 (0.58–1.20)	0.96 (0.66–1.39)	0.79 (0.54–1.15)	0.89 (0.61–1.30)	0.89 (0.61–1.30)
	English-dominant	160 (60.8)	0.59 (0.43–0.80)	0.68 (0.49–0.94)	1.15 (0.80–1.65)	1.19 (0.82–1.72)	1.19 (0.82–1.72)
Breastfeeding (if parous), ever	Mexican	51 (59.5)	0.40 (0.24–0.57)	0.44 (0.28–0.69)	0.72 (0.45–1.16)	0.77 (0.48–1.24)	0.77 (0.48–1.24)
	Spanish-dominant Bilingual	77 (35.3)	1.00	1.00	1.00	1.00	1.00
	English-dominant	46 (44.2)	1.45 (0.90–2.34)	0.57 (0.83–2.94)	1.56 (0.96–2.53)	1.76 (0.92–3.35)	1.76 (0.92–3.35)
Breastfeeding (if parous), ever	Mexican	60 (40.3)	1.23 (0.80–1.89)	0.95 (0.53–1.70)	0.94 (0.59–1.50)	0.66 (0.35–1.25)	0.66 (0.35–1.25)
	Spanish-dominant Bilingual	22 (34.9)	0.98 (0.55–1.77)	0.67 (0.30–1.47)	0.72 (0.39–1.35)	0.45 (0.19–1.02)	0.45 (0.19–1.02)
	English-dominant	466 (89.1)	1.00	1.00	1.00	1.00	1.00
Breastfeeding (if parous), ever	Mexican	144 (76.2)	0.39 (0.25–0.60)	0.38 (0.24–0.58)	0.39 (0.25–0.60)	0.37 (0.24–0.58)	0.37 (0.24–0.58)
	Spanish-dominant Bilingual	165 (63.2)	0.21 (0.14–0.31)	0.20 (0.14–0.29)	0.21 (0.14–0.22)	0.21 (0.14–0.32)	0.21 (0.14–0.32)
	English-dominant	53 (51.5)	0.13 (0.08–0.21)	0.12 (0.08–0.20)	0.13 (0.08–0.22)	0.13 (0.08–0.21)	0.13 (0.08–0.21)

(continued)

TABLE 3. (CONTINUED)

Risk factor	Acculturation level	n (%) ^a	Odds ratio (95% confidence interval)			
			Crude	Age-adjusted	Education-adjusted	Age- and education-adjusted ^b
Lifetime breastfeeding (if parous), ≥ 6 months	Mexican	391 (74.8)	1.00	1.00	1.00	1.00
	Spanish-dominant	117 (61.9)	0.55 (0.39–0.78)	0.57 (0.40–0.81)	0.54 (0.38–0.77)	0.55 (0.39–0.79)
	Bilingual	102 (39.1)	0.22 (0.16–0.30)	0.23 (0.16–0.31)	0.27 (0.19–0.38)	0.27 (0.19–0.39)
	English-dominant	23 (22.3)	0.10 (0.06–0.16)	0.10 (0.06–0.17)	0.12 (0.07–0.21)	0.12 (0.07–0.21)
						<i>P</i> _{trend} < 0.001
Age at natural menopause (if postmenopausal), ≥ 50 years	Mexican	136 (51.1)	1.00	1.00	1.00	1.00
	Spanish-dominant	31 (46.3)	0.82 (0.48–1.41)	0.81 (0.47–1.40)	0.82 (0.48–1.41)	0.81 (0.47–1.40)
	Bilingual	39 (55.7)	1.20 (0.71–2.04)	1.30 (0.76–2.23)	1.19 (0.66–2.15)	1.15 (0.63–2.10)
	English-dominant	13 (68.4)	2.07 (0.76–5.61)	2.23 (0.81–6.14)	2.05 (0.73–5.71)	2.00 (0.71–5.67)
						<i>P</i> _{trend} = 0.41
Menopausal hormone therapy use, ever	Mexican	63 (11.0)	1.00	1.00	1.00	1.00
	Spanish-dominant	19 (9.45)	0.85 (0.49–1.45)	1.02 (0.58–1.77)	0.86 (0.50–1.48)	1.07 (0.61–1.88)
	Bilingual	57 (19.4)	1.95 (1.32–2.88)	2.76 (1.82–4.18)	1.61 (1.04–2.50)	1.84 (1.16–2.93)
	English-dominant	20 (16.5)	1.60 (0.93–2.77)	2.25 (1.27–4.00)	1.31 (0.72–2.35)	1.45 (0.78–2.70)
						<i>P</i> _{trend} = 0.04
Hormone contraception use, ever	Mexican	304 (52.3)	1.00	1.00	1.00	1.00
	Spanish-dominant	99 (49.0)	0.88 (0.64–1.21)	0.77 (0.55–1.07)	0.89 (0.64–1.23)	0.78 (0.56–1.08)
	Bilingual	203 (68.8)	2.01 (1.50–2.70)	1.72 (1.27–2.34)	1.65 (1.19–2.28)	1.59 (1.14–2.22)
	English-dominant	75 (62.0)	1.48 (0.99–2.22)	1.25 (0.83–1.88)	1.19 (0.77–1.84)	1.14 (0.73–1.77)
						<i>P</i> _{trend} = 0.12

