



Antimicrobial Irrigation Difference in Implant-Based Reconstruction: Triple-Antibiotic Solution *versus* Chlorhexidine-Gluconate Based Solution

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Abstract

Background: Antimicrobial irrigation solutions are a widely practiced method to reduce infection and capsular contracture following mastectomy in implant-based breast reconstruction. This study compares the postoperative outcomes between a Triple Antibiotic Solution (TAS) and a 0.05% Chlorhexidine gluconate solution (CHG).

Methods: A multi-institutional retrospective cohort study was conducted, reviewing 304 breast reconstructions in 171 patients between April 2020 and July 2024. Patients were divided into two groups based on the use of TAS or CHG for implant irrigation. Postoperative complications, including hematoma, capsular contracture, infection, and the need for secondary procedures, were assessed at ≤ 30 and ≥ 60 days. Comparative statistical analysis was performed using t-tests, Wilcoxon rank-sum tests, Chi-squared tests, and multivariate regression.

Results: Of the 304 breast reconstructions, TAS was used in 249 breasts and CHG in 55 breasts. There were no statistically significant differences between the TAS and CHG groups in < 30 -day complications such as seroma [1 (0.4%) vs. 0 (0%), $p=0.82$], surgical site infection [2 (0.8%) vs. 0 (0%), $p=0.67$], or unplanned return to the operating room [16 (6.4%) vs. 2 (3.6%), $p=0.34$]. Similarly, no significant differences were observed between TAS and CHG groups in ≥ 60 -day outcomes, including capsular contracture [5 (2%) vs. 2 (3.6%), $p=0.37$].

Conclusion: TAS and CHG had no significant differences in rates of post-operative complications in IBR. Given the higher cost of chlorhexidine-based solutions and the absence of clear clinical superiority, TAS may present a more cost-effective option for IBR. Further prospective studies are warranted to establish optimal irrigation protocols in reconstructive breast surgery.

Introduction

Implant-based Breast Reconstruction (IBR) is a valuable reconstructive procedure for those undergoing oncologic mastectomy. Considerations of a successful operation include implant type, placement, and infection prophylaxis [1,2]. Capsular contracture, implant malposition, and implant infection following an implant placement can lead to revisions [3-5]. Past studies have reported implant-related infections for aesthetic breast augmentation to range from 1.1% to 2.5%. For IBR, the rate is cited to be slightly higher at 1% to 35% [6-8]

Efforts aimed at preventing complications such as infection or capsular contracture include utilizing irrigation solutions [2]. Studies have shown that the routine usage of antimicrobial pocket irrigation and implant soaking agents have led to decreased rates of implant-associated infection and subsequent capsular contracture [9,10]. The most widely used irrigation solutions include triple antibiotics regimens, iodine-based solutions, and 0.05% Chlorhexidine gluconate (CHG) formulations [11].

Current literature lacks evidence-based consensus on the optimal irrigation and implant-soaking solution for the reconstructive population [12]. There still remains skepticism within the surgical community that CHG formulations offer comparable efficacy to antibiotic solutions in

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preventing or reducing such complications [11] This study aimed to compare post-operative outcomes including rate of infection in a cohort of patients undergoing IBR with chlorhexidine-based solution and triple antibiotic solution.

Methods

Study design and data collection

After Institutional Board Approval (MedStar Health Research institution 2018-173), a multi-institution retrospective cohort study of consecutive patients undergoing IBR following mastectomy between April 2020 and July 2024 was conducted. All mastectomies were performed by MedStar breast surgeons (L.M.D.L.C., J.D.S., P.B.S., C.F.E., I.T.G.), and all IBR were performed by two senior authors (K.L.F. and S.J.). Patients over the age of 18 who underwent IBR for oncologic reconstruction were included. Patients undergoing breast augmentation with implants were not included.

Electronic medical records were reviewed for patient characteristics, oncologic and surgical history, operative details, and postoperative outcomes. Patient characteristics included age, sex, race, BMI, comorbidities such as hypertension or diabetes, smoking and alcohol history, and psychiatric history. Breast oncologic history was collected including chemotherapy and radiation therapy. Additional surgical details were also collected including breast resection type, incision type, implant placement location, implant texture, use of drains, use of acellular dermal matrix, lymph node dissection, and operative duration.

Patients were categorized into two groups based on the antibiotic solution used during the patients' reconstructive surgery: Triple-Antibiotic Solution (TAS) or 0.05% Chlorhexidine gluconate (CHG). Routinely, all breast pockets were irrigated first with sterile betadine and then two to three times with TAS or CHG solution. The breast implant was simultaneously bathed in the same type of irrigation solution used to irrigate the pocket. The primary outcomes are comprised of post-operative clinical outcomes. Postoperative complications such as hematoma, capsular contracture, surgical site infection, and secondary procedures were collected. Clinical records were evaluated postoperatively for complications at ≤ 30 days and ≥ 60 days. Time until follow-up was also obtained for each patient.

Statistical analysis

Comparative analyses were conducted between the TAS and CHG groups. STATA version 18.0 (StataCorp, College Station, TX) was used to conduct paired t-tests, and Wilcoxon rank-sum tests, and univariate Chi-squared tests to analyze normally distributed continuous, non-normally distributed continuous, and categorical data, respectively. Fisher's exact test was used for cell counts less than five. A P value <0.05 , determined statistical significance. To determine the effects of possible confounding variables identified in patient and surgical characteristic analysis, univariate regression analysis was performed. Variables that were identified as significant co-variables from univariate regression analysis were used to create multivariable models.

