Breast reconstruction is among the 10 most common reconstructive surgical procedures in the United States. The number of procedures has increased steadily since 2000, with nearly 100,000 procedures performed in 2013 alone. A primary, growing reason for breast reconstruction is to restore breasts to their presurgical form following mastectomy due to breast cancer. Breast reconstruction is also used to correct congenital deformities, malformations, and trauma to the breast. Natrelle silicone gel-filled breast implants (Allergan, Irvine, Calif.) are available in a range of implant options and are approved by the US Food and Drug Administration.
The Breast Implant Follow-up Study (BIFS-001) is a large, multicenter, 10-year observational study being conducted as part of a US FDA requirement to provide postapproval data on safety concerns associated with the use of silicone-filled breast implants. The primary objective of this study is to compare the long-term safety and effectiveness of Natrelle silicone breast implants with those of saline implants or national norms in subjects who underwent primary augmentation, revision-augmentation, primary reconstruction, or revision-reconstruction. This ongoing study has enrolled more than 50,000 subjects. Although primary outcome data will continue to be collected over a 10-year period, available baseline data provide an opportunity to examine the demographic, health, lifestyle, and surgical characteristics of a large population of women requesting to undergo breast reconstruction. Because the analysis of safety endpoints in BIFS-001 will involve comparisons between silicone and saline implant groups that are self-selected rather than randomly assigned, it is especially important to understand how characteristics such as health and lifestyle factors that may potentially impact safety outcomes differ at baseline.

The current analysis consequently describes demographic, health, lifestyle, and surgical characteristics for subjects enrolled in BIFS-001 who underwent either primary reconstruction or revision-reconstruction surgery and baseline rates for some of the important safety outcomes. An analysis of subjects who underwent either primary augmentation or revision-augmentation surgery is reported separately. This analysis aimed to assess how characteristics of women undergoing primary reconstruction procedures differ from those in the revision-reconstruction population and how women who select silicone implants differ at baseline from those who choose saline implants. Such differences may have important implications for the long-term safety profile of silicone implants in clinical practice.

**METHODS**

**Study Design**

BIFS-001 is a long-term observational study (ClinicalTrials.gov identification number NCT00443274) comparing outcomes between subjects who underwent primary augmentation, revision-augmentation, primary reconstruction, or revision-reconstruction procedures. Baseline information was collected upon each subject’s decision to undergo an implantation procedure. Subjects must have received unilateral or bilateral silicone or saline implants during surgery, and all silicone implants were required to be Natrelle devices. All subjects agreed at enrollment to complete follow-up questionnaires via the Internet, telephone interviews, or postal mail once annually over 10 years.

The study followed Good Clinical Practice guidelines, was approved by the institutional review board covering each participating study site, conformed with World Health Organization guidelines, and was registered with the FDA at http://www.clinicaltrials.gov (NCT00443274). All participants provided written informed consent before enrollment.

**Subjects**

This analysis includes women aged 18 years or older who desired primary reconstruction or revision-reconstruction. Women screened for study eligibility were required to be fluent in English or Spanish. Enrolled subjects then completed surgery, receiving one implant or matching implants (matching implants were both silicone or both saline).

Subjects were ineligible for study participation if they were currently implanted with saline implants after having previously received silicone breast implants. Subjects were also excluded if they were transgender or if the investigator decided that a subject was not a suitable candidate for long-term observation.

**Assessments**

Assessments recorded at baseline included demographic characteristics (age, race, height, and weight) and subject-reported health and lifestyle characteristics (marital status, education, occupation, smoking status, alcohol use, and substance abuse history). Body mass index (BMI) was calculated based on height and weight. Surgical characteristics (implant type, placement location, style and size of implant,
and incision size and site) were documented by investigators following surgery. Long-term safety outcome measures assessed at baseline included history of neurologic disease, connective tissue disease (eg, rheumatoid arthritis and fibromyalgia), cancer (basal cell/squamous cell, brain, breast, cervical/vulvar, and lung), suicide attempt or thoughts about suicide, and reproductive or lactation complications.

**Statistical Analysis**

Statistical analysis of the 10-year safety data will include comparisons of adverse event rates between women choosing silicone breast implants and those choosing saline breast implants. Therefore, baseline and surgical differences between the silicone and saline groups were analyzed. For primary-reconstruction and revision-reconstruction groups, comparisons between silicone and saline groups were based on a 2-sided z-test for continuous data and a 2-sided chi-square test for categorical data. Comparisons were also made between the primary reconstruction and revision-reconstruction populations, using descriptive statistics only.

