

Impact of Systemic Antibiotics on *Staphylococcus aureus* Colonization and Recurrent Skin Infection

Patrick G. Hogan,¹ Marcela Rodriguez,² Allison M. Spenner,² Jennifer M. Brenneisen,² Mary G. Boyle,¹ Melanie L. Sullivan,¹ and Stephanie A. Fritz¹ ¹Department of Pediatrics, Washington University School of Medicine, St Louis, Missouri; and ²Department of Pediatrics, Southern Illinois University School of Medicine, Springfield

Background. Staphylococcus aureus colonization poses risk for subsequent skin and soft tissue infection (SSTI). We hypothesized that including systemic antibiotics in the management of *S. aureus* SSTI, in conjunction with incision and drainage, would reduce *S. aureus* colonization and incidence of recurrent infection.

Methods. We prospectively evaluated 383 children with *S. aureus* SSTI requiring incision and drainage and *S. aureus* colonization in the anterior nares, axillae, or inguinal folds at baseline screening. Systemic antibiotic prescribing at the point of care was recorded. Repeat colonization sampling was performed within 3 months (median, 38 days; interquartile range, 22–50 days) in 357 participants. Incidence of recurrent infection was ascertained for up to 1 year.

Results. Participants prescribed guideline-recommended empiric antibiotics for purulent SSTI were less likely to remain colonized at follow-up sampling (adjusted hazard ratio [aHR], 0.49; 95% confidence interval [CI], .30–.79) and less likely to have recurrent SSTI (aHR, 0.57; 95% CI, .34–.94) than those not receiving guideline-recommended empiric antibiotics for their SSTI. Additionally, participants remaining colonized at repeat sampling were more likely to report a recurrent infection over 12 months (aHR, 2.37; 95% CI, 1.69–3.31). Clindamycin was more effective than trimethoprim-sulfamethoxazole (TMP-SMX) in eradicating *S. aureus* colonization (44% vs 57% remained colonized, P = .03) and preventing recurrent SSTI (31% vs 47% experienced recurrence, P = .008).

Conclusions. Systemic antibiotics, as part of acute SSTI management, impact *S. aureus* colonization, contributing to a decreased incidence of recurrent SSTI. The mechanism by which clindamycin differentially affects colonization and recurrent SSTI compared to TMP-SMX warrants further study.

Keywords. Staphylococcus aureus; systemic antibiotics; colonization; SSTI.

Skin and soft tissue infections (SSTI) rank as one of the top causes of pediatric hospitalization and account for >14 million outpatient visits each year in the United States [1–3]. Rates of pediatric hospitalization for SSTI increased by 36% between 2000 and 2012, more than for any other diagnosis [2]. *Staphylococcus aureus* is the most common cause of SSTI [4, 5]. The incidence of *S. aureus* SSTI has increased substantially since the late 1990s, largely driven by the emergence of the USA300 clone [6, 7]. Recurrence of *S. aureus* SSTI is common and may occur in >50% of pediatric patients [8–11]. *Staphylococcus aureus* at multiple sites (ie, higher burden of colonization) has been associated with higher recurrence rates [15].

Currently, there is equipoise surrounding the utilization of antibiotics in management of *S. aureus* skin abscesses. Incision

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and drainage (I&D) has been considered the mainstay of treatment for uncomplicated S. aureus skin abscesses, consistent with guidelines from the Infectious Diseases Society of America (IDSA) [16]. For patients with purulent cellulitis and skin abscesses for whom antibiotics are prescribed in the outpatient setting, the guidelines recommend empiric treatment with non-β-lactam antibiotics, including clindamycin, trimethoprim-sulfamethoxazole (TMP-SMX), a tetracycline, or linezolid [17]. Several recent studies investigating the effects of antibiotic administration on SSTI clinical cure rates have demonstrated both positive [18, 19] and negligible [8, 20, 21] effects. Other recent studies have demonstrated that patients who do not receive systemic antibiotics for SSTI, or those who do not receive an antibiotic active against the infecting organism (eg, methicillin-resistant S. aureus [MRSA]), are more likely to suffer recurrent infections [8, 21]. A recent trial of 786 patients with skin abscess randomized participants to receive clindamycin, TMP-SMX, or placebo after I&D for 10 days [22]. Those receiving antibiotics (either clindamycin or TMP-SMX) had a higher cure rate and reduced skin infections at new sites 1 month following treatment.

