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## Age at menopause and determinants of hysterectomy and menopause in a multi-ethnic community: The Hilo Women's Health Study

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### Abstract

**Objectives**—A lifespan approach was used to evaluate age at menopause, and determinants of surgical and natural menopause, in the multi-ethnic community of Hilo, Hawaii.

**Study design**—Participants aged 40–60 years (n=898) were drawn from a larger, randomly-generated sample recruited by postal questionnaires. Median age at natural menopause was computed by probit analysis. Logistic regression analysis was applied to examine determinants of hysterectomy, and Cox regression analysis was used to examine risk factors for an earlier age at menopause.

**Main outcome measures**—History of hysterectomy, Age at menopause

**Results**—Frequency of hysterectomy was 19.2% at a mean age of 40.5 years. The likelihood of hysterectomy increased with older ages, lower education, mixed ancestry, having been overweight at age 30, and married 20 years prior to survey. Median age at natural menopause was 53.0 years. Smoking and not being married 10 years before survey were associated with an earlier age at menopause.

**Conclusions**—Median age at menopause was later than the national average. Ethnicity and education were determinants of hysterectomy, but not associated with age at natural menopause. Events later in the lifespan (e.g., smoking and not being married 10 years prior to the survey) were more important than earlier events (e.g., childhood residence) in relation to age at menopause. The timing of weight gain and marital status appear to be important in relation to surgical menopause,

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#### Contributions:

Lynnette Leidy Sievert, authored the publication and carried out all analyses in close collaboration with Lorna Murphy who supervised logistic regression and Cox regression analyses; Lynn Morrison, supervised data collection and substantially edited the paper; Angela Reza, project manager and edited the paper; Daniel E. Brown, PI of the grant and substantially edited the paper.

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and the timing of marital status appears to be important in relation to the timing of natural menopause.

## Keywords

Hawaii; age at menopause; hysterectomy; BMI; ethnicity; smoking

## 1. Introduction

Ages at surgical or natural menopause are associated with long term consequences for health during the post-reproductive years. For example, earlier ages at hysterectomy with bilateral oophorectomy are associated with an elevated risk of cardiovascular mortality,<sup>1-2</sup> and a later age at natural menopause is associated with a higher risk of breast cancer.<sup>3-4</sup> A later age at natural menopause is also associated with increased longevity<sup>5-6</sup> and lower all-cause mortality.<sup>7</sup>

The state of Hawaii boasts the longest life expectancy in the nation, with a mean life expectancy of 80.5 years<sup>8</sup> vs. a national average of 78.7 years;<sup>9</sup> thus, median age at menopause in Hawaii may also be later, relative to the median ages of 51 to 52.5 years in the United States.<sup>10-12</sup> The purpose of this study was to establish the median age at natural menopause in Hilo, Hawaii, as well as the determinants of surgical and natural menopause among this multi-ethnic population. Confirming determinants in this population contributes to our knowledge about risk for health concerns in Hawaii so that women can be targeted for screening and other preventative measures.

Variation in age at menopause, both within and across populations, is related to differences in genes, early environment, adult lifestyle, and reproductive history. Estimates for heritability in age at menopause vary from 31 to 85%.<sup>13-16</sup> A number of studies have demonstrated a relationship between early ages at menopause and adverse early life events, such as severe caloric restriction during early childhood,<sup>17</sup> low weight gain during early childhood,<sup>18-19</sup> socioeconomic disadvantage,<sup>20</sup> or infectious disease burden.<sup>21</sup> Early ages at menopause are also associated with smoking,<sup>10,22</sup> nulliparity,<sup>10,23</sup> low levels of education,<sup>11</sup> being single or otherwise unmarried,<sup>24</sup> and low BMI at midlife.<sup>22,23</sup>

Ethnicity has been associated with age at menopause in some studies, but not others. In the longitudinal Study of Women's Health Across the Nation (SWAN), Japanese ancestry was at first significantly associated with a later age at natural menopause compared to non-Hispanic whites (51.8 years vs. 51.4 years).<sup>10</sup> The most recent study by SWAN researchers, using data from 1996 through 2007 (n=3253), found that Japanese participants had a later age at menopause than African-American, Chinese, and Hispanic participants in unadjusted results. However, after adjusting for smoking, self-reported health, education level, use of oral contraceptives, alcohol intake, employment, physical activity, and baseline weight, ethnic differences were no longer significant.<sup>11</sup> An independent effect of race/ethnicity on the timing of natural menopause was shown in the Multiethnic Cohort Study of women aged 45-74 (n=95,704). After adjusting for smoking status, age at menarche, parity, and BMI, native Hawaiians had an earlier menopause than non-Hispanic whites (HR 1.05, 95% CI 1.01-1.09). Japanese-Americans experienced a later menopause than non-Hispanic whites (HR = 0.93, 95% CI 0.90-0.95).<sup>23</sup>

