

# **HHS PUDIIC ACCESS**

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# Associations among personal care product use patterns and exogenous hormone use in the NIEHS Sister Study

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# Abstract

It is hypothesized that certain chemicals in personal care products may alter the risk of adverse health outcomes. The primary aim of this study was to use a data-centered approach to classify complex patterns of exposure to personal care products and to understand how these patterns vary according to use of exogenous hormone exposures, oral contraceptives (OCs) and postmenopausal hormone therapy (HT). The NIEHS Sister Study is a prospective cohort study of 50,884 US women. Limiting the sample to non-Hispanic blacks and whites (N = 47,019), latent class analysis (LCA) was used to identify groups of individuals with similar patterns of personal care product use based on responses to 48 survey questions. Personal care products were categorized into three product types (beauty, hair, and skincare products) and separate latent classes were constructed for each type. Adjusted prevalence differences (PD) were calculated to estimate the association between exogenous hormone use, as measured by ever/never OC or HT use, and patterns of personal care product use. LCA reduced data dimensionality by grouping of individuals with similar patterns of personal care product use into mutually exclusive latent classes (three latent classes for beauty product use, three for hair, and four for skin care. There were strong differences in personal care usage by race, particularly for haircare products. For both blacks and whites, exogenous hormone exposures were associated with higher levels of product use, especially beauty and skincare products. Relative to individual product use questions, latent class variables capture complex patterns of personal care product usage. These patterns differed by race and were associated with ever OC and HT use. Future studies should consider personal care product exposures with other exogenous exposures when modeling health risks.

**CONFLICT OF INTEREST** The authors declare no conflict of interest.

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### Keywords

empirical/statistical models; endocrine disruptors; epidemiology; exposure modeling; personal exposure; population based studies

# INTRODUCTION

The average American woman uses 12 personal care products a day, resulting in daily exposure to an estimated 126 unique chemicals.<sup>1</sup> Because certain chemicals in personal care products are suspected endocrine disrupters (e.g, phthalates, parabens, triclosan),<sup>2,3</sup> there is concern that exposure to personal care products may be associated with risk of breast cancer and/or may have adverse reproductive health effects. *In vitro* and animal studies have demonstrated that endocrine disrupting chemicals can mimic estrogens,<sup>4</sup> alter hormonal signaling, affect developing reproductive systems,<sup>5</sup> and/or disrupt normal mammary development.<sup>6</sup> However, there is currently no definitive evidence for the same effects in humans.<sup>7–10</sup>

The strongest evidence to support concern for endocrine disrupting chemicals comes from experimental animal studies.<sup>10</sup> However, laboratory animals are usually exposed to individual chemicals over short periods of time (often at higher doses than humans), whereas humans are typically exposed to multiple endocrine disrupting compounds simultaneously over many years. Consequently, there is concern that laboratory animal data have not addressed the patterns of exposure to these complex mixtures, which may be most relevant.<sup>11,12</sup> At the same time, more information is needed to characterize the nature of human exposure in order to design more appropriate animal and *in vitro* studies of complex mixtures as well as validate computational models of predicted exposure. To address the limitations in experimental animal studies and capture the exposure characteristics in human populations, human studies are needed.

The objective of the current study was to use data from 47,019 women in the NIEHS Sister Study to characterize patterns of personal care product use across a wide range of products. We hypothesized that individuals would be classifiable according to broad patterns of personal care product usage, with patterns differing by race. To investigate possible correlated estrogenic exposures in epidemiologic studies, we also examined the association between personal care product use and two common estrogenic medications (e.g., oral contraceptive (OC) and hormone therapy (HT) use). We used latent class analysis (LCA) to identify groups of women by patterns of product use and compared these patterns with in terms of past exogenous estrogen use.

# MATERIALS AND METHODS

#### **Study Population**

The Sister Study is a large prospective cohort study directed at identifying environmental and genetic risk factors for breast cancer. The study consists of 50,884 women who had at least one sister diagnosed with breast cancer but were cancer-free themselves at time of enrollment. Study enrollment began in 2003 and ended in 2009, and eligible women were

35–74 years of age. Baseline enrollment activities included a computer-assisted telephone interview and self-administered questionnaires that elicited information about environmental and genetic risk factors for breast cancer. The Sister Study was approved by the institutional review boards at the National Institute of Environmental Health Sciences and Copernicus Group. Written informed consent was provided by study participants. The present analysis was limited to non-Hispanic white (n = 42,558, 84% of participants) and non-Hispanic black (n = 4462, 9% of participants) women (Table 1); there were too few Hispanic (5%) and Other (3%) participants to include in this analysis.