^aNumber (and percent) of women within each acculturation group for each risk factor.^bAge treated as a continuous variable and education as an ordinal variable with categories coded as 0 (< high school), 1 (high school), and 2 (> high school).

model, this was largely explained by education; similar attenuation was observed for nulliparity in English-dominant MA women. Conversely, our findings suggest that the association between breastfeeding and acculturation is not attributable to education, suggesting that this behavior may be largely culturally driven. Thus, our study contributes to the existing literature on acculturation in helping to understand its influence beyond the effects of education as an indicator of SES.

One unexpected finding in our study is the observation that after adjustment for education, MA bilingual and English-dominant women were significantly less likely to have had a first birth after age 30 compared to women in Mexico. However, the results also show that Spanish-dominant women have the lowest proportion of these later births, although the association became non-significant once education was taken into account. These results are not related to the choice of the age cut-point given that when we dichotomized the variable at 25 instead of 30 years, we observed similar results (data not shown). This association needs to be examined in future studies similar to ours.

It is difficult to place our results in context of existing literature, since there is a paucity of data on the association between breast cancer hormonal and reproductive factors and acculturation from studies that include participants residing in the country of origin, and because few studies assessed the spectrum of risk factors examined here. Two published reports^{34,35} showed that US-born Hispanics in California had a higher prevalence of breast cancer risk factors (i.e., hormonal, reproductive factors and obesity) than their foreign-born counterparts. Keegan et al.,³⁴ further noted that this pattern is consistent with the corresponding differences in breast cancer incidence rates in the state. Our results are consistent with published reports on breastfeeding, which show that women born outside the US are more likely to breastfeed than those who are born in the US^{36–41} and that language acculturation also plays a role in whether women choose to breastfeed, particularly among MAs. Results of studies that considered SES variables in their models show that less-acculturated women are more likely to breastfeed than those who are more acculturated.^{39,42–44} Our findings show a substantially lower prevalence and duration of breastfeeding in MA women versus women residing in Mexico, and we also observed a gradient of decreasing breastfeeding with increasing language acculturation in MA women. Our results are also largely in agreement with those of John et al., who assessed migration history, acculturation, and breast cancer risk among Hispanic women (predominantly of Mexican descent).³⁵ Their study found that less-acculturated, Spanish-dominant women had a lower breast cancer risk than their more acculturated, English-speaking counterparts and attributed these differences to adoption of a “Western lifestyle” by the highly acculturated women, which has also been suggested elsewhere in the literature.⁴⁵ Factors contributing to this trend include higher education, no or limited breastfeeding, early age at menarche, low parity, and later age at first pregnancy.³³ In the Multiethnic Cohort study, Pike et al.⁷ showed that the breast cancer risk factor distribution resulted in a predicted multivariate relative risk (RR) of 0.81 for U.S-born Hispanic and 0.72 for non-US-born Hispanic women, compared with NHWs. The models took into account reproductive and hormonal factors as well as body weight and alcohol. In fact, the observed multivariate-

adjusted RR for US-born Hispanic women explained nearly all the difference (RR=0.95); however, for non-US-born Hispanics, the adjusted RR was 0.84, suggesting additional contributing factors. It is important to note that no breastfeeding information was available in this multiethnic study, which could be a key explanatory variable for the lower risk in non-US-born Hispanic women.