Ethnicity has also been associated with variation in rates of hysterectomy. In a 1977 study of menopause among women aged 35 to 60 in Hawaii, menopausal women of European descent were more likely to have undergone "surgery for female disorders" compared to Japanese-Americans (52% vs. 37%).<sup>25</sup> More recently, SWAN results found that Asian-Americans reported a lower rate of hysterectomy compared with non-Hispanic whites.<sup>26</sup> In a

study of California school teachers, Asians/Pacific Islanders were not more likely to have surgery for uterine fibroids than non-Hispanic whites.<sup>27</sup>

Hilo, on the Big Island of Hawaii, was once a port associated with a sugar cane industry that attracted thousands of workers of different ethnicities to Hawaii.<sup>28</sup> Hilo is now a multi-ethnic community of 46,165 residents. Japanese-Americans are the largest ethnic group (21%), followed by European-Americans (19%), Native Hawaiians (11%), and Filipinos (7%). Thirty-two percent of people classify themselves as members of two or more ethnic groups (2010 census). This population provides the opportunity to study different ethnic groups residing in the same general locality, although there are socioeconomic and cultural differences, e.g., in diet and activity patterns.<sup>29</sup> Hilo is also of interest because of a previous study that showed a high frequency of hysterectomies. In a study of ambulatory blood pressure among Japanese-American and European-American school teachers (n=120), 66% of the 47 postmenopausal participants (29 Japanese-Americans and 18 European-Americans) had undergone a hysterectomy, 15 with ovaries left intact and 16 with ovaries removed.<sup>30</sup>

The purpose of this study was to consider age at menopause in the multi-ethnic community of Hilo, HI, and to examine determinants of surgical and natural menopause. This study applied a lifespan approach<sup>31,32</sup> to examine early life events (e.g., where women grew up – Hawaii, U.S. mainland, not U.S.), level of education, and later life events such as smoking habits and marital status. In a lifespan perspective, biological and socio-cultural trajectories are intertwined across time,<sup>33–35</sup> and every point in the lifespan can be studied as both predictive of later aging and the result of cumulative processes.<sup>36</sup>

Age at hysterectomy and age at natural menopause are both influenced by biological and socio-cultural variables. For example, a twin study carried out in the UK found similar estimates of heritability for hysterectomy (0.59, 95% CI 0.43–0.72) and for the main indications for hysterectomy, fibroids (0.69) and menorrhagia (0.55).<sup>15</sup> In a more recent study, the decision to undergo a hysterectomy was associated with fibroids and uterine bleeding, and also with the importance of sexual relationships and attitudes, e.g., whether the uterus is “useless” to women who have completed childbearing.<sup>37</sup> A lower level of education and income has also been associated with increased rates of hysterectomy.<sup>38</sup>

Age at natural menopause is determined by the number of undeveloped eggs present in ovaries at birth and the rate of loss of ovarian follicles across the lifespan.<sup>39,40</sup> We do not have birth weight to assess possible effects of maternal health on the number of undeveloped eggs present at birth; therefore, we focus on determinants that may influence the rate of loss of ovarian follicles across the lifespan. In addition to identifying determinants of age at surgical and natural menopause, we are interested in when, during the lifespan, such variables may have had an influence.

## 2. Methods

### 2.1 Study sample

For this study, women aged 40 to 60 (n=898) were drawn from the larger Hilo Women’s Health Study. This study involved two phases: a postal survey about general health and menopause, and a follow up clinical study of hot flashes and blood pressure among a subsample of 200 women. The current analysis is based on the postal survey.

Eight-page health questionnaires were mailed to property lots in Hilo based on tax map key numbers chosen by random assignment (see Sievert et al. for further details).<sup>41</sup> Surveys were mailed to 7,207 households, and 1824 surveys were completed and returned for a

household return rate of 28.5%. All women gave written informed consent and the study was approved by the University of Hawaii Human Studies Program.

To determine ethnicity, women were asked to check off all groups “that are included in your background” from a list of ethnicities, and were asked to include the percentage of each ethnicity where this was known. For these analyses, ethnicity was categorized into 4 groups: European-American, Japanese-American, Mixed, and Other. The category European-American was made up of women who claimed 100% European ancestry on the questionnaire. The category Japanese-American was limited to women who claimed 100% Japanese ancestry. Although Hawaiian Japanese-Americans are primarily third and fourth generation, and native English-speakers, the Japanese-American community in Hawaii has intermarried less than other ethnic groups.<sup>42</sup> Therefore, a high proportion of women with 100% Japanese ancestry is present.<sup>41</sup> In the mixed ancestry group, the majority indicated Hawaiian (60%) and/or European (66%) ancestry. Mixed ancestry also included Chinese (43%), Japanese (25%), Filipino (23%), Hispanic (12%), Native American (12%), Korean (4%), Pacific Islander (4%), and African-American (2%) backgrounds. Women were categorized as “Other” if they described their background as 100% one ethnicity besides European or Japanese.

## 2.2 Variables of interest

Body mass index (BMI) at time of survey was calculated from self-reported height and weight. Past BMI was calculated from recalled, self-reported weight at ages 20, 30, and 40 in combination with self-reported height at time of survey. For multivariate models, BMI was categorized as underweight/normal (<25 kg/m<sup>2</sup>), overweight (25 to 30 kg/m<sup>2</sup>), and obese (>30 kg/m<sup>2</sup>).