**RESULTS**

**Subjects**

A total of 56,616 eligible subjects were enrolled in the BIFS-001 study from February 2007 through March 2010 (silicone, n = 41,163; saline, n = 15,453) at 1116 sites. Of these, 5637 subjects underwent breast reconstruction procedures, including 4887 (86.7%) who underwent primary reconstruction and 750 (13.3%) who underwent revision-reconstruction. Most subjects in both the primary reconstruction and revision-reconstruction groups received silicone implants (96.2% and 94.1%, respectively).

### Demographics

The median age in subjects in the primary reconstruction group was 50 years at implantation (range, 18–86 years), and the majority of subjects (82.9%) were white. Approximately half of the subjects (48.4%) had a normal BMI, whereas 46.2% were overweight or obese. Within the primary reconstruction group, silicone and saline groups differed statistically with regard to several demographic characteristics. Subjects selecting silicone implants were significantly older ($P = 0.0024$) and had a lower mean BMI ($P = 0.0259$) than subjects receiving saline implants (Table 1). Race/ethnicity distribution also differed significantly between implant groups ($P = 0.0342$), with a greater percentage of subjects in the silicone group versus the saline group being white and a greater percentage of subjects in the saline group versus the silicone group being Hispanic.

Relative to the primary reconstruction group, subjects in the revision-reconstruction group were older, with a median age difference of approximately 3 years (50 vs 53 years; Table 1). When divided by

| Characteristic                  | Primary Reconstruction | Revision-Reconstruction | $P$   | $P$   |
|--------------------------------|------------------------|-------------------------|-------|-------|
|                                | Silicone ($n = 4701$)  | Saline ($n = 186$)      |       |       |
|                                |                        |                         |       |       |
| Age, y (SD)                    | 50.1 (10.97)           | 47.6 (13.09)            | 0.0024|       |
| Median (min–max)               | 50 (18–86)             | 49 (18–80)              |       |       |
| Age at implantation, n (%)     |                        |                         | <0.0001|       |
| 18–21 y                        | 74 (1.6)               | 11 (5.9)                |       |       |
| 22–29 y                        | 83 (1.8)               | 10 (5.4)                |       |       |
| 30–39 y                        | 534 (11.4)             | 25 (13.4)               |       |       |
| 40–49 y                        | 1603 (34.1)            | 53 (28.5)               |       |       |
| 50–59 y                        | 1495 (31.8)            | 52 (28.0)               |       |       |
| 60–69 y                        | 734 (15.6)             | 30 (16.1)               |       |       |
| ≥70 y                          | 176 (3.7)              | 3 (1.6)                 |       |       |
| Unknown                        | 2 (<0.1)               | 2 (1.1)                 |       |       |
| Mean BMI, kg/m² (SD)           | 25.8 (5.4)             | 26.7 (5.9)              | 0.0259|       |
| BMI <18.5, %                   | 2.3                    | 1.0                     |       |       |
| BMI 18.5–24.9, %               | 48.5                   | 45.2                    |       |       |
| BMI ≥25, %                     | 46.1                   | 47.8                    |       |       |
| Unknown, %                     | 3.1                    | 6.5                     |       |       |
| Race, n (%)                    |                        |                         |       |       |
| White                          | 3907 (83.1)            | 144 (77.4)              | 0.0342| 0.1754|
| Hispanic                       | 263 (5.6)              | 17 (9.1)                | 28 (4.0)|       |
| Black                          | 235 (5.0)              | 7 (3.8)                 | 36 (5.1)|       |
| Asian                          | 103 (2.2)              | 3 (1.6)                 | 15 (2.1)|       |
| Other                          | 103 (2.2)              | 8 (4.3)                 | 10 (1.4)|       |
| Unknown                        | 90 (1.9)               | 7 (3.8)                 | 12 (1.7)|       |

Table 1. Subject Demographics by Procedure and Type of Implant
10-year intervals, most subjects in both the primary reconstruction and revision-reconstruction groups received implants between the ages of 40 and 59 (Fig. 1). The mean BMI and BMI distribution were similar in the primary reconstruction and revision-reconstruction groups, although the percentage of subjects who were overweight/obese was lowest in the saline revision-reconstruction group. Among subjects undergoing primary reconstruction or revision-reconstruction, the age distribution at implantation differed significantly between the silicone and saline groups ($P < 0.0001$, primary reconstruction; $P = 0.0135$, revision-reconstruction). Subjects between the ages of 40 and 59 were more likely to receive silicone than saline implants, whereas subjects aged 39 years and younger were more likely to receive saline implants.