It is unknown whether systemic antibiotics decolonize the nares and skin. Oral antibiotic agents generally do not achieve suitable concentrations in the nares to effectively decolonize this site [23]; the impact of systemic antibiotics on *S. aureus* at other

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Correspondence: S. A. Fritz, Department of Pediatrics, Washington University School of Medicine, 660 S Euclid Ave, CB 8116, St Louis, MO 63110 (fritz.s@wustl.edu).

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common sites of colonization, such as the axillae or inguinal folds, is largely undefined. However, multiple studies have evaluated oral antibiotic regimens as part of multipronged MRSA eradication strategies. In both Greece and the Netherlands, MRSA SSTI management algorithms that include systemic antibiotics have been successful in reducing carriage of MRSA [24, 25]. In hospitalized patients in Canada, a decolonization treatment bundle including chlorhexidine body washes, intranasal mupirocin, and oral rifampin and doxycycline, compared with no treatment, significantly reduced MRSA colonization at 3 and 8 months following treatment [26]. Other studies, however, have shown antibiotics to be variably effective decolonizing agents and linked with increased antimicrobial resistance [27-29]; thus, prescription of oral antibiotics as part of decolonization efforts (in the absence of acute SSTI) has not been encouraged by the IDSA [17].

In the present study, we sought to illuminate the link between oral antibiotic therapy and reduction in recurrent SSTI. We tested the hypothesis that inclusion of guideline-recommended oral antibiotics in the management of *S. aureus* skin abscesses in addition to I&D would result in eradication of *S. aureus* colonization as well as a decreased incidence of recurrent SSTI.

METHODS

Study Design and Participant Recruitment

The cohort for this study is an aggregate of participants <21 years old with community-onset S. aureus SSTI and S. aureus colonization whose data were collected prospectively from 2008-2016 during one of several studies [10, 11, 22, 30], in addition to nascent participants recruited for this study. Participants were recruited from St Louis, Missouri (n = 376; St Louis Children's Hospital and pediatric practices affiliated with the Washington University Pediatric and Adolescent Ambulatory Research Consortium) and Springfield, Illinois (n = 7; St John's Children's Hospital, Memorial Medical Center, and Southern Illinois University pediatric clinics). Across study populations, patient characteristics were similar. All participants presented with acute, community-onset SSTI for which an I&D procedure was performed. For participants receiving oral antibiotics, the class, dose, and duration of antibiotic therapy varied. Children with immunodeficiency, those hospitalized within the previous 14 days, or those who performed decolonization measures (with mupirocin ointment, chlorhexidine gluconate, or bleach baths) in the prior month were excluded. This study was approved by the institutional review boards at Washington University and Southern Illinois University. Informed consent was obtained for each participant.

Baseline Screening and Longitudinal Data and Sample Collection

Across all study populations, patients with SSTI were swabbed by study personnel for detection of *S. aureus* colonization in the anterior nares, axillae, and inguinal folds at time of presentation and I&D procedure (baseline screening). Those with S. aureus SSTI and S. *aureus* colonization (n = 383) were then enrolled in a longitudinal study. Three-hundred fifty-seven of these participants underwent a repeat colonization swab at a follow-up visit (median, 38 days; interquartile range, 22-50 days). Each set of colonization swabs was collected by study team personnel during in-person longitudinal study visits. At the time of collection of each set of colonization swabs, a detailed questionnaire was administered and medical records were reviewed to ascertain antibiotic use (class and duration) and decolonization with topical antimicrobials. Most participants were prescribed decolonization measures (eg, intranasal mupirocin, chlorhexidine body washes, and/or dilute bleach water baths; Table 1), as previously described [10, 11]. Subsequent follow-up surveys were completed in person and by mail or telephone (depending on the study) up to 5 times over 12 months to ascertain incidence of recurrent SSTI. Infections were confirmed by review of medical records when available.