Variables examined in relation to ages at surgical and natural menopause included: age at time of survey; ethnicity; where a woman grew up (Hawaii, U.S. mainland, not U.S.); level of education (some or completed high school, some or completed two year or four year college, some or completed postgraduate degree); financial comfort (struggling, OK, comfortable, well-off); marital status at time of survey, marital status 10 years ago, marital status 20 years ago; parity (parous or nulliparous); smoking habits at time of survey (yes/no); BMI at time of survey and at ages 20, 30, and 40; age at menarche; use of birth control pills; and use of hormone therapy (HT).

## 2.3 Age at hysterectomy and determinants

Recalled age at hysterectomy was used to determine mean age at hysterectomy. Across all women aged 40 to 60 years, history of hysterectomy (yes/no) was examined through bivariate logistic regression in relation to ethnicity, demographic variables, lifestyle factors, BMI, weight change, and hormone use. Choice of variables was guided by univariate analyses and review of the literature. Women classified as other ethnicity (N=54) were excluded from the multivariable analyses.

Marital status at time of survey, marital status 10 years ago, and marital status 20 years ago were each examined in the model along with other demographic variables. Only marital status 20 years ago was significant in the univariate analyses. After considering models with BMI at ages 30, 40, and at the time of the survey, the best fit was the model that included BMI at age 30. The final model included ethnicity, age at time of survey, age at menarche, BMI at age 30, marital status 20 years ago, parity, level of education, and smoking status.

Where a woman grew up was not included in the model because it was highly associated with ethnicity ( $p < 0.001$ ), e.g., 66% of the European-Americans grew up on the mainland, while 96% of the Japanese-Americans and 87% of the women of mixed descent grew up in

Hawaii. Use of HT was not included in the model because HT use was viewed as a consequence, not determinant, of having had a hysterectomy. Based on the results of the logistic regression, the analysis was repeated among the mixed ethnicity subsample to examine whether percent Hawaiian ancestry was a determinant of having undergone a hysterectomy while controlling for demographic and lifestyle variables. Hosmer and Lemeshow tests for goodness of fit were carried out for all logistic regression models.

## 2.4 Age at natural menopause and determinants

For comparisons within this study, and across other studies, mean recalled ages at natural menopause for the entire sample and for women of European, Japanese, and mixed descent were calculated. In addition, median ages at natural menopause were computed by probit analysis. Post-menopausal women were defined as those who had not menstruated for at least 12 months for the calculation of mean recalled age at menopause, for median ages at menopause, and for Cox regression analyses carried out to determine risk of an earlier age at menopause.<sup>11,43</sup> Women with recalled ages at menopause prior to 40 years of age (n=6) were excluded from this and other analyses.

Mean recalled ages at natural menopause were examined by t-test or ANOVA analyses in relation to where a woman grew up (HI, US mainland, not US); level of education; financial comfort; current marital status (married, not married), marital status 10 years ago, and marital status 20 years ago; parous or nulliparous status; smoking status; BMI (normal, overweight, obese) at time of survey, at age 20, at age 30, and at age 40; change in BMI from age 20 to 40 (<5 kg/m<sup>2</sup>, 5+ kg/m<sup>2</sup>); and use of birth control or HT. Pearson's correlation was used to examine recalled ages at natural menopause in relation to height, age at menarche, age at first pregnancy, and age at last pregnancy.

Cox regression analyses were used to calculate hazard ratios (HR) for the event of not having menstruated for at least 12 months (postmenopausal status). Variables included in a Cox regression analysis precede the event, or are proxies for behaviors that most likely preceded the event (e.g., smoking). Pre-menopausal and naturally post-menopausal women were included in the analysis, with age at most recent menstrual period entered as the covariate of interest. BMI at ages 20, 30, and 40 were examined in separate models. Marital status 10 years ago, 20 years ago, and at time of survey were examined in separate models. The models with current marital status and marital status 20 years ago were not significant; therefore, we included marital status 10 years ago in the final model. Level of education and financial status were also examined separately. Neither were significant variables, although financial status somewhat improved the significance of the model. In addition to age at most recent menstrual period, the final model included ethnicity, financial comfort, smoking status, parity, BMI at age 30, and marital status 10 years ago. All analyses were carried out using SPSS 16.0 GP.

## 3. Results

### 3.1 Sample characteristics

The sample was characterized by women of European-American (24%), Japanese-American (29%), and mixed ethnic descent (41%) who grew up in Hawaii (73%). The majority attended or completed college, described themselves to be financially OK or comfortable, married, with an average of 2 children. Sample characteristics are shown in Table 1.

### 3.2 Age and determinants of hysterectomy

Nineteen percent of women reported a history of hysterectomy. Mean age at hysterectomy was 40.5 years (s.d. 6.8) with a range in age from 22.7 to 58.4 years (n=148). Ninety-four

percent of women were still menstruating at the time of their hysterectomy, 22% had one ovary removed and 43% reported the removal of both ovaries.