**Lifestyle Characteristics**

Between the primary reconstruction and revision-reconstruction groups, the lifestyle characteristics were generally similar (Table 2). Among all subjects, approximately 67% were married and more than 75% had attended college. Further, almost half of the subjects in both the primary reconstruction and revision-reconstruction groups held a professional occupation. About 50% of the subjects in both groups were nonsmokers, and fewer than 15% of subjects were current smokers. More than 86% of all subjects consumed 3 or fewer alcoholic drinks per week, including approximately 28% of subjects who did not drink at all. In both groups, fewer than 3% of subjects undergoing primary reconstruction or revision-reconstruction had ever been treated for substance abuse. In both the primary recon-
struction and revision-reconstruction groups, no significant differences were found between subjects receiving silicone versus saline implants with regard to any of the lifestyle factors examined, with the exception of marital status for women undergoing revision-reconstruction ($P = 0.0170$). Among women undergoing revision-reconstruction, a greater percentage of those receiving silicone versus saline implants were married, whereas a greater percentage of those receiving saline versus silicone implants were widowed.

### Medical History
Baseline rates of selected medical diagnoses that are safety endpoints in BIFS-001 are presented for primary reconstruction and revision-reconstruction groups in Table 3. Among subjects undergoing primary reconstruction, there was a significantly higher overall rate of previous cancer diagnoses for the silicone group versus the saline group ($P < 0.0001$). The rate of breast cancer was significantly higher in the silicone group compared with the saline group at baseline ($P < 0.0001$), but no between-group differences were observed in previous basal cell/squamous cell carcinoma, brain, lung, or cervical/vulvar cancer rates. The percentages of women who reported previous suicide attempts or who had thoughts about suicide were similar in the silicone and saline groups. The percentage of subjects who were ever pregnant was similar between the silicone (89.2%) and saline (86.5%) groups. However, more than twice as many women in the silicone group who had been pregnant had tried to breastfeed ($P < 0.0001$) and, of those, the proportion of women who had difficulties in breastfeeding was significantly higher at baseline in women who chose saline versus silicone implants.