#### **Personal Care Product Assessment**

Detailed self-reported use of 48 personal care products was collected during the baseline phase of the study (Supplementary Material 1) by inquiring about frequency of use (5-level-response options) during the previous 12 months. The five response options varied according to intended use of the product. For example, the response options for a product intended to be used regularly (e.g., hand lotion) included: (1) did not use, (2) used less than once a month, (3) used 1–3 times per month, (4) 1–5 times per week, (5) >5 times per week. Response options for products that are used less often (e.g., hair dye) included: (1) did not use, (2) 1–2 times a year, (3) every 3–4 months, (4) every 5–8 weeks, (5) once a month or more. To identify latent classes of personal care product use, each of three product categories were analyzed separately to identify latent classes for each category: (1) beauty products (e.g., lipstick, mascara, nail polish), (2) hair products (e.g., hair spray, hair relaxers), and (3) skincare products (e.g. facial lotion, hand lotion).

#### Latent Classes

LCA was used to identify groups of individuals with similar patterns of personal care product use. LCA is a data reduction tool that describes variability among multiple, correlated, observed variables in terms of a fewer number of unobserved variables called latent classes. It has been used for identifying patterns of exposure when the exposure is a complex combination of separate factors.<sup>13</sup> Personal care products were categorized into three product types,<sup>14</sup> and separate latent classes were constructed for each type: (1) beauty products, (2) hair products, and (3) skincare products. To reduce dimensionality, improve interpretability of the model, and improve classification and precision we used a method based on Dean et al.<sup>15</sup> to select the variables that were most useful for distinguishing among latent classes (i.e., 10% difference in posterior probabilities between classes). Variables that were determined not to contribute to the distinction between latent classes were removed from the model.

We fit a sequence of LCA models starting with two classes and increasing the number of classes for each model (up to six). To identify an optimal but minimal number of classes, Akaike's information criterion (AIC), the Bayesian information criterion (BIC), and entropy were considered.<sup>13</sup> A smaller AIC and BIC and higher entropy for a particular model suggests a better model fit.<sup>13</sup> The AIC and BIC continue to diminish with increasing numbers of latent classes; however, to avoid overfitting sparsely populated categories, we identified an optimal number of classes through a combination of the change in information criteria (i.e., a leveling off of differences) and minimum size of the latent class (10%). A

summary of the fit statistics and entropy are shown in Table 2. The classes were described and labeled based on item-response probabilities<sup>15</sup> and ability to identify which variables were driving each class.<sup>13</sup>

To assign participants class membership, we used a common classify-analyze approach referred to as the maximum-probability assignment rule, where individuals are assigned to the class in which they have the highest posterior probability of membership.<sup>16</sup> Item-response probabilities provided the basis on which each latent class was interpreted.<sup>17</sup> The correlation among use patterns for the three product types was evaluated by Spearman correlation coefficients of the item-response variables' posterior probabilities.

# Statistical Analysis Describing Association Between Personal Product Use and Exogenous Hormone Use

We examined associations between personal product use and OC use (ever, never) for the entire sample and the association between personal care product use and HT use (never/ever) among women >50 years. We used logistic regression stratified by race to estimate age-adjusted prevalence differences (PD) and 95% confidence intervals in exogenous hormone use associated with latent class membership for the three different product types. OC use and HT use were coded as binary variables so they could be allocated as outcome variables in the logistic models. As an additional analysis we examined the associations between personal product use and duration of OC and HT use (5 vs < 5 years).

# RESULTS

#### Latent Class Descriptions

Based on fit statistics and parsimony (Table 2), three latent classes were identified as optimal for both the beauty and hair product groups; the skincare product group had four classes. Figure 1 indicates the specific personal care products and the item-response probabilities for each product. Analysis of fourteen initial beauty product usage items resulted in three latent classes (infrequent users; moderate users; frequent users) with nine contributing product items: mascara, lipstick, foundation, nail polish, perfume, eye shadow, eyeliner, blush, and make-up remover (Figure 1, Table 3). Analysis of fifteen hair product usage items resulted in three latent classes (infrequent users of hair products other than shampoo/conditioner; users of pomade and hair straightener; frequent users of hair products other than pomade and hair straightener) with six contributing product items: pomade, hair straightener, conditioner, hair spray, hair gel, and shampoo. Finally, analysis of nineteen skincare product usage items resulted in four latent classes (infrequent users; moderate users; frequent users; talcum powder users) with nine contributing product items: cleansing cream, anti-aging cream, body lotion, hand lotion, face cream, foot cream, petroleum jelly, talcum powder applied under arms, and talcum powder applied elsewhere. Short descriptions of the classes were created based on the item-response probabilities.