Table 1. Participant Characteristics (N = 383)

| Characteristic | No. (%) |
|--|---------------------------|
| Age, y, median (range) | 3.0 (0.5–20.3) |
| Female sex | 216 (56) |
| Race | |
| White | 143 (37) |
| African American or biracial | 237 (62) |
| Asian | 3 (1) |
| Baseline infecting organism | |
| MRSA | 309 (81) |
| MSSA | 74 (19) |
| Baseline colonization status | |
| MRSA | 241 (63) |
| MSSA | 105 (27) |
| MRSA and MSSA ^a | 37 (10) |
| Anatomic site of baseline S. aureus colonization | |
| Nares only | 80 (21) |
| Axillae only | 16 (4) |
| Inguinal folds only | 108 (28) |
| Nares and axillae | 21 (5) |
| Nares and inguinal folds | 74 (19) |
| Axillae and inguinal folds | 19 (5) |
| Nares, axillae, and inguinal folds | 65 (17) |
| Decolonization measures prescribed over the longitudinal | study period ^b |
| None | 80 (21) |
| Intranasal mupirocin only | 40 (10) |
| Chlorhexidine body washes only | 1 (0.3) |
| Dilute bleach water baths only | 4 (1) |
| Intranasal mupirocin and chlorhexidine body washes | 223 (58) |
| Intranasal mupirocin and dilute bleach water baths | 35 (9) |

Abbreviations: MRSA, methicillin-resistant *Staphylococcus aureus*; MSSA, methicillinsusceptible *Staphylococcus aureus*.

^aAt different anatomic sites.

^bOne hundred thirteen of 357 (32%) participants with follow-up colonization swabs were prescribed decolonization measures between samplings.

Microbiology

Antibiotic susceptibility data for each infecting isolate was obtained from the medical record. Colonization swabs were inoculated to tryptic soy broth with 6.5% sodium chloride and incubated overnight at 35°C. Broth was plated to trypticase soy agar with 5% sheep's blood and incubated for 24-48 hours. Confirmatory testing for S. aureus (by colony morphology, catalase activity, latex agglutination, and Gram stain) and antibiotic susceptibility testing (by Kirby-Bauer disk diffusion, including testing for inducible clindamycin resistance) were performed in accordance with published guidelines [31].

Statistical Analysis

Data were analyzed with IBM SPSS for Windows version 23. The χ^2 test was performed to compare the proportion of participants colonized with S. aureus or reporting a recurrent infection at follow-up between those receiving antibiotics recommended for purulent SSTI at the point of care (ie, clindamycin, TMP-SMX, a tetracycline, or linezolid, per IDSA guidelines for empiric therapy of purulent SSTI [17]) and those not prescribed guideline-recommended antibiotics. Incidence of recurrent SSTI was also compared between participants remaining colonized at repeat sampling and those not remaining colonized. Cox proportional hazards regression analyses were used to determine whether receipt of guideline-recommended antibiotics at the time of index SSTI was independently associated with (1) remaining colonized with S. aureus at follow-up sampling and (2) incidence of recurrent SSTI for up to 1 year. Other covariates included in the regression analyses were age, race, methicillin susceptibility of the SSTI isolate (MRSA vs methicillin-susceptible S. aureus), prescription of decolonization measures for baseline SSTI, and burden (ie, number of anatomical sites) of S. aureus colonization at baseline. Staphylococcus aureus colonization status at follow-up sampling was also included in the recurrent infection regression analysis. All tests of significance were 2-tailed, and a *P* value of <.05 was considered significant.

RESULTS

The 383 participants with S. aureus SSTI and S. aureus colonization had a median age of 3.0 years (range, 0.5-20.3 years), and 56% were female. A majority (62% [n = 237]) of participants were African American or biracial, while 37% (n = 143) were white and 1% (n = 3) were Asian; 2% (n = 8) were of Hispanic or Latino origin. Of 383 participants colonized with S. aureus at the time of presentation with SSTI requiring I&D (ie, at baseline screening), 204 (53%) were colonized at 1 anatomic site, 114 (30%) at 2 sites, and 65 (17%) at all 3 sites (Table 1).