Results

A total of 304 breasts in 171 patients met inclusion criteria. TAS was used in 249 IBR (81.9%) and CHG was used in 55 IBR (18.1%). At the time of surgery, median age was 51 (Interquartile Range [IQR], 9) for the TAS group versus 49 (IQR, 12) for the CHG group ($p=0.73$). Median body mass index was similar ($p=0.86$) between TAS [25 (IQR

6.87)] and CHG [25.64 (IQR 7.64)] groups. Distribution of race varied significantly between groups, with both the TAS and CHG cohort identifying as White ($p=0.004$). Median Charlson Comorbidity Index for TAS and CHG groups were both 2 (IQR, 1). Patient characteristics are summarized in Table 1.

Table 2 summarizes surgical and oncologic histories between the TAS and CHG cohorts. A higher number of breasts in the TAS cohort, compared to the CHG cohort, had prior breast reduction ($n=2$, 0.8% vs. $n=0$, 0%, $p=0.02$) and prior breast reconstruction ($n=27$, 10.8% vs. $n=4$, 7.3%, $p=0.02$). Prior lumpectomy in breasts between the TAS and CHG cohorts was 9 (3.6%) vs. 2 (3.6%), respectively ($p=0.02$). There was a lower proportion of breasts in the TAS cohort that had previous excisional biopsy compared to the CHG cohort ($n=11$, 4.4% vs. $n=7$, 12.73%, $p=0.02$). Concerning chemotherapy history, of the TAS cohort, 31 (12.9%) breasts received neoadjuvant chemotherapy, 91 (38%) received adjuvant chemotherapy, and 17 (7.1%) received both. Of the CHG cohort, 7 (13.5%) breasts received neoadjuvant chemotherapy, 6 (11.5%) received adjuvant chemotherapy, and 2 (3.9%) received both. There was a significant difference in chemotherapy history between TAS and CHG groups ($p=0.001$). Radiation history was similar between TAS and CHG groups: respectively, 8 (3.5%) vs. 1 (1.9%) breast received neoadjuvant radiation, 62 (25.9%) vs. 6 (11.5%) received adjuvant radiation, and 1 (0.4%) vs. 0 (0%) received both ($p=0.09$).

Operative details

Details of the index surgical procedure (Mastectomy and subsequent IBR) are summarized in Table 3. Mastectomy type varied across groups ($p=0.001$): of the TAS breasts, 156 (61.5%) received nipple sparing mastectomy, 78 (31.6%) received a skin sparing mastectomy, 5 (2%) received a simple mastectomy, 2 (0.8%) received a modified radical mastectomy and of the CHG breasts, 44 (80%) received a nipple sparing mastectomy, 6 (10.9%) received a skin sparing mastectomy, and 5 (9.1%) received a simple mastectomy. Regarding the incision type, of the TAS cohort, 202 (81.1%) of breasts had an inframammary fold incision, 15 (6%) had an elliptical incision, 24 (9.6%) had a wise pattern incision, and 1 (0.4%) had a circumareolar incision. Of the CHG cohort, 51 (92.7%) of breasts had an inframammary fold incision and 4 (7.27%) had a wise pattern incision. There was no significant difference between incision types ($p=0.19$). Mastectomy weight was similar between TAS and CHG breasts: 305 grams (IQR 304.5) vs. 242.5 grams (IQR 262), respectively ($p=0.14$). Drains were placed in all patients of the TAS and CHG cohorts: 249 (100%) vs. 55 (100%), respectively.

Duration of surgery was similar between TAS and CHG cohorts: 205 minutes (IQR 100) vs. 157 minutes (IQR 170), respectively. The majority of IBR were performed as Direct-to-Implant (DTI) for both cohorts: 226 (90.8%) of TAS breasts, and 50 (90.91%) of CHG breasts. Most implants were of the brand MENTOR (Mentor Worldwide, city, state): 242 (97.2%) of TAS cohort implants and 55 (100%) of CHG cohort implants. The majority of the implant type was silicone for both cohorts: 239 (95.9%) implants in the TAS group and 55 (100%) implants in the CHG group. Most of the implants' texture was smooth: 247 (99.2%) implants in the TAS cohort and 55 (100%) implants in the CHG cohort. Additionally, most implants in both cohorts were placed in the pre-pectoral plane: 217 (87.2%) implants in the TAS group and 52 (94.6%) implants in the CHG group ($p=0.09$). ADM was used at a significantly higher rate in the TAS group compared to CHG group: 217 (87.1%) vs. 15 (27.2%), respectively ($p=0.01$). The

Table 1: Patient characteristic and demographic information of implant-based reconstruction. Bold p-values indicate statistical significance ($p < 0.05$).