### Table 2. Lifestyle Characteristics of Subjects by Procedure and Type of Implant

| Characteristic                  | Primary Reconstruction |          | Revision-Reconstruction |          |
|--------------------------------|------------------------|----------|-------------------------|----------|
|                                | Silicone ($n = 4701$)  | Saline ($n = 186$) | $P$          | Silicone ($n = 706$)  | Saline ($n = 44$) | $P$          |
| Education, $n (%)$             | High school*           | 963 (20.5) | 44 (23.7) | 0.1079 | 181 (25.6) | 10 (22.7) | 0.2223 |
|                                | Some college           | 1357 (28.9) | 59 (31.7) |          | 202 (28.6) | 15 (34.1) |          |
|                                | College graduate       | 1319 (28.1) | 45 (24.2) |          | 177 (25.1) | 15 (34.1) |          |
|                                | Some postgraduate      | 260 (5.5)  | 8 (4.3)   |          | 43 (6.1)   | 1 (2.3)   |          |
|                                | Postgraduate degree    | 675 (14.4) | 20 (10.8) |          | 85 (12.0)  | 1 (2.3)   |          |
|                                | Unknown                | 127 (2.7)  | 10 (5.4)  |          | 18 (2.5)   | 2 (4.5)   |          |
| Marital status, $n (%)$        | Married                | 3176 (67.6) | 118 (63.4) | 0.1433 | 478 (67.7) | 26 (59.1) | 0.0170 |
|                                | Never married          | 400 (8.5)  | 22 (11.8) |          | 55 (7.8)   | 0          |          |
|                                | Separated/divorced     | 735 (15.6) | 23 (12.4) |          | 111 (15.7) | 7 (15.9)  |          |
|                                | Cohabitating with partner | 83 (1.8)   | 4 (2.2)   |          | 12 (1.7)   | 2 (4.5)   |          |
|                                | Widowed                | 196 (4.2)  | 11 (5.9)  |          | 37 (5.2)   | 7 (15.9)  |          |
|                                | Other/unknown          | 111 (2.4)  | 8 (4.3)   |          | 13 (1.8)   | 2 (4.5)   |          |
| Occupation, $n (%)$            | Professional           | 2266 (48.2) | 85 (45.7) | 0.3536 | 313 (44.3) | 21 (47.7) | 0.9520 |
|                                | Homemaker              | 801 (17.0) | 29 (15.6) |          | 136 (19.3) | 7 (15.9)  |          |
|                                | Clerical               | 769 (15.1) | 30 (16.1) |          | 99 (14.0)  | 6 (13.6)  |          |
|                                | Unemployed             | 287 (6.1)  | 9 (4.8)   |          | 52 (7.4)   | 5 (11.4)  |          |
|                                | Trade                  | 140 (3.0)  | 6 (3.2)   |          | 24 (3.4)   | 1 (2.3)   |          |
|                                | Manual labor           | 120 (2.6)  | 4 (2.2)   |          | 15 (2.1)   | 1 (2.3)   |          |
|                                | Student                | 73 (1.6)   | 7 (3.8)   |          | 11 (1.6)   | 1 (2.3)   |          |
|                                | Unknown                | 305 (6.5)  | 16 (8.6)  |          | 56 (7.9)   | 2 (4.5)   |          |
| Smoking history, $n (%)$       | Nonsmoker              | 2780 (59.1) | 99 (53.2) | 0.0666 | 374 (53.0) | 21 (47.7) | 0.5298 |
|                                | Ex-smoker              | 1520 (32.3) | 63 (33.9) |          | 247 (35.0) | 15 (34.1) |          |
|                                | Current smoker         | 295 (6.3)  | 15 (8.1)  |          | 72 (10.2)  | 6 (13.6)  |          |
|                                | Unknown                | 106 (2.3)  | 9 (4.8)   |          | 13 (1.8)   | 2 (4.5)   |          |
| Alcohol consumption, $n (%)$   | Never drink            | 1344 (28.6) | 59 (31.7) | 0.3239 | 195 (27.6) | 11 (25.0) | 0.9246 |
|                                | 1-6 drinks/mo          | 2005 (42.7) | 84 (45.2) |          | 283 (40.1) | 19 (43.2) |          |
|                                | 1-3 drinks/wk          | 739 (15.7) | 26 (14.0) |          | 119 (16.9) | 8 (18.2)  |          |
|                                | 1 drink/d              | 284 (6.0)  | 6 (3.2)   |          | 59 (8.4)   | 4 (9.1)   |          |
|                                | ≥2 drinks/d            | 121 (2.6)  | 1 (0.5)   |          | 25 (3.5)   | 0          |          |
|                                | Unknown                | 208 (4.4)  | 10 (5.4)  |          | 25 (3.5)   | 2 (4.5)   |          |
| Treated for substance abuse, $n (%) | Yes                   | 87 (1.9)   | 1 (0.5)   | 0.2667 | 15 (2.1)   | 0          | 0.4787 |
|                                | No                    | 4342 (92.4) | 171 (91.9) |          | 660 (93.5) | 41 (93.2) |          |
|                                | Unknown                | 272 (5.8)  | 14 (7.5)  |          | 31 (4.4)   | 3 (6.8)   |          |

*Includes vocational schools.
The rate of breast-related disease was also significantly higher in the silicone group \((P = 0.002)\). As observed in the primary reconstruction subjects, revision-reconstruction subjects who selected silicone implants had a significantly higher overall rate of previ-