Of 383 participants with an S. aureus SSTI requiring I&D and S. aureus colonization, 355 (93%) were empirically prescribed a guideline-recommended antibiotic. Most participants received clindamycin (n = 220 [57%]) or TMP-SMX (n = 199 [52%]); 19 (5%) received vancomycin, 12 (3%) received a β-lactam (penicillin, amoxicillin, amoxicillin-clavulanate, piperacillin-tazobactam, ceftriaxone, cephalexin, or cefadroxil), and 27 (7%) received no systemic antibiotics. For 81 (21%) participants, the SSTI treatment course included >1 class of antibiotics (median, 1; range, 0-4). The antibiogram for all recovered S. aureus isolates is included in Table 2.

Of 383 participants with S. aureus SSTI and S. aureus colonization, follow-up colonization sampling was performed in 357; of these, 178 (50%) remained colonized at repeat sampling. Within this cohort, among 331 participants prescribed guideline-recommended empiric antibiotics at the point of care, 48% (n = 158) remained colonized with *S. aureus*, while 77% (n = 20)of 26 participants not prescribed guideline-recommended empiric antibiotics remained colonized (P = .004) (Figure 1).

| Type of <i>S. aureus</i> | No. of <i>Staphylococcus</i> aureus Isolates | % Susceptible | | | | | | | | | | |
|---------------------------------------|---|------------------|------------------|-----|---------|-----|-----|-----|-----|-----|-----|-----|
| | | MET ^a | CLI ^b | ERY | TMP-SMX | RIF | TET | CIP | LZD | CPT | MUP | VAN |
| Overall <i>S. aureus</i> ^c | 1301 | 29 | 85 | 17 | 100 | 100 | 97 | 55 | 100 | 100 | 100 | 100 |
| Infecting | 383 | 19 | 90 | 9 | 100 | 100 | 96 | 42 | 100 | 100 | 100 | 100 |
| Colonizing | 918 | 33 | 83 | 21 | 99 | 100 | 97 | 56 | 100 | 100 | 100 | 100 |
| MRSA | 925 | 0 | 85 | 6 | 99 | 100 | 97 | 44 | 100 | 100 | 100 | 100 |
| Infecting | 309 | 0 | 90 | 5 | 100 | 100 | 97 | 35 | 100 | 100 | 100 | 100 |
| Colonizing | 616 | 0 | 83 | 6 | 99 | 100 | 97 | 45 | 100 | 100 | 100 | 100 |
| MSSA | 376 | 100 | 85 | 46 | 100 | 100 | 97 | 79 | 100 | 100 | 100 | 100 |
| Infecting | 74 | 100 | 88 | 27 | 100 | 100 | 95 | 80 | 100 | 100 | 100 | 100 |
| Colonizing | 302 | 100 | 84 | 51 | 100 | 100 | 97 | 79 | 100 | 100 | 100 | 100 |

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Data may represent multiple S. aureus isolates recovered from the same participant.

Abbreviations: CIP, ciprofloxacin; CLI, clindamycin; CPT, ceftaroline; ERY, erythromycin; LZD, linezolid; MET, methicillin; MRSA, methicillin-resistant Staphylococcus aureus; MSSA, methicillin-susceptible Staphylococcus aureus; MUP, mupirocin; RIF, rifampin; TET, tetracycline; TMP-SMX, trimethoprim-sulfamethoxazole; VAN, vancomycin.

^aAs predicted by cefoxitin or oxacillin testing.

^bClindamycin-susceptible isolates exhibiting inducible clindamycin resistance (ie, D-test positive) were considered clindamycin resistant.

°lsolates underwent susceptibility testing to varying antimicrobials; therefore, % susceptible is out of <1301 for the following: CLI (n = 1283), ERY (n = 1282), TMP-SMX (n = 1246), RIF (n = 991), TET (n = 1006), CIP (n = 931), LZD (n = 947), CPT (n = 90), MUP (n = 64), VAN (n = 1158).