#### **Product-Use Patterns by Race**

Among non-Hispanic women in our study population, 91% were white and 9% were black (Table 4). Race was most strongly associated with differences in haircare product classes;

only 3% of white women were in the "users of pomade and hair straightener" group, whereas over two-thirds (67%) of black women were. There were also some differences in beauty product classes: white women were more frequent users than black women. The moderate lotion user category was the most common skincare class among both black and white women users (Table 4). Because of racial differences in population distribution across product classes, we report results separately by race in the subsequent analyses.

#### Patterns of Class Membership Across Product Categories (Beauty, Hair, Skin)

After considering different types of personal care products in separate classification schemas, we also evaluated whether product usage class in one category (e.g., hair products) predicted use in another category (e.g., beauty products). As shown in Figure 2, the product types for whites tended to be correlated, i.e., infrequent users of one product type tended to be infrequent users of the other product types, but correlation coefficients were modest. The highest correlation of posterior probabilities of class membership was between infrequent users of beauty and skincare products (r = 0.45). The next strongest correlation was between frequent users of beauty products and hair products (r = 0.39). For black women the correlations among usage patterns for the different product types were low except for skin and beauty (r=0.39).

#### **Exogenous Hormonal Exposures and Product Use**

The prevalence of OC use history was 85% for whites and 86% for blacks; the prevalence of HT among women over age 50 was 51% for whites and 48% for blacks. The percentage of women (all ages) who had ever used OC therapy and/or HT was 93% (N= 39,388) among white women and 91% (4449) among black women. Exogenous hormonal exposures (Figure 3) showed strong associations with product use. White and black women who had ever taken OCs were more likely to be "moderate users" or "frequent users" of beauty products than to be "infrequent users" of beauty products. Black women who had used OCs were also more likely to be "moderate users" of skincare products than to be "infrequent users". White and black women over age 50 who had ever received HT were more likely to be "frequent users" of beauty products than to be "infrequent users" of skincare products. White women who had ever received HT were substantially more likely to be "moderate users" of skincare products. White women who had ever received HT were substantially more likely to be "moderate users" of skincare products or "talcum powder users" than to be "infrequent users". When we used 5 vs < 5 years of OC or HT exposure as the variable of interest, the pattern of results was essentially the same as when we used ever/never, though the associations tended to be somewhat attenuated (Supplementary tables).

# DISCUSSION

We found that use of LCA could identify a relatively small number of subgroups of women with distinct personal care product use and that personal care product use varied by race. We also showed that the women with the highest use of personal care products are more likely to have used the common exogenous hormone medications, OC, or HT. Previous studies have examined correlation structure between specific personal care products,<sup>14,18,19</sup> but such studies were not aimed at reducing the complexity of individual product usage patterns, nor

did these studies evaluate associations between personal care product use and other exposures.

Some key challenges have impeded progress in understanding the relationship between personal care product use and health outcomes in humans. First, publically available data on personal care product usage patterns in the United States typically lack large sample populations or include only specific types of product users.<sup>20–23</sup> Currently the only population-based studies of personal care product use that have a comparison group of infrequent personal care product users are limited to other countries,<sup>18,19</sup> or small populations within the United States.<sup>14</sup> Second, studies that have collected data on product use have described correlations between use patterns for only a limited numbers of products. Third, studies to date have not placed the personal care product exposures in context of other exposures, particularly those in relevant biological pathways.

Our analysis addresses some of these challenges. First, we used a large, nationwide study of personal care product use and other environmental exposures to study patterns of exposure. Second, we took a broad approach to characterizing exposure, starting with data on 48 different personal care products included in the Sister Study's questionnaire. Then we used LCA to identify a subset of 24 of the 48 that provided unique information on pattern of product use within three product-use categories (beauty, hair, and skin). The result was a set of exposure variables that grouped women with similar product use. Finally, we used the LCA-identified use patterns to examine the association between personal care product use and history of using either OC or HT. With this data reduction method, we could investigate associations between product use and other factors with control for potential confounders and without the severe multiple-testing problems that arise when each product is examined separately.