Patient Characteristics	Antimicrobial Solution			p value
	Total Population	Triple Antibiotic Solution	0.05% Chlorhexidine-Gluconate Solution	
	n = 171	n = 141 (82.5)	n = 30 (17.5)	
Race				0.004
African American	45 (26.3%)	37 (26.2%)	8 (26.7%)	
White	94 (54.9%)	83 (58.9%)	11 (36.7%)	
Hispanic	1 (0.58%)	1 (0.1%)	0 (0%)	
Asian	7 (4.09%)	6 (0.4%)	1 (3.3%)	
Other	24 (14%)	14 (9.93%)	10 (33%)	
Age, years				
(median (IQR))	50 (18)	51 (9)	49 (12)	0.73
Body Mass Index				
(median (IQR))	25.27 (7.22)	25.1 (7.19)	25.93 (7.51)	0.86
Charlson Comorbidity Index				
(median (IQR))	2 (1)	2 (1)	2 (1)	0.83
Diabetes Mellitus	0 (0%)	0 (0%)	0 (0%)	0.65
Metastatic malignancy	8 (4.68%)	8 (5.67%)	0 (0%)	0.234
Congestive Heart Failure	5 (2.92%)	4 (2.84%)	1 (3.33%)	0.624
Chronic Obstructive Pulmonary Disease	2 (1.17%)	2 (1.42%)	0 (0%)	0.679
Cerebrovascular Accident	2 (1.17%)	2 (1.42%)	0 (0%)	0.679
Dementia	2 (1.17%)	2 (1.42%)	0 (0%)	0.679
Connective Tissue Disease	7 (4.09%)	6 (4.26%)	1 (3.33%)	0.645
Pneumonia <30 days before Surgery	1 (0.58%)	1 (0.71%)	0 (0)	0.825
Hypertension	36 (21.05%)	31 (22%)	5 (16.7%)	0.354
Dyslipidemia	39 (22.81%)	32 (22.7%)	7 (23.3%)	0.554
End Stage Renal Disease	3 (1.75%)	1 (0.71)	2 (6.67)	0.08
Obesity	39 (22.81%)	33 (23.4%)	6 (20%)	0.687
Marijuana use	7 (4.09%)	6 (4.26%)	1 (3.33%)	0.645
Former Smoker	30 (17.54%)	25 (17.73%)	5 (16.67%)	0.564
Current Smoker	9 (5.26%)	5 (3.55%)	4 (13.33%)	0.052
>2 Alcohol drinks/day	4 (2.34%)	4 (2.84%)	0 (0%)	0.46
Active Oral Steroid use	14 (8.19%)	12 (8.51%)	2 (6.67%)	1
Psychiatric Diagnosis	60 (35.09%)	52 (36.88%)	8 (26.67%)	0.19
Major Depressive Disorder	35 (20.47%)	31 (21.9)	4 (13.3)	0.21
Generalized Anxiety Disorder	41 (23.98%)	35 (24.8)	6 (20)	0.38
Active Psychiatric Medication	48 (28.07%)	43 (30.5)	5 (16.7)	0.09

majority of IBR used Dermacell (Stryker): 154 (61.9%) of breasts in the TAS cohort compared to 10 (18.2%) of breasts in the CHG cohort.

Implant outcomes and complications

Incidence of ≤ 30 -day complications are summarized in Table 4. Between groups, there was no significant difference in ≤ 30 -day seroma formation [1 (0.4%) vs. 0 (0%), $p=0.82$], hematoma formation [8 (3.2%) vs. 0 (0%), $p=0.2$], dehiscence [5 (2%) vs. 0 (0%)], delayed healing [13 (5.2%) vs. 2 (3.6%), $p=0.47$], mastectomy flap necrosis [13 (5.2%) vs. 2 (3.6%), $p=0.47$], cellulitis [7 (2.8%) vs. 0 (0%), $p=0.24$], surgical site infection [2 (0.8%) vs. 0 (0%), $p=0.67$], or unplanned return to the operating room [16 (6.4%) vs. 2 (3.6%), $p=0.34$], respectively. Table 4 also displays incidence of ≥ 60 -day complications. Likewise,

there was no significant difference in ≥ 60 -day seroma formation [4 (1.6%) vs. 0 (0%), $p=0.45$], hematoma formation [1 (0.4%) vs. 0 (0%), $p=0.82$], dehiscence [9 (3.6%) vs. 1 (1.8%), $p=0.43$], mastectomy flap necrosis [4 (1.6%) vs. 0 (0%), $p=0.45$], cellulitis [4 (1.6%) vs. 0 (0%), $p=0.447$], surgical site infection [6 (2.4%) vs. 1 (1.8%), $p=0.63$], unplanned implant removal [20 (8.1%) vs. 1 (1.8%), $p=0.08$], capsular contracture [5 (2%) vs. 2 (3.6%), $p=0.37$], respectively. There were no cases of red breast syndrome or breast implant-associated anaplastic large-cell lymphoma at any time point. Time to follow up was similar between groups: TAS patients' follow up was 31 days (IQR 15) and CHG patients' follow up was 32 days (IQR 14), ($p=0.896$).

Results of univariate regression analysis of all collected variables on ≤ 30 -day postoperative complications, ≥ 60 -day postoperative

Table 2: Surgical and oncologic history of implant-based reconstruction, by breast. Bold p-values indicate statistical significance (p<0.05).