### Table 3. Selected Medical History by Procedure and Type of Implant*

| Characteristic                        | Primary Reconstruction |          |          |          |          |          |          |
|---------------------------------------|------------------------|----------|----------|----------|----------|----------|----------|
|                                       | Silicone \((n = 4650)\) | Saline \((n = 182)\) |          | Silicone \((n = 700)\) | Saline \((n = 43)\) |          |          |
| Neurologic diagnoses, \(n (\%)\)      | 77 (1.7)               | 1 (0.5)  | 0.2453   | 11 (1.6) | 0 (0.0)  | 0.4076   |
| Cancer diagnoses, \(n (\%)\)         | 3986 (85.7)            | 126 (69.2)| \(<0.0001\) | 537 (76.6) | 23 (53.5) | 0.0007   |
| Basal cell carcinoma or squamous cell carcinoma | 213 (4.6)  | 6 (3.3)  | 0.4140   | 51 (7.3) | 1 (2.3)  | 0.2159   |
| Brain                                 | 3 (0.1)                | 0        | 0.7318   | 1 (0.1)  | 0        | 0.8041   |
| Breast                                | 3703 (79.6)            | 122 (67.0)| \(<0.0001\) | 491 (70.1) | 21 (48.8) | 0.0034   |
| Cervical/vulvar                       | 50 (1.1)               | 3 (1.6)  | 0.4665   | 7 (1.0)  | 0        | 0.5100   |
| Lung                                  | 12 (0.3)               | 0        | 0.4926   | 2 (0.3)  | 0        | 0.7256   |
| Mental health, \(n (\%)\)            | 45 (1.0)               | 1 (0.5)  | 0.5686   | 11 (1.6) | 0        | 0.4076   |
| Attempted suicide                     | 300 (6.5)              | 11 (6.0) | 0.8260   | 44 (6.3) | 0        | 0.0901   |
| Thoughts about suicide                | 3900 (89.2)            | 147 (86.5)| 0.2537   | 598 (91.7) | 38 (90.5) | 0.7781   |
| Ever pregnant, \(n (\%)\)            | 1665 (42.6)            | 30 (20.4) | \(<0.0001\) | 223 (37.3) | 7 (18.4)  | 0.0189   |
| Attempted suicide                     | 929 (55.8)             | 10 (33.3) | 0.0142   | 127 (57.0) | 5 (71.4)  | 0.4456   |
| Mastitis                              | 215 (12.9)             | 7 (23.3)  | 0.0936   | 27 (12.1) | 1 (14.3) | 0.8622   |
| Low milk production                   | 297 (17.8)             | 6 (20.0)  | 0.7593   | 32 (14.3) | 0        | 0.2801   |
| Infertility, \(n (\%)\)              | 432 (11.1)             | 14 (9.5)  | 0.5611   | 46 (7.7)  | 3 (7.9)  | 0.9638   |
| Breast-related disease, \(n (\%)\)   | 1870 (47.8)            | 47 (32.0) | 0.0002   | 310 (51.8) | 13 (34.2) | 0.0351   |

*Based on the number of subjects who filled out the baseline questionnaire.
†Of those ever pregnant.
‡Of those who tried to breastfeed.

### Table 4. Surgical Characteristics by Procedure and Type of Implant*

| Parameter                        | Primary Reconstruction |          |          |          |          |          |          |
|----------------------------------|------------------------|----------|----------|----------|----------|----------|----------|
|                                  | Silicone \((n = 4701)\) | Saline \((n = 324)\) |          | Silicone \((n = 1211)\) | Saline \((n = 71)\) |          |          |
| Bilateral implants, \(n (\%)\)   | 3439 (73.2)            | 138 (74.2)| 0.7537   | 505 (71.5) | 27 (61.4) | 0.1496   |
| Implant, \(n\)                   | 8140 (n = 324)         | 324      | NA       | 324      | NA       | NA       |
| Submuscular–partial               | 4890 (60.1)            | 165 (50.9)| 0.0012   | 682 (56.3) | 30 (42.3) | 0.1100   |
| Submuscular–complete              | 2728 (33.5)            | 141 (43.5)|          | 401 (33.1) | 32 (45.1) |          |
| Subglandular                      | 445 (5.5)              | 13 (4.0)  |          | 110 (9.1)  | 7 (9.9)  |          |
| Unknown                           | 77 (0.9)               | 5 (1.5)   |          | 18 (1.5)   | 2 (2.8)  |          |
| Surface type, \(n (\%)\)         | 7488 (92.0)            | 235 (72.5)| NA       | 1066 (88.3) | 57 (80.3) | NA       |
| Textured                          | 649 (8.0)              | 89 (27.5) | NA       | 142 (11.7) | 14 (19.7) | NA       |
| Unknown                           | 3 (<0.1)               | 0         | 0        | 0         | 0        |          |
| Implant style, \(n (\%)\)        | 514 (6.3)              | NA       | NA       | 101 (8.3)  | NA       | NA       |
| 15                                | 1934 (23.8)            | NA       | 344 (28.4) | NA       | NA       |          |
| 20                                | 4141 (50.9)            | NA       | 452 (37.3) | NA       | NA       |          |
| 40                                | 32 (0.4)               | NA       | 18 (1.5)  | NA       | NA       |          |
| 45                                | 422 (5.2)              | NA       | 47 (3.9)   | NA       | NA       |          |
| 110                               | 81 (1.0)               | NA       | 25 (2.1)  | NA       | NA       |          |
| 115                               | 277 (3.4)              | NA       | 63 (5.2)  | NA       | NA       |          |
| 120                               | 363 (4.5)              | NA       | 90 (7.4)  | NA       | NA       |          |
| 68HP                              | 60 (18.5)              | NA       | 14 (19.7) | NA       | NA       |          |
| 68LP                              | 11 (3.4)               | NA       | 2 (2.8)   | NA       | NA       |          |
| 68MP                              | 85 (26.2)              | NA       | 28 (39.4) | NA       | NA       |          |
| 168                               | 24 (7.4)               | NA       | 7 (9.9)   | NA       | NA       |          |
| 360LF                             | 52 (16.0)              | NA       | 4 (5.6)   | NA       | NA       |          |
| 468                               | 31 (9.6)               | NA       | 8 (11.3)  | NA       | NA       |          |
| Unknown                           | 376 (4.6)              | 61 (18.8) | 71 (5.9)  | 8 (11.3)  |          |          |