In univariate analyses, women with a history of hysterectomy were more likely to have grown up in Hawaii, had lower levels of education, were more likely to have been married 20 years ago, were more likely to have children, more likely to be overweight in the past and at the time of survey, and more likely to be taking HT (Table II). In addition, women with a history of hysterectomy were older at time of survey (51.7 years, s.d. 5.4, vs. 49.4 years, s.d. 5.4,  $p < 0.001$ ) and reported an earlier age at menarche (12.2 years, s.d. 1.6, vs. 12.5 years, s.d. 1.7,  $p = 0.006$ ). See also Figure I.

Results from the final logistic regression analysis are shown in Table III. In this model, women with mixed ancestry were more likely to have undergone a hysterectomy compared to women of European descent (OR 2.10, 95% CI 1.23–3.59). Probability of hysterectomy increased with age (OR 1.09, 95% CI 1.04–1.13), and women who were overweight at age 30 were almost twice as likely to have undergone a hysterectomy (OR 1.87, 95% CI 1.13–3.09). Women who were married 20 years ago were also nearly twice as likely to have undergone a hysterectomy (OR 1.94, 95% CI 1.29–3.16). Lower levels of education increased the likelihood of a hysterectomy, but smoking, parity and age at menarche did not (Table III).

Mixed ethnicity was a determinant of hysterectomy and 60% of women of mixed ethnicity claimed Hawaiian background. Therefore, we examined the determinants of hysterectomy among only women of mixed ethnicity to assess whether self-reported percent Hawaiian ancestry was a determinant of hysterectomy, controlling for variables significant in the larger model. Age and marital status 20 years ago remained significant in this second model; however, Hawaiian ancestry examined as a continuous variable or in quartiles, was not a significant determinant of the likelihood of having undergone a hysterectomy (not shown.)

### 3.3 Age and determinants of natural menopause

Mean recalled age at natural menopause was 49.4 years (s.d. 3.4). Median age at natural menopause by probit analysis was 53.0 years ( $n = 568$ ). As Table IV shows, there were no significant differences across ethnic groups in mean or median ages at menopause.

In univariate analyses, post-menopausal women were older, less likely to be of mixed ethnic descent, and had an earlier age at menarche (Table I). An earlier recalled age at natural menopause was associated with lower levels of financial comfort, smoking, and unmarried status 20 years ago (but not unmarried status 10 years ago or at time of survey) (Table V). Change in BMI from ages 20 to 40 approached significance, so that women who gained 5 or more  $\text{kg/m}^2$  had an earlier age at natural menopause ( $p = 0.051$ ). There was no significant correlation between mean age at natural menopause and height, age at menarche, age at first birth, or age at last birth. The correlation between age at natural menopause and age at birth of last child approached significance ( $r = -.157$ ,  $p = 0.080$ ,  $n = 125$ ).

In the final Cox survival model for age at natural menopause (Table VI), only smoking (HR 1.75, 95% CI 1.12–2.74) and not being married 10 years ago (HR 1.54, 95% CI 1.06–2.25) were significantly associated with the likelihood of an earlier age at menopause. Being overweight at age 30 approached significance for an association with an earlier age at natural menopause (HR 1.50, 95% CI 0.962–2.338).



## 4. Discussion

### 4.1 Age and determinants of hysterectomy

In this study of ages at surgical and natural menopause in the multi-ethnic community of Hilo, Hawaii, frequency of hysterectomy was 19.2%. This frequency of hysterectomy is somewhat higher than the 17.3% found in a survey carried out in 2005, the same year as our survey, in the county of Hawaii, where Hilo is located.<sup>44</sup> It could be that the relatively low rate of questionnaire return (28.5%), and the large percentage of respondents with mixed ancestry (41%), may have biased our sample toward women with hysterectomies. Among women in Hilo, Hawaii, we found that the likelihood of having undergone a hysterectomy increased with older ages, lower levels of education, mixed ancestry, having been overweight at age 30, and having been married 20 years before survey.

The lifespan perspective provides a framework for looking at early life events in relation to later health outcomes. In addition, it offers clues with regard to cohort effects.<sup>33</sup> When the Hilo sample was limited to post-menopausal women (n=390), 42.3% had undergone a hysterectomy. This frequency is lower than the frequency reported in an earlier study among school teachers in Hilo where Brown et al.<sup>30</sup> found that 66% of postmenopausal women had undergone a hysterectomy. The difference in rates of hysterectomy between the two surveys may reflect changes in the delivery of health care over time among the few gynecologists practicing in the Hilo area. Alternatively, the difference may represent an occupational or socioeconomic effect because the earlier study was limited to school teachers.<sup>30</sup>

With a focus on early life events, mixed ancestry, growing up in Hawaii, and lower levels of education were associated with having had a hysterectomy. Compared to European-Americans, women of mixed ancestry were more than twice as likely to have undergone a hysterectomy (OR 2.10, 95% CI 1.23–3.59). There may be a genetic component to risk of hysterectomy;<sup>15</sup> however, a relationship between percent Hawaiian ancestry and likelihood of hysterectomy could not be demonstrated. Instead, in this population, it is more likely that ancestry, place of birth, and education are associated. Women of mixed ethnicity were more likely to have grown up in Hawaii. In addition, women of mixed ancestry have significantly lower levels of education compared to European- and Japanese-American women (p<0.001). Women who grew up in Hawaii have significantly lower levels of education compared to women who grew up outside of Hawaii (p<0.001). For example, only 19% of women who grew up in Hawaii have some post-graduate education compared to 36% of women who grew up on the U.S. mainland.