Surgical and Oncologic History	Antimicrobial Solution			p value
	Total Breasts (n = 304)	Triple Antibiotic Solution (n = 249)	0.05% Chlorhexidine-Gluconate Solution (n = 55)	
Surgical History				0.022
Breast Reduction	2 (3.13%)	2 (0.8%)	0 (0%)	
Breast Recon	2 (3.13%)	27 (10.84%)	4 (7.27%)	
Lumpectomy	11 (17.19%)	9 (3.61%)	2 (3.62%)	
Excisional Biopsy	18 (28.13%)	11 (4.42%)	7 (12.73%)	
Chemo				0.001
Adjuvant	97 (33.33%)	91 (38%)	6 (11.54%)	
Neoadjuvant	38 (13.06%)	31 (12.97%)	7 (13.46%)	
Both	19 (6.53%)	17 (7.11%)	2 (3.85%)	
XRT				0.09
Neoadjuvant	9 (3.09%)	8 (3.35%)	1 (1.92%)	
Adjuvant	68 (23.37%)	62 (25.94%)	6 (11.54%)	
Both	1 (0.34%)	1 (0.42%)	0 (0%)	

Table 3: Operative details of implant-based reconstructions, by breast. Bold p-values indicate statistical significance (p<0.05).

Operative Details	Antimicrobial Solution			p value
	Total Breasts (n = 304)	Triple Antibiotic Solution (n = 249)	0.05% Chlorhexidine-Gluconate Solution (n = 55)	
Initial breast surgery				0.001
Nipple Sparing Mastectomy	200 (65.79%)	156 (62.7%)	44 (80%)	
Skin Sparing Mastectomy	84 (27.63%)	78 (31.3%)	6 (10.91%)	
Simple Mastectomy	10 (3.29%)	5 (2.02%)	5 (9.09%)	
Modified Radical Mastectomy	2 (0.66%)	2 (0.81%)	0 (0%)	
Other	8 (2.63%)	8 (3.24%)	0 (0%)	
Incision type				0.19
Inframamry fold	253 (83.22%)	202 (81.12%)	51 (92.73%)	
Elliptical	15 (4.93%)	15 (6.02%)	0 (0%)	
Wise Pattern	28 (9.21%)	24 (9.64%)	4 (7.27%)	
Other	7 (2.3%)	7 (2.81%)	0 (0%)	
Circumareolar	1 (0.33%)	1 (0.4%)	0 (0%)	
Sentinel Lymph Node Biopsy	112 (34.54%)	88 (35.3%)	17 (30.91%)	0.48
Axillary Lymph Node Dissection	20 (6.57%)	24 (9.64%)	3 (5.45%)	0.48
Drains Usage	304 (100%)	249 (100%)	55 (100%)	0.81
Operation duration (minutes)	192 (105.5)	205 (100)	157 (170)	0.002
Direct to implant	276 (90.79%)	226 (90.76%)	50 (90.91%)	0.605
Implant brand				1
Allergen	3 (0.99%)	3 (1.2%)	0 (0%)	
Mentor	297 (97.7%)	242 (97.2%)	55 (100%)	
Sientra	2 (0.66%)	2 (0.8%)	0 (0%)	
Other	2 (0.66%)	2 (0.8%)	0 (0%)	
Implant type				0.13
Silicone	294 (96.71%)	239 (95.98%)	55 (100%)	
Saline	10 (3.29%)	10 (4.02%)	0 (0%)	
Implant texture				0.67
Smooth	302 (99.34%)	247 (99.2%)	55 (100%)	
Textured	2 (0.66%)	2 (0.8%)	0 (0%)	

Implant Placement				0.09
Pre-pectoral	269 (88.49%)	217 (87.15%)	52 (94.55%)	
Retro-pectoral	35 (11.51%)	32 (12.85%)	3 (5.45%)	
Acellular Dermal Matrix				0.01
Surgimed	60 (19.74%)	60 (24.1%)	0 (0%)	
Alloderm	8 (2.63%)	3 (1.2%)	5 (9.09%)	
Dermacell	164 (53.95%)	154 (61.54%)	10 (18.18%)	

Table 4: 30-day and 60-day postoperative complications, by breast. Bold p-values indicate statistical significance ($p < 0.05$).

Complication Details	Antimicrobial Solution			p value
	Total Breasts	Triple Antibiotic Solution	0.05% Chlorhexidine-Gluconate Solution	
<30 days post op complication	51 (16.78%)	46 (18.47%)	5 (9.09%)	0.09
Seroma	1 (0.33%)	1 (0.4%)	0 (0%)	0.82
Hematoma	8 (2.63%)	8 (3.21%)	0 (0%)	0.2
Dehiscence	5 (1.64%)	5 (2.01%)	0 (0%)	0.37
Delayed healing	14 (4.61%)	13 (5.2%)	2 (3.64%)	0.47
Mastectomy flap necrosis	15 (4.93%)	13 (5.22%)	2 (3.64%)	0.47
Red breast syndrome	0 (0%)	0 (0%)	0 (0%)	0
Cellulitis	7 (2.3%)	7 (2.81%)	0 (0%)	0.244
Surgical Site Infection	2 (0.66%)	2 (0.8%)	0 (0%)	0.67
Unplanned Return to OR	18 (5.92%)	16 (6.43%)	2 (3.64%)	0.335
>60 day postop complication	32 (10.56%)	29 (11.69%)	3 (5.45%)	0.13
Seroma	4 (1.32%)	4 (1.61%)	0 (0%)	0.447
Hematoma	1 (0.33%)	1 (0.4%)	0 (0%)	0.818
Dehiscence	10 (3.3%)	9 (3.63%)	1 (1.82%)	0.431
Mastectomy flap necrosis	4 (1.32%)	4 (1.61%)	0 (0%)	0.447
Red breast syndrome	0 (0%)	0 (0%)	0 (0%)	0
Cellulitis	4 (1.32%)	4 (1.61%)	0 (0%)	0.447
Surgical Site Infection	7 (2.31%)	6 (2.42%)	1 (1.82%)	0.63
Unplanned implant removal	21 (6.93%)	20 (8.06%)	1 (1.82%)	0.078
Capsular contracture	7 (2.31%)	5 (2.02%)	2 (3.64%)	0.073
Degree of capsular contracture (mean (stdev))	1.86 (± 1.07)	1.4 (± 0.89)	3 (± 0)	0.062
Breast Implant Associated ALCL	0 (0%)	0 (0%)	0 (0%)	0
Cancer Recurrence	6 (1.98%)	6 (2.42%)	0 (0%)	0.3
Days to follow up (median (IQR))	31 (15)	31 (15)	32 (14)	0.896