However, with currently available data we cannot specify a commonality of chemical exposures, only a commonality of which products tend to be used together by different groups of women. Information about specific chemicals or ingredients in personal care products was not captured in the questionnaire.

The use of LCA was a key strength of our analysis. As opposed to a variable centered approach that considers how variables are related to each other, LCA is a data-centered approach that considers how variables are grouped within individuals. LCA can reduce and organize large, multifaceted data sets and create manageable categorical data elements to summarize complex patterns.<sup>13</sup> LCA has been used to capture complex exposures in a variety of research settings. For example, it has been used to organize and describe subgroups of weight loss strategies and disordered eating among women<sup>17</sup> and to identify subgroups of emerging sexual behaviors among adolescents.<sup>24</sup> LCA has also been used to identify substance use behavior among adolescents to inform programs that could be targeted for or tailored to the different population subgroups that are expected to show the strongest response.<sup>25</sup>

Our results were consistent with some previous studies. We observed race-related patterns of hair product use similar to those observed in previous smaller studies.<sup>9,26–28</sup> The class that

was characterized by use of pomade and hair straightener contained the majority of black women, but only 3% of white women. In a small study with 10 black and 206 white participants from California<sup>14</sup> African American women were more likely to have their hair treated permanently (including chemical straightening or relaxing). Although the National Health and Nutrition Examination Survey (NHANES) 1999–2000 did not collect questionnaire information on frequency of product use, a NHANES analysis of 2540 participant samples found that compared with non-Hispanic white women, African American women had higher urinary levels of monoethyl phthalate, a phthalate found in personal care products.<sup>9</sup> The authors of the NHANES suggest that these differences were likely due to differences in hair texture and cultural practices. Although our analyses did not link product exposure with specific internal dose markers, future application of LCA approaches, or the patterns identified here, could be used to link specific exposure biomarkers with self-reported exposure information.

To consider personal care product use in association with health outcomes, it is important to integrate these exposures with other biologically relevant and risk-related exposures. Having observed correlation within personal care product use classes (e.g., between heavy users of haircare products and heavy users of beauty products), we tested whether personal care product categories are associated with other relevant exposure patterns. OC and HT are examples of key exposures to exogenous estrogens. There were statistically significant associations between beauty and skincare product latent classes and OC and HT. Therefore, when personal care product use is being evaluated as a potential risk factor for hormonally mediated conditions, we encourage researchers to consider possible confounding by OC and HT use.

Our analysis has several key strengths. LCA provides an objective method of distinguishing between groups of women on the basis of their patterns of personal care product use and, thus, potential chemical exposures. The number of latent classes is determined, in part, by data-based metrics, and is small relative to the number of product use items. The component product-use probabilities are complex and objectively discerning overall product-use data from casual inspection would be difficult. The NIEHS Sister Study provided a large dataset for this purpose.

LCA provides an objective means of reducing data dimensionality; however, there are some limitations. For example, the classes can be difficult to interpret. Labels were assigned to different classes based on our observation and interpretation of the probability based weights for class membership. Although there is some subjectivity in choosing the shorthand label descriptors for different classes, the precise item-response probabilities are provided in Supplementary tables. Another limitation was that we could not consider them all of the personal care products together; owing to the large number of variables we had to break LCA into three different product category models: beauty, hair, and skin products. Also, the categories we identified with LCA may not be generalizable to other populations. Therefore it is important that these methods are replicated in other populations. Finally, our models only adjusted for age and not for other possible confounders of the personal care product use and exogenous estrogen association. However, the goal of this research was not to quantify the independent effects of personal care product use on estrogen exposure, but rather to

illustrate a method for identifying important covariates for future studies of personal care product-health outcome associations.