The effect of mixed ethnicity and lower levels of education persisted in logistic regression results after controlling for age at time of survey and other demographic and lifestyle variables. The relationship between lower levels of education and higher frequencies of hysterectomy has been found in studies in a number of countries.<sup>45,46</sup> In a U.S. study of 7139 women aged 25 to 54 in 15 states, frequency of hysterectomy ranged from 26.1% among women with some high school education to 10.1% among college graduates. After controlling for income, women without a high school degree were twice as likely to have undergone a hysterectomy compared to women with a college degree or beyond (OR 2.1, 95% CI 1.7–2.7).<sup>38</sup> Factors that may influence hysterectomy rates in relation to levels of education include access to information, access to health care resources, ability to communicate with physicians, and attitudes toward surgical procedures.<sup>25,45</sup>

With regard to later life events, marital status 20 years ago, but not marital status at time of survey or 10 years prior to survey, was associated with history of hysterectomy. Women who were married 20 years ago were twice as likely to have undergone a hysterectomy (OR 1.94, 95% CI 1.20–3.16). The women in this study have a mean age of 50.0 years. Mean age

at hysterectomy was 40.5 years. It is interesting, then, that it was not marital status at the time of hysterectomy, but marital status on average 10 years prior to hysterectomy – around the age of 30 – that was associated with a higher likelihood of hysterectomy. As Figure I shows, fewer women were married 20 years prior to the survey. Marriage may have increased the likelihood of health insurance. In addition, if heavy bleeding or pain from fibroids affected sexual relationships,<sup>37</sup> then married women may have been more likely to seek a surgical treatment. The answer is not that women married 20 years prior to the survey were more likely to have had children, because women married 10 years ago and at the time of survey were also significantly more likely to have had children compared to unmarried women at those periods of time. Parity did not remain a significant determinant of hysterectomy after controlling for other demographic and lifestyle factors.

Finally, with regard to hysterectomy, women who were overweight at the age of 30 were more likely to have undergone a hysterectomy. Univariate and logistic regression results point to BMI at age 30, rather than ages 20, 40, or at time of survey, as having relevance for the likelihood of undergoing a hysterectomy. This indicates BMI 10 years prior to the mean age at hysterectomy (Figure I). BMI is related to various health concerns that result in hysterectomy. For example, in a study of 80,204 California teachers, the risk of surgery for fibroids was higher among overweight women (RR 1.23, 95% CI 1.10–1.38) and obese women (RR 1.27, 95% CI 1.11–1.46) compared to women of normal body size, after controlling for numerous demographic and lifestyle factors.<sup>27</sup> There is also a strong association between obesity and endometrial cancer,<sup>47</sup> and an association between obesity and pelvic organ prolapse.<sup>48</sup>

#### 4.2 Age and determinants of natural menopause

Mean recalled ages at menopause are generally earlier than median ages at menopause computed by probit analysis, as is the case here (49.4 years vs. 53.0 years.) This is to be expected because of the truncation of data in a cross-sectional study. Among women aged 40 to 60, those who will have a later age at menopause continue to menstruate and, therefore, cannot contribute their age at menopause to the sample. In a cross-sectional study, probit analysis is the best measure for determining age at menopause because it does not rely on recall. Women are asked their age at time of survey and whether or not they have menstruated within the past 12 months. The age at which 50% of the women have menstruated in the past 12 months, and 50% of the women have not is the median age at menopause.<sup>43</sup>

In Hilo, Hawaii, the median age at menopause is 53.0 years, which is higher than the median reported for the U.S.<sup>11,12</sup> This later age at menopause is consistent with a higher life expectancy in Hawaii, and consistent with the relatively high level of education seen in the sample.

There were no significant differences across ethnic groups in mean or median ages at menopause, consistent with the most recent finding from SWAN.<sup>11</sup> From a lifespan perspective, where a woman grew up was also without significance, although this may be because migrants to Hawaii tend to be of equal or greater socioeconomic status than women living in Hawaii. Quite different results are found when migration takes place, for example, between Bangladesh and London. In a recent study, Bangladeshi immigrants had a significantly earlier age at natural menopause compared to their London neighbors of European ancestry.<sup>9</sup> In Hilo, smoking was significantly associated with the likelihood of an earlier age at natural menopause. The relationship between smoking and age at menopause has been well-substantiated.<sup>49</sup> In Cox survival analyses we found no association between ethnicity, financial comfort, parity, or BMI and age at natural menopause.