complications, and capsular contracture is displayed supplemental Table 1. Operative duration was a significant co-variate to ≤ 30 day and ≥ 60 -day postoperative complications. Simple mastectomy was a significant co-variate to ≥ 60 -day postoperative complication and capsular contracture. Type of irrigation solution was not a significant variable for either < 30 day or > 60 -day complications. Results of multivariate regression analysis is displayed in supplemental table 2. ADM use was not a significant variable for < 30 -day complications, > 60 -day complications, or capsular contracture.

Discussion

In our study of 304 breasts undergoing IBR for oncologic purposes, we determined that there were no significant differences between TAS and CHG in terms of post-operative complications or rate of infection. This study contributes to the ongoing discussion regarding implant irrigation solution. The use of antimicrobial

irrigation solutions is ubiquitous throughout surgical specialties and their effectiveness is well reported [13-15]. Specific to breast reconstruction, antimicrobial implant soaking agents and breast pocket irrigation are methods used to prevent infection, capsular contracture, and return to the operating room, amongst other postoperative complications. Both TAS and CHG solutions have been shown to be effective antimicrobial irrigation agents in various implant-based surgeries but have not been compared in the setting of IBR [13,16].

With an incidence of 10.6%, capsular contracture leads to excessive pain and deformity and possible need for revision [3] Adams et al. demonstrated that the use of triple-antibiotic solution reduced incidence of grade III/IV capsular contracture in patients receiving implants for aesthetic and reconstructive indications [17]. In our study, we found relatively low rates of this complication (total 2.31%; TAS

Supplementary Table 1:

	Supplement Table 1: Univariate Regression Analysis		
	≤30 Day Complications	≥60 Day Complications	Capsular Contracture
Age	0.012 (0.527)	0.011 (0.634)	0.012 (0.796)
Diabetes Mellitus	0.513 (0.369)	0.979 (0.326)	-
Body Mass Index	0.029 (0.251)	0.011 (0.76)	0.125 (0.191)
Hyperlipidemia	0.167 (0.7)	0.514 (0.396)	0.79 (0.515)
Surgical History			
Breast Reduction	-	-	-
Breast Recon	-	-	-
Lumpectomy	-	-	-
Excisional Biopsy	1.215 (0.315)	-	-
Chemotherapy			
Adjuvant	0.517 (0.585)	1.21 (0.447)	-
Neoadjuvant	1.172 (0.427)	-	-
Both	-	-	-
Radiation Therapy			
Neoadjuvant	-	-	-
Adjuvant	0.601 (0.556)	1.21 (0.447)	-
Both	-	-	-
Direct to Implant	0.066 (0.927)	-	-
Initial breast surgery			
NSM	-	-	-
SSM	0.001 (0.998)	1.506 (0.005)	0.062 (0.959)
Simple Mastectomy	0.343 (0.737)	2.487 (0.022)	3.832 (0.046)
MRM	-	-	-
Other	-	-	-
Incision type			
Inframammary fold	-	1.172 (0.142)	-
Elliptical	1.236 (0.309)	-	-
Wise Pattern	-	0.76 (0.216)	-
Other	0.63 (0.222)	-	2.354 (0.074)
Circumareolar	0.294 (0.796)	-	-
Axillary Lymph Node Dissection	0.637 (0.355)	0.112 (0.894)	1.304 (0.381)
Operative duration	0.005 (0.022)	0.007 (0.01)	0.002 (0.768)
Implant Brand			
Allergen	-	-	-
Mentor	2.072 (0.267)	0.12 (0.941)	-
Sientra	-	-	-
Other	-	-	-
Implant Type	0.99 (0.44)	1.458 (0.159)	-
Implant Placement	0.837 (0.097)	0.099 (0.889)	-
Acellular Dermal Matrix			
Surgimed	0.471 (0.442)	0.955 (0.326)	0.537 (0.783)
Alloderm	1.548 (0.114)	2.19 (0.106)	-
DermaCell	0.685 (0.186)	0.875 (0.344)	1.223 (0.483)
Antimicrobial solution	0.716 (0.159)	0.713 (0.267)	1.054 (0.255)

Significance set at p<0.05. Categorical variables listed as Odds Ratio (p-value) and continuous variables listed as Coefficient (p-value). Statistical analysis completed using linear logistic or linear regression, as appropriate.