Strengths and limitations

Among the strengths of our study is the inclusion of women from the country of origin as a comparison group to assess possible cultural effects; we could not find such a comparison in the literature. Furthermore, by including only women of Mexican descent in our study, we are able to exclude heterogeneity that is present in some studies of Hispanics in the US, which assume the existence of a homogeneous culture across all sub-groups. Our clinic-based study was completed with very high response rates (>90%)²⁵ minimizing sampling bias. As noted previously, adjustment for education is also a strength of our analytical approach. Even though we did not collect other SES variables, such as income, education is a commonly used variable, which is considered a fundamental aspect of SES.^{46,47} From the perspective of understanding associations of SES with health, education provides cognitive resources that may increase receptivity to health promotion messages and foster better access to health services. However, we recognize that education does not fully capture all aspects of SES. For example, income may provide additional information regarding access to tangible resources and health promoting environments, which may be relevant to understanding breast cancer risk and other aspects of health.^{48,49} Further, little is known about the use of education or other markers of SES in low- or middle-resource countries.²⁴ Future research that takes a more comprehensive view to conceptualizing SES in efforts to understand its influence on health among women born outside of the US, or to examine the impact of acculturation across women of Mexican descent, may be informative. Similarly, although language is a common indicator of acculturation, it does not fully capture all aspects of this complex process. An additional limitation of the study includes the cross-sectional design, which does not allow for assessment of temporality between acculturation and risk factors. As well, our study had a modest number of English-dominant MA women ($n = 123$), resulting in less precise estimates of association for this group. Furthermore, the *Ella* Study is a case-only study, which does not permit us to test associations in a non-diseased population or calculate risk of developing breast cancer. Rather, our study provides a framework for an analytical approach for use in binational studies. However, given that we have adequate annotation of the tumor marker profiles^{25,31} we will be able to assess the role of reproductive and hormonal factors in relation to breast tumor subtypes in the context of acculturation in future studies.

Conclusions

Results of the present study show that education, language acculturation, and country of residence are associated with several key risk or protective factors for breast cancer. By considering education in the analysis, our study addresses

some of the criticisms of research concerning language-based acculturation measures, which assert that this metric primarily captures SES. Measuring culture, particularly related to health behavior and outcomes, is complex but clearly relevant to understanding disease prevention and outcome. If our results are replicated in a disease-free population, the continued assessment of sociocultural contextual factors (SES, nativity, acculturation) related to breast cancer risk for women of Mexican descent, both in Mexico and in the US, will be important in understanding the factors underlying breast cancer risk in this population. In addition, the findings support the utility of educational interventions focused on MA women to promote the protective behavior of breastfeeding associated with more traditional Mexican culture.

Given the continued rise in globalization in Mexico and other low- and middle-resource countries, more women will continue to enter the workforce, resulting in more reproductive control by these women, delayed childbearing and lower breastfeeding rates. National survey data for Mexico show that breastfeeding rates are dropping substantially, including in rural areas.⁵⁰ Such changes, along with longer life expectancy and continued low breast cancer screening rates, will undoubtedly further increase the breast cancer burden in Mexico and other less developed countries. As has been suggested previously,⁴⁵ early detection of breast cancer must be a primary goal worldwide. Simultaneously, population-wide educational efforts must be implemented towards meeting this goal along with risk reduction to consider modifiable risk factors, such as promotion of breastfeeding.

Acknowledgments

We are indebted to Erin Ashbeck, Rachel Garcia, and Fang Wang for their contribution. We thank Malaika Tobias for assistance in manuscript preparation. This research was supported by NIH/NCI grants UO1CA153086, CA023074-2953, and CA116199-02S1; the Avon Foundation; and the Susan G. Komen for the Cure® (KG090934).

Disclosure Statement

No competing financial interests exist for any of the authors.