*\(n\) values refer to the number of implants.
NA, not applicable.
ous cancer diagnoses ($P = 0.0007$) and a higher rate of previous breast cancer diagnosis ($P = 0.0034$) at baseline, with rates of other cancer types included in the analysis not differing between the silicone and saline groups (Table 3). There was no difference between implant groups in the number of subjects reporting previous suicide attempts or thoughts about suicide. Among revision-reconstruction subjects, 91.7% of the silicone group and 90.5% of the saline group had ever been pregnant; of those, twice as many women in the silicone group compared with the saline group had tried to breastfeed ($P = 0.0189$). The baseline rate of breast-related disease in subjects undergoing revision-reconstruction was higher for those receiving silicone versus saline implants ($P = 0.0351$).

**Surgical Characteristics**

The majority of subjects received bilateral breast implants (Table 4). For those who underwent primary reconstruction, 73.2% and 74.2% received bilateral silicone and saline implants, respectively. Similarly, among subjects undergoing revision-reconstruction, 71.5% and 61.4% received bilateral silicone and saline implants, respectively. The majority of implants for primary reconstruction and revision-reconstruction were placed submuscularly, with partial submuscular placement occurring more frequently than complete submuscular placement for silicone implants (60.1% vs 33.5%, primary reconstruction; 56.3% vs 33.1%, revision-reconstruction). By contrast, saline implants were more evenly distributed between partial and complete submuscular placement for both primary reconstruction (50.9% vs 43.5%) and revision-reconstruction (42.3% vs 45.1%), respectively. The distribution of implant locations differed significantly only for subjects undergoing primary reconstruction ($P = 0.0012$), in whom subglandular placement was more common and complete submuscular placement was less common for silicone versus saline implants. Although submuscular placement was also more common in subjects who underwent revision-reconstruction procedures, the frequency of subglandular placement of both silicone and saline implants was higher compared with that in primary reconstruction procedures (9.1% and 9.9% vs 5.5% and 4.0%, respectively).

**Implant Styles and Sizes**

The majority of silicone and saline implants were smooth in both the primary reconstruction and revision-reconstruction groups (Table 4). All silicone implants were round; Natrelle shaped implants were not available at the time of this study. The most frequently used silicone implant styles in both the primary reconstruction and revision-reconstruction groups were, respectively, Natrelle style 20 (50.9% and 37.3%), which is considered high profile, and style 15 (23.8% and 28.4%), which is considered midrange profile. The most frequently used saline implant styles in both indications were, respectively, the moderate-profile Natrelle style 68MP (26.2% and 39.4%) and the high-profile style 68HP (18.5% and 19.7%).