Supplementary Table 2:

	Supplemental Table 2: Confounding Variable Multivariate Regression Analysis		
	<30 Day Complications	>60 Day Complications	Capsular Contracture
Initial breast surgery			
Skin Sparing Mastectomy	1.164 (0.663)	5.479 (0.001)	1.569 (0.634)
Simple Mastectomy	1.876 (0.461)	12.876 (0.002)	24.775 (0.003)
Operative duration	1.005 (0.024)	0.005 (0.021)	1.006 (0.164)
Implant Placement	1.741 (0.296)	0.555 (0.334)	-
Acellular Dermal Matrix Use	1.059 (0.252)	0.057 (0.753)	1.177 (0.654)
Antimicrobial Solution	0.650 (0.394)	0.430 (0.556)	1.715 (0.617)
Significance set at p<0.05. Categorical variables listed as Odds Ratio (p-value) and continuous variables listed as Coefficient (p-value). Statistical analysis completed using multivariate logistic or linear regression, as appropriate.			

2.02%; CHG 3.64%). Studies report a multitude of factors contribute to this complication, including concurrent implant infection, prior radiation or chemotherapy, and patient demographics such as age and comorbidities [18,19]. TAS has been previously shown to reduce incidence of capsular contracture three-fold in IBR [17]. However, in our current study we demonstrate similar outcomes between TAS and CHG use in regard to seroma, hematoma, dehiscence, surgical site infection, and capsular contracture. With the use of both solutions, we achieved very low complication rates overall.

In addition to antimicrobial solutions used, other factors may have a more important effect on the incidence of short and long-term postoperative complications. Amongst both TAS and CHG cohorts, the most common incision performed was along the inframammary fold. Literature has demonstrated that the inframammary fold offers a stable anatomical location for incision placement that minimizes the incidence of trauma and inflammation, factors known to contribute to capsular contracture [20,21]. In addition, the vast majority of IBR performed in both TAS and CHG cohorts had implants placed in the pre-pectoral space. Placement of implants in the pre-pectoral space has been shown to have comparable rates of capsular contracture, prosthesis failure, and animation deformity when compared to subpectoral IBR [10,22].

In general, ADMs are used to decrease implant exposure, decrease rippling, and may enhance aesthetic results [23]. Additionally, ADMs are thought to alter the reactive processes at the tissue-implant interface, potentially reducing rates or delaying onset of contracture [3]. However, ADMs usage may lead to increased risk of seroma formation, infection, or red breast syndrome [23,24]. While the usage of ADM was significantly higher in the TAS cohort, there were no significant differences in notable postoperative outcomes. Given that the literature presents conflicting views regarding the incidence of infection with ADM while simultaneously highlighting its efficacy in reducing capsular contracture, its usage alone may not be sufficient in being associated with complications in our cohort [25,26]. Furthermore, our regression analysis did not yield any significance in ADM usage to short or long-term postoperative outcomes.

Patients that are female, obese, have preexisting diabetes mellitus, smoking, and history neoadjuvant chemotherapy or radiation, and prior breast therapy have been associated with increased infection rates [27-29]. Despite a balanced cohort with no significant difference in incidence of diabetes, smoking, obesity, in TAS or CHG patients, significantly more TAS patients underwent both neoadjuvant and adjuvant chemotherapy or just adjuvant chemotherapy compared to CHG patients. Furthermore, there were significantly more patients

who underwent radiation (neoadjuvant, adjuvant, or both) therapy in the TAS cohort. These differences may contribute to the higher rates of postoperative complications found in the TAS cohort, although the difference in complication rate was not significant.

While implant type, whether silicone or saline, has not been significantly associated with infection, longer operation duration, lymph node dissection, and direct usage of implants during reconstruction have been shown to have a positive association [30]. Though the difference was not statistically significant, the median duration of surgery was 205 mins in the TSA group and 157 mins for the CHG group. Regardless, longer surgical times have been linked to increased risk of infection due to prolonged tissue exposure and anesthetic use, which could partially explain the higher, yet non-significant, postoperative complication rates in the TAS cohort [31]. On our regression analysis, we found that operative duration was a significant co-variate to both short and long-term postoperative complications, supporting that operative duration may have played a role in incidence of complications. Moreover, literature supports the association between lymph node dissection and seroma formation and delayed wound healing [32,33]. Patients in the TAS cohort did have more axillary lymph node dissections than the CHG cohort which may have also contributed to the TAS cohort's higher postoperative complication rates of seroma and dehiscence. However, there was no statistically significant difference in rates of axillary lymph node dissection nor short- and long- term seroma or dehiscence rates. While a favored method for immediate aesthetic outcomes, DTI has also been linked to higher rates of infection and capsular contracture in part due to the absence of a tissue expander phase [34]. While DTI reconstruction was extremely common in both cohorts (90.8% TAS, 90.91% CHG), the low rates of infection in both groups may suggest that both irrigation solutions offer effective and sufficient prophylaxis in the context of DTI.