References

- Jemal A, Bray F, Center MM, Ferlay J, Ward E, Forman D. Global cancer statistics. *CA Cancer J Clin* 2011;61:69–90.
- Chavarri-Guerra Y, Villarreal-Garza C, Liedke PE, et al. Breast cancer in Mexico: A growing challenge to health and the health system. *Lancet Oncol*. 2012;13:e335–343.
- Porter PL. Global trends in breast cancer incidence and mortality. *Salud Publica Mex* 2009;51:S141–146.
- Bosetti C, Malvezzi M, Chatenoud L, Negri E, Levi F, La Vecchia C. Trends in cancer mortality in the Americas, 1970–2000. *Ann Oncol* 2005;16:489–511.
- de la Vara-Salazar E, Suarez-Lopez L, Angeles-Llerenas A, Torres-Mejia G, Lazcano-Ponce E. [Breast cancer mortality trends in Mexico, 1980–2009]. *Salud Publica Mex* 2011; 53:385–393.
- Howe LD, Galobardes B, Matijasevich A, et al. Measuring socio-economic position for epidemiological studies in low- and middle-income countries: A methods of measurement in epidemiology paper. *Int J Epidemiol* 2012;41:871–886.
- Pike MC, Kolonel LN, Henderson BE, et al. Breast cancer in a multiethnic cohort in Hawaii and Los Angeles: Risk factor-adjusted incidence in Japanese equals and in Hawaiians exceeds that in whites. *Cancer Epidemiol Biomarkers Prev* 2002;11:795–800.
- Fejerman L, Romieu I, John EM, et al. European ancestry is positively associated with breast cancer risk in Mexican women. *Cancer Epidemiol Biomarkers Prev* 2010;19:1074–1082.
- Slattery ML, John EM, Torres-Mejia G, et al. Genetic variation in genes involved in hormones, inflammation and energetic factors and breast cancer risk in an admixed population. *Carcinogenesis* 2012;33:1512–1521.
- Fejerman L, Chen GK, Eng C, et al. Admixture mapping identifies a locus on 6q25 associated with breast cancer risk in US Latinas. *Hum Mol Genet* 2012;21:1907–1917.
- Fejerman L, John EM, Huntsman S, et al. Genetic ancestry and risk of breast cancer among US Latinas. *Cancer Res* 2008;68:9723–9728.
- Key TJ, Verkasalo PK, Banks E. Epidemiology of breast cancer. *Lancet Oncol* 2001;2:133–140.
- Martinez ME, Cruz GI, Brewster AM, Bondy ML, Thompson PA. What can we learn about disease etiology from case-case analyses? Lessons from breast cancer. *Cancer Epidemiol Biomarkers Prev* 2010;19: 2710–2714.
- Cabassa LJ. Measuring acculturation: Where we are and where we need to go. *Hisp J Behav Sci* 2003;25:127–146.
- Carvajal SC, Granillo TM. Acculturation theory, practice and new developments. In: Leong F, ed. *Encyclopedia of counseling: Cross-cultural counseling and psychotherapy*. Thousand Oaks, CA: Sage Publications Inc., 2008.
- Alegria M. The challenge of acculturation measures: What are we missing? A commentary on Thomson & Hoffman-Goetz. *Soc Sci Med* 2009;69:996–998.
- Matsudaira T. Measures of psychological acculturation: A review. *Transcult Psychiatry* 2006;43:462–487.
- Abraido-Lanza AF, Armbrister AN, Florez KR, Aguirre AN. Toward a theory-driven model of acculturation in public health research. *Am J Public Health* 2006;96:1342–1346.
- Arcia E, Skinner M, Bailey D, Correa V. Models of acculturation and health behaviors among Latino immigrants to the US. *Soc Sci Med* 2001;53:41–53.
- Thomson MD, Hoffman-Goetz L. Defining and measuring acculturation: A systematic review of public health studies with Hispanic populations in the United States. *Soc Sci Med* 2009;69:983–991.
- Hunt LM, Schneider S, Comer B. Should “acculturation” be a variable in health research? A critical review of research on US Hispanics. *Soc Sci Med* 2004;59:973–986.
- Carter-Pokras O, Bethune L. Defining and measuring acculturation: a systematic review of public health studies with Hispanic populations in the United States. A commentary on Thomson and Hoffman-Goetz. *Soc Sci Med* 2009;69:992–995; 999–1001.
- Adler NE, Newman K. Socioeconomic disparities in health: Pathways and policies. *Health Affairs* 2002;21:60–76.
- Howe LD, Galobardes B, Matijasevich A, et al. Measuring socio-economic position for epidemiological studies in low- and middle-income countries: a methods of measurement in epidemiology paper. *Int J Epidemiol* 2012; 41:871–886.
- Martinez ME, et al. Comparative study of breast cancer in Mexican and Mexican-American women. *Health* 2010; 2:1040–1048. Available at: <http://www.scirp.org/journal/>