From a lifespan perspective, marriage 20 years before survey was related to age at menopause in univariate results, and marriage 10 years ago was related to age at menopause in the Cox regression analysis after controlling for other demographic and lifestyle variables. Marriage at all points in time was significantly related to financial comfort, with married women significantly more likely to describe themselves as comfortable. Married women may have a later age at menopause because of the relationship between higher income and marital status,<sup>12</sup> because of sexual activity, or because of the pheromonal influence of living with a man.<sup>24</sup> One difficulty in pinpointing the relationship between marital status and age at menopause is the fluidity of the former. For example, 21% of the women who were married 20 years ago were not married at the time of the survey, and 56% of the women who were not married 20 years were married at the time of the survey.

### 4.3 Limitations

One limitation of this postal survey is the need to rely on self-reported heights and weights. In general women tend to over-report their height and under-report their weight,<sup>50</sup> however, if everyone tends to under- or over-report in the same direction, then the error will not substantially affect study results. Weight tends to increase with age,<sup>51</sup> and this increase is apparent in Table I and Figure I. As shown in Table I, BMI increases with age (21.4 kg/m<sup>2</sup>, 23.2 kg/m<sup>2</sup>, 25.1 kg/m<sup>2</sup>, and 26.7 kg/m<sup>2</sup>) as would be expected if we had access to measurements rather than self-report.

As noted, the relatively low rate of questionnaire return (28.5%), and the large percentage of respondents with mixed ancestry (41%), may have biased our sample toward women with hysterectomies. With regard to the determinants of hysterectomy, these results may not be generalizable outside of Hilo or Hawaii because of regional variation in rates of hysterectomy.<sup>52</sup>

## 5. Conclusions

The purpose of this study was to consider age at menopause in the multi-ethnic community of Hilo, Hawaii, and to examine determinants of surgical and natural menopause. This study applied a lifespan approach to examine early life events (e.g., where women grew up) and later life events such as the timing of weight gain and the timing of marriage.

Women who were older, married 20 years before the survey, overweight, or of mixed ethnicity were more likely to have undergone a hysterectomy, perhaps because of factors related to a lower level of education such as access to information, access to health care resources, better communication with physicians, or attitudes toward surgical procedures.<sup>25,45</sup> The rate of hysterectomies may also be a reflection of the dominant healthcare paradigm in Hilo one to two decades ago.

With regard to age at menopause, it appears that events later in the lifespan (e.g., smoking and not being married 10 years prior to the survey) were more important than events earlier in the lifespan (e.g., childhood residence) in relation to age at menopause.

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	30 years ago	20 years ago	10 years ago	At survey
Mean age	20.0 yrs	30.0 yrs	40.0 yrs	50.0 yrs
Median age at menopause				50.3 yrs
% naturally postmenopausal <sup>a</sup>		1%	13%	31%
Mean age at hysterectomy			40.5 yrs	
% married	--	67%**	76%*	72%
Mean BMI (kg/m <sup>2</sup> )				
Normal %	90%	77%	60%	46%
Overweight %	8%	16%*	24%	30%
Obese %	2%	8%	15%	24%

**Figure 1.**

The timing of various life events among women in Hilo, Hawaii aged 40 to 60 at survey (n=898).

\* p<0.05, \*\* p<0.01

Significance based on logistic regression results.

<sup>a</sup> Among naturally post-menopausal women at time of survey (n=191), for whom age at menopause is known. Twenty years ago, all but 2 women were pre-menopausal. Ten years ago 82% were pre-menopausal, 5% were peri-menopausal, and 13% were postmenopausal.

**Table I**

Sample characteristics with (1) comparison of pre-menopausal vs. naturally post-menopausal and (2) comparison of naturally post-menopausal vs. hysterectomy status

	All N=898 <sup>a</sup>	Pre-menopausal N=468	Naturally post-menopausal N=225	History of hysterectomy N=165
Age at time of survey (yrs) Mean and (s.d.)	50.0 (5.4)	47.0 (4.3) <sup>** b</sup>	54.5 (3.7)	51.7 (5.4) <sup>** c</sup>
Ethnicity				
European	24%	23% <sup>* b</sup>	29%	16%
Japanese	29%	28%	34%	29%
Mixed (Hawaiian)	41%	43%	30%	49%
Other	6%	6%	6%	6%
Where she grew up				
Hawaii	73%	72%	67%	81% <sup>** c</sup>
Mainland, U.S.	21%	21%	27%	15%
Not U.S.	6%	7%	6%	4%
Level of education				
Some or completed HS	16%	15%	13%	19% <sup>** c</sup>
Some or completed college	60%	60%	56%	66%
Some or completed grad sch	24%	25%	31%	16%
Financial Status				
Struggling	16%	15%	16%	17%
OK	46%	47%	43%	48%
Comfortable	35%	35%	38%	33%
Well-off	3%	3%	4%	2%
Percent married				
At time of survey	72%	75% <sup>* b</sup>	67%	72%
10 years ago	76%	77% <sup>* b</sup>	69%	79% <sup>* c</sup>
20 years ago	67%	57% <sup>** b</sup>	75%	81%
Age at menarche (yrs)				
Mean and (s.d.)	12.5 (1.7)	12.6 (1.7) <sup>* b</sup>	12.3 (1.6)	12.2 (1.6)
Mean parity				
Mean and (s.d.)	2.1 (1.3)	2.1 (1.4)	2.1 (1.4)	2.2 (1.1)
Percent nulliparous	13%	13%	13%	7%
Smoking	16%	17%	14%	15%
Height at time of survey (cm) Mean and (s.d.)	63.0 (2.9)	63.1 (2.8)	62.8 (2.9)	62.8 (3.1)