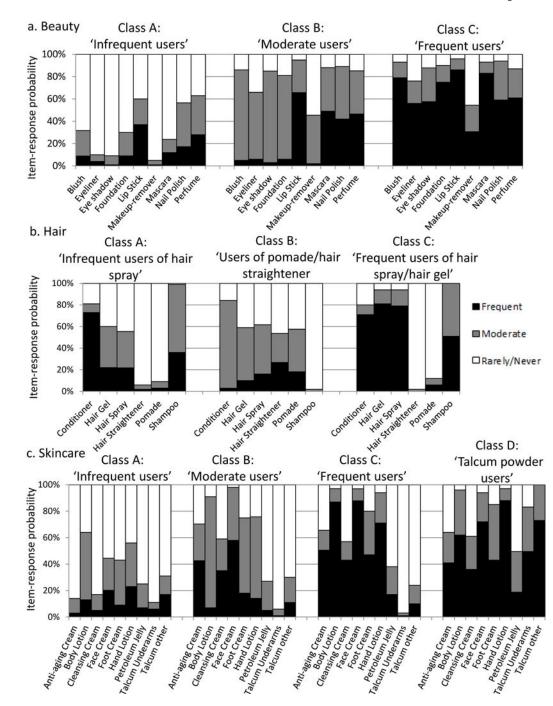
# CONCLUSION

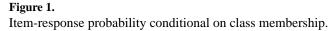
One active area of environmental health research is the investigation of associations between personal care product use and health outcomes.<sup>29–31</sup> This necessitates a thorough understanding of how exposures vary within a population and co-vary with other exposures. We used LCA to identify patterns of personal care product use among a nationwide group of women and found that for white women, those with the highest level of exposure to personal care products also tended to have used exogenous hormone medications. Understanding and accounting for such relationships is critical as researchers explore associations between personal care product use and health outcomes. Future studies on personal care product exposures and health impacts should consider common hormonal treatments as potential confounders.

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# A. White women

	Hair A	Hair B	Hair C	Skin A	Skin B	Skin C	Skin D
Beauty A	0.34	0.19	-0.37	0.45	0.00	-0.15	-0.26
Beauty B	0.08	0.10	-0.09	0.07	0.18	0.05	-0.13
Beauty C	-0.36	-0.22	0.39	-0.42	-0.08	0.13	0.34
Hair A				0.24	0.04	-0.10	-0.17
Hair B				0.17	0.07	0.00	-0.17
Hair C				-0.26	-0.04	0.10	0.20

# B. Black women

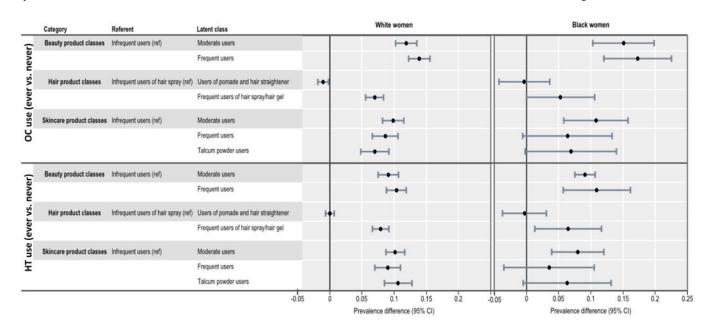
	Hair A	Hair B	Hair C	Skin A	Skin B	Skin C	Skin D
Beauty A	-0.01	0.03	-0.09	0.39	-0.02	-0.11	-0.20
Beauty B	-0.02	0.03	0.00	-0.11	0.14	0.09	0.00
Beauty C	0.03	-0.05	0.10	-0.35	-0.07	0.08	0.25
Hair A				-0.06	-0.02	-0.01	0.05
Hair B				0.07	0.02	0.00	-0.05
Hair C				-0.11	-0.03	0.05	0.06

r=-1.0	r=0	r=1.0

# Figure 2.

Spearman correlation coefficients of posterior probabilities of latent class membership for beauty products, hair products, and skincare products among (a) white and (b) black women r = Spearman Correlation Coefficient.

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#### Figure 3.

Prevalence differences (PDs) and 95% confidence intervals (CIs) for oral contraceptive (OC) and post-menopausal hormone therapy (HT) by race.

OC: Oral contraceptives.

HT: Post-menopausal hormone therapy

Statistically significant results are those where the 95% CI excludes a prevalence difference of zero.

Descriptive characteristics of sample population.