- PaperInformation.aspx?paperID=2632 Accessed March 18, 2013.
26. Garcia RZ, Carvajal SC, Wilkinson AV, et al. Factors that influence mammography use and breast cancer detection among Mexican-American and African-American women. *Cancer Causes Control* 2012;23:165–173.
 27. Marin G, Gamba RJ. A new measurement of acculturation for Hispanics: The bidimensional acculturation scale for Hispanics (BAS). *Hisp J Behav Sci* 1996;18:297–316.
 28. Hamilton AS, Hofer TP, Hawley ST, et al. Latinas and breast cancer outcomes: Population-based sampling, ethnic identity, and acculturation assessment. *Cancer Epidemiol Biomarkers Prev* 2009;18:2022–2029.
 29. Mainous AG, Diaz VA, and Geesey ME. Acculturation and healthy lifestyle among Latinos with diabetes. *Ann Fam Med* 2008;6:131–137.
 30. Gail MH, Brinton LA, Byar DP, et al. Projecting individualized probabilities of developing breast cancer for white females who are being examined annually. *J Natl Cancer Inst* 1989;81:1879–1886.
 31. Cruz GI, Martinez ME, Natarajan L, et al. Hypothesized role of pregnancy hormones on HER2+ breast tumor development. *Breast Cancer Res Treat* 2013;137:237–246.
 32. Schedin P. Pregnancy-associated breast cancer and metastasis. *Nat Rev Cancer* 2006;6:281–291.
 33. Gallo LC, de los Monteros KE, Allison M, et al. Do socioeconomic gradients in subclinical atherosclerosis vary according to acculturation level? Analyses of Mexican-Americans in the multi-ethnic study of atherosclerosis. *Psychosom Med* 2009;71:756–762.
 34. Keegan TH, John EM, Fish KM, Alfaro-Velcamp T, Clarke CA, Gomez SL. Breast cancer incidence patterns among California Hispanic women: Differences by nativity and residence in an enclave. *Cancer Epidemiol Biomarkers Prev* 2010;19:1208–1218.
 35. John EM, Phipps AI, Davis A, Koo J. Migration history, acculturation, and breast cancer risk in Hispanic women. *Cancer Epidemiol Biomarkers Prev* 2005;14:2905–2913.
 36. Noble L, Hand I, Haynes D, McVeigh T, Kim M, Yoon JJ. Factors influencing initiation of breast-feeding among urban women. *Am J Perinatol* 2003;20:477–483.
 37. Bonuck KA, Freeman K, and Trombley M. Country of origin and race/ethnicity: impact on breastfeeding intentions. *J Hum Lact* 2005;21:320–326.
 38. Celi AC, Rich-Edwards JW, Richardson MK, Kleinman KP, Gillman MW. Immigration, race/ethnicity, and social and economic factors as predictors of breastfeeding initiation. *Arch Pediatr Adolesc Med* 2005;159:255–260.
 39. Gibson-Davis CM, Brooks-Gunn J. Couples' immigration status and ethnicity as determinants of breastfeeding. *Am J Public Health* 2006;96:641–646.
 40. Merewood A, Brooks D, Bauchner H, MacAuley L, Mehta SD. Maternal birthplace and breastfeeding initiation among term and preterm infants: A statewide assessment for Massachusetts. *Pediatrics* 2006;118:e1048–1054.
 41. Singh GK, Kogan MD, Dee DL. Nativity/immigrant status, race/ethnicity, and socioeconomic determinants of breastfeeding initiation and duration in the United States, 2003. *Pediatrics* 2007;119:S38–46.
 42. Ahluwalia IB, D'Angelo D, Morrow B, McDonald JA. Association between acculturation and breastfeeding among Hispanic women: Data from the Pregnancy Risk Assessment and Monitoring System. *J Hum Lact* 2012;28:167–173.
 43. Gibson MV, Diaz VA, Mainous AG, 3rd, Geesey ME. Prevalence of breastfeeding and acculturation in Hispanics: Results from NHANES 1999–2000 study. *Birth* 2005;32:93–98.
 44. Sussner KM, Lindsay AC, Peterson KE. The influence of acculturation on breast-feeding initiation and duration in low-income women in the US. *J Biosoc Sci* 2008;40:673–696.
 45. Porter P. "Westernizing" women's risks? Breast cancer in lower-income countries. *N Engl J Med* 2008;358:213–216.
 46. Hadden WC. Annotation: The use of educational attainment as an indicator of socioeconomic position. *Am J Public Health* 1996;86:1525–1526.
 47. Liberatos P, Link BG, Kelsey JL. The measurement of social class in epidemiology. *Epidemiol Rev* 1988;10:87–121.
 48. Krieger N, Williams, DR, Moss NE. Measuring social class in US public health research: Concepts, methodologies, and guidelines. *Annu Rev Public Health* 1997;18:341–378.
 49. Lynch J, Kaplan G. Socioeconomic position. In: Berkman L, Kawachi I, eds. *Social epidemiology*. New York: Oxford University Press, 2000:13–35.
 50. Gutierrez JP, Rivera J, Shamah T, Oropeza C, Ávila MH. Encuesta Nacional de Salud y Nutrición 2012: Resultados Nacionales. 2012 Institution Cuernavaca, Mexico April 16 2013. Available at: <http://ensanut.insp.mx/informes/ENSANUT2012ResultadosNacionales.pdf>

Address correspondence to:

Jesse Nodora, DrPH
 Moores UCSD Cancer Center
 3855 Health Sciences Drive, #0901
 La Jolla, CA 92093

E-mail: jnodora@ucsd.edu