Expanding from a medical perspective, the cost considerations of irrigation solution choice should also be considered given ambiguous surgical advantage. CHG tends to be more expensive than triple-antibiotic solutions. Research from other institutions has shown triple antibiotic solution to cost \$20.43 per Bacitracin (50,000U) + cefazolin (1g) + gentamicin (80mg) mixed with 1-L normal saline, while chlorhexidine costs \$35.95 for a 950 mL solution [35]. A separate consideration is that CHG does not require mixing solutions together and is thus easier to use. Surgical staff unfamiliar with mixing the triple antibiotic solution may lead to a time consuming and confusing process to achieve an appropriate irrigation solution. This barrier to use can be exacerbated if a surgeon does not work with the

same dedicated surgical staff or operates at multiple hospitals. Given similar prevention in complication rates, cost and efficiency may be given more weight when considering which irrigation solution to use in practice.

Adverse side effects of the solutions are uncommon yet play an important role in decision-making when providing optimal patient care. In February of 2017, the United States Food and Drug Administration (FDA) issued a warning regarding a rare allergic reaction to Chlorhexidine-containing skin aseptic products [36]. Specifically, the FDA had identified 52 cases of anaphylaxis with use of CHG products applied to the skin, with 43 cases reported worldwide between January 1969 and June 2015 [36]. Given the potentially severe and devastating allergic reactions to CHG or antibiotics in TAS, caution must continue to be taken to minimize preventable adverse reactions when either irrigation solution is used. In our study, there was no incidence of allergic reactions to either TAS or CHG.

Limitations

The nature of this study design being a retrospective multi-institution study is an inherent limitation. There may be inconsistencies in documentation or patient follow-up in the electronic medical record which may influence data collection and subsequent analysis. Additionally, the sample size of patients in the CHG cohort was smaller compared to the TAS cohort which may amplify the detection of significant difference between cohorts. However, our study found no significant difference in our primary outcomes between both cohorts.

Conclusion

Routine breast pocket irrigation and implant soaking continues to be used as effective infection prophylaxis in implant-based breast reconstructive surgery. We found that there was no significant difference between use of triple antibiotic solution and chlorhexidine-based solution in rates of short- and long-term postoperative infection, capsular contracture, and return to the operating room.

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References

- Adams WP, Jr., Mallucci P. Breast augmentation. *Plast Reconstr Surg.* 2012;130:597e-611e.
- Ooi A, Song DH. Reducing infection risk in implant-based breast-reconstruction surgery: challenges and solutions. *Breast Cancer (Dove Med Press).* 2016;8:161-72.
- Headon H, Kasem A, Mokbel K. Capsular Contracture after Breast Augmentation: An Update for Clinical Practice. *Arch Plast Surg.* 2015;42:532-43.
- Gabriel A, Sigalove S, Sigalove NM, et al. Prepectoral Revision Breast Reconstruction for Treatment of Implant-Associated Animation Deformity: A Review of 102 Reconstructions. *Aesthet Surg J* 2018;38(5):519-26.
- Mesa F, Catano S, Tuberquia O. Study of Infections in Breast Augmentation Surgery with Implants in 9,691 Patients over 5 Years. *Plast Reconstr Surg Glob Open.* 2021;9:e3752.
- Alderman AK, Wilkins EG, Kim HM, Lowery JC. Complications in postmastectomy breast reconstruction: two-year results of the Michigan Breast Reconstruction Outcome Study. *Plast Reconstr Surg.* 2002;109:2265-74.
- Hvilsom GB, Holmich LR, Henriksen TF, Lipworth L, McLaughlin JK, Friis S. Local complications after cosmetic breast augmentation: results from the Danish Registry for Plastic Surgery of the Breast. *Plast Surg Nurs.* 2010;30:172-9.
- Olsen MA, Nickel KB, Fox IK, Margenthaler JA, Ball KE, Mines D, et al. Incidence of Surgical Site Infection Following Mastectomy With and Without Immediate Reconstruction Using Private Insurer Claims Data. *Infect Control Hosp Epidemiol.* 2015;36:907-14.
- Adams Jr WP, Culbertson EJ, Deva AK, Magnusson MR, Layt C, Jewell ML, et al. Macrot textured Breast Implants with Defined Steps to Minimize Bacterial Contamination around the Device: Experience in 42,000 Implants. *Plast Reconstr Surg.* 2017;140:427-31.
- Federica G, Tommaso F, Alessia C, Agostino C, Florian B, Antonio G, et al. Use of Antimicrobial Irrigation and Incidence of Capsular Contracture in Breast Augmentation and Immediate Implant-Based Breast Reconstruction. *Aesthetic Plast Surg.* 2023;47:2345-50.
- Papadakis M. Wound irrigation for preventing surgical site infections. *World J Methodol.* 2021;11:222-7.
- Epps MT, Langsdon S, Pels TK, Lee TM, Thurston T, Brzeziński MA. Antimicrobial Irrigation and Technique during Breast Augmentation: Survey of Current Practice. *Plast Reconstr Surg Glob Open.* 2019;7:e2310.
- Hemmingsen MN, Bennedsen AK, Kullab RB, Weltz TK, Larsen A, Ørholt M, et al. Antibiotic Implant Irrigation and Deep Infection: A Retrospective Study of 1508 Patients Undergoing Breast Reconstruction with Implants. *Plast Reconstr Surg.* 2024;154:5-13.
- Li J, Wang N, Zhang J. Case Report: Antibiotic Irrigation and Drainage Tube for Managing Chronic Suppurative Otitis Media After Cochlear Implantation. *Ear Nose Throat J.* 2024;1455613241238829.
- Yildiz AK, Bayraktar A, Kacan T, Demir DO, Gokkurt Y, Keseroglu BB, et al. A new protocol for renal collecting system sterilization with antibiotic irrigation during lithotripsy in retrograde intrarenal surgery: a prospective, comparative study. *World J Urol.* 2024;42:229.
- Namnoon JD, Largent J, Kaplan HM, Oefelein MG, Brown MH. Primary breast augmentation clinical trial outcomes stratified by surgical incision, anatomical placement and implant device type. *J Plast Reconstr Aesthet Surg.* 2013;66:1165-72.
- Adams Jr WP, Rios JL, Smith SJ. Enhancing patient outcomes in aesthetic and reconstructive breast surgery using triple antibiotic breast irrigation: six-year prospective clinical study. *Plast Reconstr Surg.* 2006;118:46S-52S.
- Bachour Y, Ritt M. Risk factors for developing capsular contracture in women after breast implant surgery: A systematic review of the literature. *J Plast Reconstr Aesthet Surg.* 2018;71:e68.
- Dancey A, Nassimizadeh A, Levick P. Capsular contracture - What are the risk factors? A 14 year series of 1400 consecutive augmentations. *J Plast Reconstr Aesthet Surg.* 2012;65:213-8.
- Jacobson JM, Gatti ME, Schaffner AD, Hill LM, Spear SL. Effect of incision choice on outcomes in primary breast augmentation. *Aesthet Surg J.* 2012;32:456-62.
- Stevens WG, Nahabedian MY, Calobrace MB, Harrington JL, Capizzi PJ, Cohen R, et al. Risk factor analysis for capsular contracture: a 5-year Sientra study analysis using round, smooth, and textured implants for breast augmentation. *Plast Reconstr Surg.* 2013;132:1115-23.
- Escandón JM, Weiss A, Christiano JG, Langstein HN, Escandón L, Prieto PA, et al. Prepectoral versus subpectoral two-stage implant-based breast reconstruction: U.S. medical center experience and narrative review. *Ann Transl Med* 2023;11:411.
- Podsednik Gardner A, Nunez A, De la Garza M. Red Breast Syndrome and