In the primary reconstruction group, the most common implant size range for both silicone and saline implants was 400–499 cm$^3$ (Fig. 2A). More specifically, the most commonly used implant size was 450–474 cm$^3$, followed by 400–424 cm$^3$, for both silicone and saline implants. The most common implant size range in the revision-reconstruction group was 300–399 cm$^3$ for both silicone and saline implants (Fig. 2B). More specifically, the most commonly used implant size was 350–374 cm$^3$ followed by 375–399 cm$^3$ for silicone implants and a tie between 350–374 cm$^3$ and 475–499 cm$^3$ for saline implants. Size distributions differed significantly between the silicone and saline implant groups in subjects undergoing primary reconstruction ($P = 0.0096$) but did not differ significantly in subjects undergoing revision-reconstruction ($P = 0.3186$).

The percentage of women selecting saline implants was higher than the percentage selecting silicone for implant sizes from 200 cm$^3$ up to 499 cm$^3$ for primary reconstruction and for implant sizes up to 499 cm$^3$ for revision-reconstruction. When subjects were divided by 10-year age intervals, the majority of subjects aged 50 years or older who underwent primary reconstruction received either 400–499 cm$^3$ or 500–599 cm$^3$ silicone or saline implants, whereas subjects aged 49 years or younger were more likely to receive either 300–399 cm$^3$ or 400–499 cm$^3$ for both silicone and saline implants. None of the subjects aged 18–21 years received saline implants for revision-reconstruction procedures.

When subjects were analyzed by BMI category, an association was observed between subject weight and implant size for those who underwent primary reconstruction procedures. Subjects who were underweight or within normal weight primarily received implants with volumes between 300 and 499 cm$^3$. By contrast, overweight subjects mostly received 500- to 699-cm$^3$ implants, whereas obese subjects primarily received 700-cm$^3$ to greater than 800-cm$^3$ implants. No size differences were noted between subjects receiving silicone or saline implants. A similar but less prominent trend was observed in subjects who underwent revision-reconstruction procedures.
Incision Sizes and Sites

In the primary reconstruction group, the majority of silicone implants required incision sizes of 5.0–5.9 cm or 6.0–6.9 cm (Fig. 3). Most saline implants required incision sizes of 4.0–4.9 cm or 5.0–5.9 cm. This trend was also evident in the revision-reconstruction group, wherein silicone implants more often required incision sizes of 5.0–5.9 or 6.0–6.9 cm. By contrast, the majority of saline implants required somewhat smaller incision sizes of 3.0–3.9 or 4.0–4.9 cm. The most frequently used anatomical incision site in both primary reconstruction and revision-reconstruction groups was the mastectomy scar, followed by inframammary incision. Although these were the 2 most common incision sites for both implant types in each indication, the percentage of subjects with each incision site differed significantly between silicone and saline implants for both primary and revision-reconstruction procedures (both \( P < 0.0001 \); Table 5).

DISCUSSION

The current analysis provides a detailed examination of the demographics, lifestyle, health, and surgical characteristics of a large number of subjects undergoing primary reconstruction and revision-reconstruction breast implant procedures. The majority of women in these populations received silicone implants. Women who underwent primary reconstruction procedures were slightly younger than those who underwent revision-reconstruction procedures, although other baseline demographics were similar. However, based on percentages, Hispanics were the second most common ethnic group to undergo primary reconstruction, whereas blacks were the second most common racial group to undergo revision-reconstruction. The majority of subjects who underwent primary reconstruction used a mastectomy scar as the incision site, with either silicone or saline implants placed most frequently in a partial or complete submuscular position.
However, subjects who underwent revision-reconstruction procedures were just as likely to have saline implants placed using an inframammary incision as a mastectomy scar incision. Subjects who received saline implants for either primary reconstruction or revision-reconstruction generally had smaller incision sizes compared with those who received silicone implants, most likely because saline implants are not filled until after insertion.\(^9,10\) In addition, subjects who underwent revision-reconstruction procedures received slightly smaller-sized silicone and saline implants than did those who underwent primary reconstruction procedures.

Because subjects in this observational study were not randomly assigned but were self-selected into the silicone or saline implant groups, it will be critical to identify any differences between the populations choosing silicone versus saline implants so that these differences can be controlled statistically in the long-term safety analyses. Several significant differences observed between the silicone and saline groups at baseline could potentially influence...
the 10-year findings. Compared with women who chose saline implants, women who selected silicone implants for primary reconstruction were substantially older and had a lower mean BMI. Ethnicity patterns also differed significantly in this population, with a greater proportion of Hispanic subjects receiving saline versus silicone implants. Among subjects undergoing revision-reconstruction, those choosing saline implants were less likely to be married and were more likely to be widowed than were women receiving silicone implants. Such demographic and lifestyle characteristics may be associated with differential rates of BIFS-001 safety outcomes, including cancer,11–13 rheumatic and neurologic diseases,14,15 suicide attempts,16,17 and postoperative complications, such as capsular contracture and rippling.18