	All N=898 <sup>a</sup>	Pre-menopausal N=468	Naturally post-menopausal N=225	History of hysterectomy N=165
<b>BMI</b>				
At age 20	21.4 (3.3)	21.4 (3.4)	21.3 (3.1)	21.9 (3.2)
At age 30	23.2 (4.3)	23.2 (4.3)	22.7 (4.3)	24.1 (4.3) <sup>** c</sup>
At age 40	25.1 (5.4)	25.3 (5.6) <sup>** b</sup>	24.1 (4.6)	26.1 (5.7) <sup>** c</sup>
At time of survey	26.7 (6.2)	26.5 (5.9)	26.5 (6.2)	27.9 (7.0) <sup>* c</sup>
<b>Birth control pill</b>				
Never	19%	18% <sup>** b</sup>	20%	20%
Past	77%	75%	79%	80%
Current	4%	7%	1%	1%
<b>Hormone therapy</b>				
Never	71%	92% <sup>** b</sup>	52%	41%
Past	18%	4%	34%	36%
Current	10%	4%	14%	24%

\*  
p<0.05

\*\*  
p<0.01

<sup>a</sup> unable to classify all participants by hysterectomy and menopausal status

<sup>b</sup> Comparison of pre-menopausal vs. naturally post-menopausal status

<sup>c</sup> Comparison of naturally post-menopausal vs. history of hysterectomy

**Table II**

History of hysterectomy in relation to potential determinants among women aged 40 to 60 in Hilo, Hawaii

	N	% having had a hysterectomy	P value*
<b>Ethnicity</b>			
European	201	12.9	0.012
Japanese	250	18.8	
Mixed (Hawaiian)	347	23.3	
<b>Where she grew up</b>			
Hawaii	623	21.3	0.027
Mainland, U.S.	183	13.1	
Not U.S.	51	13.7	
<b>Level of education</b>			
Low	132	23.5	0.012
Medium	518	20.8	
High	211	12.3	
<b>Financial Status</b>			
Struggling	132	20.5	0.681
OK	396	19.7	
Comfortable	297	17.8	
Well-off	26	11.5	
<b>Married 20 years ago</b>			
Married 20 years ago	548	23.4	<0.001
Not married 20 years ago	275	11.3	
<b>Married 10 years ago</b>			
Married 10 years ago	613	19.7	0.276
Not married 10 years ago	197	16.2	
<b>Married at time of survey</b>			
Married at time of survey	621	19.3	0.956
Not married at time of survey	238	19.2	
<b>Parous</b>			
Parous	755	20.3	0.028
Nulliparous	106	11.3	
<b>Not smoking</b>			
Not smoking	719	19.3	0.796
Smoking	136	18.4	
<b>BMI at age 20</b>			
Normal	732	18.9	0.122
Overweight	70	28.6	
Obese	15	13.3	
<b>BMI at age 30</b>			
Normal	608	17.6	
Overweight	129	27.9	

	N	% having had a hysterectomy	P value*
Obese	60	25.0	0.017
BMI at age 40			
Normal	467	16.5	
Overweight	193	23.3	
Obese	121	24.8	0.036
BMI at time of survey			
Normal	372	15.9	
Overweight	246	19.5	0.049
Obese	202	24.3	
BMI change from 20 to 40			
<5 kg/m <sup>2</sup>	566	18.4	
5+ kg/m <sup>2</sup>	220	22.3	0.215
Birth control pill use			
Never	158	20.3	
Past	661	19.8	0.902
Hormone therapy use			
Never	597	11.1	
Past	150	38.7	
Current	90	43.3	<0.001

\* ANOVAs or t-tests

**Table III**

Results of logistic regression for history of hysterectomy (n=712)

	Odds ratio	95% CI	p-value
<b>Ethnicity</b>			
European (ref)			
Japanese	1.360	0.766–2.415	0.294
Mixed (Hawaiian)	2.102	1.232–3.587	0.006
<b>Age</b>			
Age	1.087	1.043–1.132	<0.001
<b>Age at menarche</b>			
Age at menarche	0.901	0.792–1.025	0.113
<b>BMI at age 30</b>			
Normal (ref)			
Overweight	1.868	1.130–3.088	0.015
Obese	1.705	0.860–3.380	0.126
<b>Married 20 years ago</b>			
Married 20 years ago	1.943	1.195–3.159	0.007
Not married 20 years ago (ref)			
<b>Nulliparous (ref)</b>			
Parous	1.377	0.682–2.780	0.372
<b>Education</b>			
High school or less	1.871	0.903–3.877	0.092
Some college or degree	1.932	1.067–3.253	0.013
Some post-grad or degree (ref)			
<b>Current smoker no (ref)</b>			
Current smoker yes	0.847	0.534–1.673	0.945

**Table IV**

Age at natural menopause among women of European, Japanese, and Mixed ancestry (age at menopause >40).  
No significant differences across ethnic groups in mean or median ages at menopause.