- Acellular Dermal Matrix. *Plast Reconstr Surg Glob Open* . 2023;11:e5062.
24. Johnson AC, Colakoglu S, Siddikoglu D, Li A, Kaoutzanis C, Cohen JB, et al. Impact of Dermal Matrix Brand in Implant-Based Breast Reconstruction Outcomes. *Plast Reconstr Surg*. 2022;150:17-25.
 25. Samuels K, Millet E, Wong L. Efficacy of Acellular Dermal Matrix Type in Treatment of Capsular Contracture in Breast Augmentation: A Systematic Review and Meta-Analysis. *Aesthet Surg J*. 2023;44:26-35.
 26. Phillips BT, Bishawi M, Dagum AB, Bui DT, Khan SU. A systematic review of infection rates and associated antibiotic duration in acellular dermal matrix breast reconstruction. *Eplasty*. 2014;14:e42.
 27. Francis SH, Ruberg RL, Stevenson KB, Beck CE, Ruppert AS, Harper JT, et al. Independent risk factors for infection in tissue expander breast reconstruction. *Plast Reconstr Surg*. 2009;124:1790-6.
 28. Peled AW, Itakura K, Foster RD, Hamolsky D, Tanaka J, Ewing C, et al. Impact of chemotherapy on postoperative complications after mastectomy and immediate breast reconstruction. *Arch Surg* 2010;145:880-5.
 29. Kaoutzanis C, Gupta V, Winocour J, Shack B, Grotting JC, Higdon K. Incidence and Risk Factors for Major Surgical Site Infections in Aesthetic Surgery: Analysis of 129,007 Patients. *Aesthet Surg J* 2017;37:89-99.
 30. Basile AR, Basile F, Basile AV. Late infection following breast augmentation with textured silicone gel-filled implants. *Aesthet Surg J*. 2005;25:249-54.
 31. Ravi B, Jenkinson R, O' Heireamhoin S, Austin PC, Aktar S, Leroux TS, et al. Surgical duration is associated with an increased risk of periprosthetic infection following total knee arthroplasty: A population-based retrospective cohort study. *E Clin Med*. 2019;16:74-80.
 32. Nagayama A, Kitagawa Y. Managing Seroma Formation Post Breast Surgery through Somatostatin Analogs. *JMA J*. 2023;6:282-3.
 33. Nzenwa IC, Iqbal HA, Hardie C, Smith GE, Matteucci PL, Totty JP. Wound complications following surgery to the lymph nodes: A protocol for a systematic review and meta-analysis. *PLoS One* 2022;17:e0272490.
 34. Jeon HB, Lee M, Roh TS, Jeong J, Ahn SG, Bae SJ, et al. Complications Including Capsular Contracture in Direct-to-Implant Breast Reconstruction With Textured Anatomical Versus Smooth Round Implants: A Single Center Retrospective Analysis. *J Breast Cancer*. 2023;26:25-34.
 35. Bamba R, Tran PC, Mailey BA, Lin J, DeBrock W, Dawson S, et al. Comparison of Breast Reconstruction Outcomes Using Oxychlorosene versus Triple Antibiotic Solution for Pocket Irrigation. *Plast Reconstr Surg Glob Open*. 2022;10:e3975.
 36. FDA. Chlorhexidine gluconate: Drug safety communication - Rare but serious allergic reactions. 2017.