In the primary reconstruction population, the implant distributions were significantly skewed toward a larger size for silicone implants relative to saline implants. Although requiring confirmation, larger implant size may be a risk factor for breast implant rupture.19 Significant differences between the silicone and saline groups were also found among both primary and revision-reconstruction subjects for baseline rates of BIFS-001 outcomes, including previous cancer diagnoses, breast-related disease, and breastfeeding history and difficulties. There were also significant differences noted between silicone and saline implantation with regard to the percentage of subjects with each incision site.

The baseline data from BIFS-001 offer an opportunity to compare characteristics between a substantial cross-section of real-world reconstruction patients and populations previously enrolled in prospective, long-term clinical trials for silicone and/or saline implants. Subject demographic data and surgical characteristics reported for other breast reconstruction studies reveal similar baseline findings compared with those of the current study. In studies that reported demographic information, the majority of subjects were white, married, and had attended college; most subjects who underwent primary reconstruction procedures were aged in their late 40s, with those who had revision-reconstruction procedures being, on average, approximately 4 to 5 years older.20–26 With respect to surgical characteristics, the majority of subjects in these studies had breast implants placed submuscularly, using the mastectomy scar or other scar for the incision site.20–24,26 Incision sizes between 6.0 and 9.0 cm were favored for reconstruction procedures.23,24 Of the studies conducted solely using silicone implants, the majority (>70%) of subjects elected to undergo bilateral reconstruction procedures.22–24 One previously published study compared the baseline demographic, lifestyle, and surgical characteristics of subjects undergoing postmastectomy breast reconstruction with silicone \((n = 306)\) versus saline \((n = 176)\) implants.25 In that study, patients selecting silicone implants were significantly older at implantation (mean, 53.7 vs 51.3 years; \(P = 0.017\)) and were more likely to have bilateral placement (52.8% vs 39.9%; \(P = 0.008\)). No significant differences were observed between implant groups with regard to marital status, education level, or occupation, whereas the current analysis did identify differences in ethnicity between subjects choosing silicone versus saline implants for primary reconstruction.

BIFS-001 confirms many previously reported findings. However, as the sample size is considerably larger than the aforementioned studies, it offers an opportunity for surgeons to examine in greater depth the baseline characteristics of women receiving silicone or saline implants for breast reconstruction procedures. In addition, the BIFS-001 study assesses categories not described in recent breast reconstruction studies, including subject’s medical history, history of substance abuse, the distribution of implant sizes by age and by BMI, and the distribution of incision sizes. In regard to BMI and implant size, there was a direct association between larger silicone and saline implant sizes and higher BMI for both breast reconstruction procedures, but this association was less prominent for subjects who underwent revision-reconstruction.

Several study limitations deserve mention. A primary objective of BIFS-001 is to compare the long-term safety of Natrelle silicone breast implants with that of saline implants. However, women who choose to receive silicone versus saline breast implants may potentially differ in clinically important ways that can affect safety outcomes. This analysis of the baseline characteristics of subjects selecting silicone or saline implants for primary reconstruction or revision-reconstruction addresses this limitation by identifying demographic and clinical differences between the implant types that must be addressed in all future safety analyses. Additionally, some of the data in the BIFS-001 study, such as lifestyle information, were self-reported. However, it is likely that the robust overall sample size may mitigate any reporting bias. The sample size for subjects with saline implants was small because individuals seeking reconstruction or revision-reconstruction typically favor silicone implants. Another limitation is that a substantial majority of the subjects were white \((n = 4690)\), and relatively small numbers of Hispanic \((n = 310)\), black \((n = 282)\), and Asian \((n = 122)\) subjects were included. This observation may reflect disparities in rates of postmastectomy reconstruction.27–29
CONCLUSIONS

In conclusion, data derived from the BIFS-001 study offer surgeons an opportunity to make informed decisions and provide counseling with respect to the most appropriate implant attributes and surgical approaches for their patients who desire breast implants for primary or revisionary breast reconstruction. Further, these results identify baseline factors that will be critical when analyzing and interpreting the long-term safety outcomes for silicone-filled implants in BIFS-001.

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