Group	N	Mean recalled age at menopause (years, s.d.)	N	Median age at menopause by probit analysis (95% Fieller Bounds)
European	53	49.3 (3.3)	153	53.1 (52.0–54.3)
Japanese	61	50.1 (3.4)	179	53.0 (52.1–54.0)
Mixed	57	48.9 (3.4)	236	52.8 (52.0–53.8)
All	171	49.4 (3.4)	568	53.0 (52.4–53.6)

**Table V**

Recalled age at natural menopause in relation to potential determinants (Women with menopause before age 40 excluded)

	N	Mean age at menopause (s.d)	p-value
<b>Ethnicity</b>			
European	53	49.0 (3.3)	
Japanese	61	50.1 (3.4)	
Mixed (Hawaiian)	57	48.9 (3.4)	
Other	11	48.4 (2.9)	0.167
<b>Where she grew up</b>			
Hawaii	122	49.4 (3.5)	
Mainland, U.S.	50	49.3 (3.3)	
Not U.S.	12	49.6 (2.0)	0.965
<b>Level of education</b>			
Low	22	49.4 (2.4)	
Medium	102	49.1 (3.6)	
High	61	49.8 (3.2)	0.452
<b>Financial Status</b>			
Struggling	27	48.1 (3.8)	
OK	77	49.0 (3.4)	
Comfortable	71	50.1 (3.2)	
Well-off	8	49.9 (1.6)	0.033
<b>Married 20 years ago</b>			
Married 20 years ago	133	49.7 (3.2)	
Not married 20 years ago	45	48.3 (3.7)	0.012
<b>Married 10 years ago</b>			
Married 10 years ago	122	49.6 (3.3)	
Not married 10 years ago	53	49.0 (3.4)	0.227
<b>Married at time of survey</b>			
Married at time of survey	123	49.5 (3.2)	
Not married at survey	61	49.1 (3.6)	0.462
<b>Parous</b>			
Parous	161	49.5 (3.3)	
Nulliparous	24	48.7 (3.8)	0.273
<b>Not smoking</b>			
Not smoking	159	49.6 (3.3)	
Smoking	24	47.8 (3.6)	0.015
<b>BMI at age 20</b>			
Normal	163	49.5 (3.4)	
Overweight	11	48.5 (2.4)	
Obese	2	46.6 (7.4)	0.357
<b>BMI at age 30</b>			



	N	Mean age at menopause (s.d)	p-value
Normal	139	49.4 (3.4)	
Overweight	25	49.5 (2.7)	
Obese	9	48.2 (3.5)	0.566
<b>BMI at age 40</b>			
Normal	121	49.6 (3.1)	
Overweight	35	49.1 (3.2)	
Obese	16	47.6 (3.8)	0.073
<b>BMI at time of survey</b>			
Normal	89	49.6 (3.1)	
Overweight	48	49.3 (3.4)	
Obese	39	49.0 (3.9)	.591
<b>BMI change from 20 to 40</b>			
<5 kg/m <sup>2</sup>	139	49.6 (3.1)	
5+ kg/m <sup>2</sup>	35	48.3 (4.0)	0.051
<b>Birth control pill use</b>			
Never	33	48.7 (4.0)	
Past	148	49.5 (3.2)	0.216
<b>Hormone therapy use</b>			
Never	93	49.4 (3.4)	
Past	63	49.6 (3.0)	
Current	26	48.5 (4.1)	0.377

Table VI

Cox survival model for age at natural menopause (n=545)

	Unadjusted HR	Adjusted HR	95% Confidence interval	P-value	P-trend
Ethnic group					
European (ref)					
Japanese	0.86	0.951	0.646–1.402	0.802	
Mixed (Hawaiian)	0.89	0.874	0.587–1.301	0.506	
Financial comfort					0.407
Struggling	1.112	0.907	0.370–2.222	0.831	
OK	0.660	0.630	0.279–1.425	0.267	
Comfortable	0.691	0.703	0.309–1.601	0.402	
Well off (ref)					
BMI at age 30					0.197
Normal (ref)					
Overweight	1.501	1.500	0.962–2.338	0.073	
Obese	1.185	1.168	0.584–2.333	0.661	
Smoking					
No (ref)					
Yes	1.810	1.750	1.116–2.744	0.015	
Marital status 10 yrs ago					
No	1.719	1.543	1.060–2.245	0.024	
Yes (ref)					
Parous (ref)					
Nulliparous	1.383	1.209	0.761–1.919	0.422	