	Ν	%
Age: 5-year categories		
35-39 years	1838	4%
40-44 years	4016	9%
45-49 years	7031	15%
50-54 years	9015	19%
55–59 years	9396	20%
60-64 years	7227	15%
64–69 years	5489	12%
70-74 years	2885	6%
Race/ethnicity		
Non-Hispanic White	42,453	91%
Non-Hispanic Black	4452	9%
Menopausal status		
Missing	264	0.6%
Pre-menopausal	16,550	35%
Post-menopausal	30,091	64%
Highest level of education completed		
Missing	4	0
< High school	369	0.8%
HS or equivalent	6574	14%
Some college but no degree	9185	20%
Associate, technical, or bachelor's degree	19,323	41%
More than bachelor's	11,450	24%
Oral contraceptive use		
Missing	36	0.1%
Never	7161	15%
Ever	39,708	85%
Post-menopausal hormone therapy		
Missing	155	0.3%
Never	23,545	50%
Ever	23,205	49%
Geographic location		
Missing	34	0.1%
Northeast	8165	17%
Midwest	13,234	28%
South	15,574	33%
West	9906	21%

# Indicators of fit for latent class analysis.

	AIC	BIC	Entropy
Beauty classes			
2	78,666	78,993	0.77
3	35,717	36,210	0.83
4	29,716	30,378	0.79
5	24,355	25,184	0.74
6	22,393	23,390	0.71
Hair classes			
2	14,239	14,459	0.96
3	7936	8271	0.67
4	5237	5686	0.61
5	3416	3980	0.64
6	2046	2725	0.67
Skin classes			
2	48,055	48,381	0.72
3	38,337	38,831	0.71
4	30,150	30,812	0.73
5	26,769	27,598	0.69
6	24,092	25,089	0.67

Abbreviations: AIC, Akaike information criterion; BIC, Bayesian information criterion.

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Latent class labels and descriptions by product category.

Category/class	Label	Description		
Beauty product classes	A Infrequent users	Infrequent use of eye shadow, eyeliner, mascara, foundation, and blush; relatively (to the other classes) infrequent use of make-up remover, perfume, an lipstick.		
	B Moderate users	Intermediate use of eye shadow, eyeliner, mascara, foundation, blush, make-up remover, perfume, and lipstick (relative to the other classes).		
	C Frequent users	Frequent use of eye shadow, eyeliner, mascara, foundation, blush, make-up remover, nail polish, and lipstick.		
Hair product classes	A Infrequent users of hair spray	Relatively infrequent use of hair spray, hair gel compared with Hair-C (similar to class Hair-B); frequent use of shampoo, conditioner; infrequent use of pomade and hair straightener		
	B Users of pomade and hair straightener	Infrequent use of shampoo, hair gel; intermediate use of pomade, hair straightener, and hair spray		
	C Frequent users of hair spray/ hair gel	Frequent use of hair spray, hair gel, shampoo, conditioner; infrequent use of pomade and hair straightener		
Skincare product classes	A Infrequent users	Infrequent use of lotions, creams, talcum powder		
	B Moderate users	Intermediate use of lotions, creams; infrequent use of talcum powder		
	C Frequent users	Frequent use of face creams and lotions; infrequent use of talcum powder		
	D Talcum powder users	Second most frequent use of lotions; most frequent use of talcum powder		

# Latent class distribution by race.

	Class descriptor	White	Black	Total
		N (%)	N (%)	N (%)
Beauty products				
Total		42,558	4461	47,019
Beauty A	Infrequent users	9231 (22%)	1282 (29%)	10,513 (22%)
Beauty B	Moderate users	16,010 (38%)	2156 (48%)	18,166 (39%)
Beauty C	Frequent users	16,762 (39%)	746 (17%)	17,508 (37%)
Missing		555 (1%)	277 (6%)	832 (2%)
Hair products				
Total		42,558	4461	47,019
Hair A	Infrequent users of hair spray	20,950 (49%)	1011 (23%)	21,961 (47%)
Hair B	Users of pomade and hair straightener	1182 (3%)	2989 (67%)	4171 (9%)
Hair C	Frequent users of hair spray and hair gel	19,659 (46%)	173 (4%)	19,832 (42%)
Missing		767 (2%)	288 (6%)	1055 (2%)
Skincare products	\$			
Total		42,558	4461	47,019
Skin A	Infrequent users	7954 (19%)	812 (18%)	8766 (19%)
Skin B	Moderate users	18,617 (44%)	2192 (49%)	20,810 (44%)
Skin C	Frequent users	10,271 (24%)	551 (12%)	10,822 (23%)
Skin D	Talcum powder users	5158 (12%)	628 (14%)	5786 (12%)
Missing		558 (1%)	278 (6%)	836 (2%)