Of the 357 participants, 269 were prescribed either clindamycin (n = 142) or TMP-SMX (n = 127). Whereas 44% (n = 62) of those receiving clindamycin for their SSTI remained colonized with *S. aureus* at follow-up, 57% (n = 72) of those receiving TMP-SMX remained colonized (P = .03).

In a Cox proportional hazards regression analysis adjusting for age, race, methicillin susceptibility of the SSTI isolate, prescription of decolonization measures, and number of anatomic sites of baseline *S. aureus* colonization (ie, burden of colonization), participants prescribed guideline-recommended antibiotics for SSTI at the point of care were less likely to remain colonized at their second sampling (adjusted hazard ratio [aHR], 0.49; 95% CI, .30–.79) (Figure 2). Additionally, decolonization measures prescribed for baseline SSTI were independently associated with reduced prevalence of colonization at follow-up (aHR, 0.29; 95% CI, .20–.40). Older children were also less likely to remain colonized at their second sampling (each additional year of age associated with aHR of 0.97; 95% CI, .95–.99).

All 383 participants with *S. aureus* SSTI and *S. aureus* colonization were followed longitudinally (for up to 12 months) to ascertain incidence of recurrent SSTI. Of these, 161 (42%) reported a recurrent infection: 40% (143/355) of those prescribed guideline-recommended antibiotics for SSTI at the point of care reported a recurrent SSTI, whereas 64% (18/28) of those not prescribed guideline-recommended antibiotics reported a recurrence (P = .01). Of the 383 participants, 291 were prescribed either clindamycin (n = 156) or TMP-SMX (n = 135). While 31% (n = 49) of those prescribed clindamycin reported a recurrent SSTI, 47% (n = 63) of those prescribed TMP-SMX reported a recurrence (P = .008).

We next tested a direct link between the effects of antibiotics (given at the time of SSTI) on subsequent *S. aureus* colonization and on recurrent SSTI. Of 357 children who had both repeat colonization sampling and longitudinal follow-up to ascertain recurrent SSTI, those remaining colonized with *S. aureus* at follow-up sampling were more likely to report a recurrent SSTI (101/178 [57%]) than those who were not colonized (54/179 [30%], P < .001) (Figure 1).

In a Cox proportional hazards regression analysis adjusting for age, race, methicillin susceptibility of the SSTI isolate, prescription of decolonization measures, burden of baseline *S. aureus* colonization, and colonization status at repeat sampling, participants prescribed guideline-recommended antibiotics for SSTI at the point of care were less likely to have a recurrent SSTI in the year following enrollment (aHR, 0.57; 95% CI, .34–.94) (Figure 3). Additionally, participants remaining colonized with *S. aureus* at repeat sampling were more likely to report a recurrent infection over 12 months (aHR, 2.37; 95% CI, 1.69–3.31) than those not colonized at follow-up sampling.

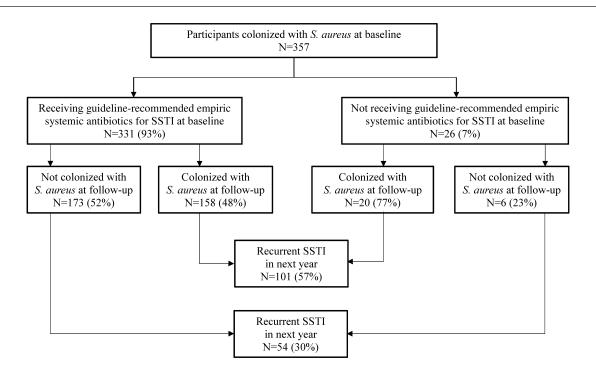


Figure 1. Participants with *Staphylococcus aureus* skin and soft tissue infection (SSTI) and colonization receiving and not receiving guideline-recommended empiric systemic antibiotics for SSTI in conjunction with incision and drainage. Participants receiving guideline-recommended empiric systemic antibiotics for SSTI were less likely to remain colonized with *S. aureus* (P=.004 by χ^2 testing) than those not prescribed guideline-recommended antibiotics. Participants remaining colonized with *S. aureus* at the follow-up sampling were more likely to report a recurrent SSTI (P<.001 by χ^2 testing) than those not colonized at follow-up.

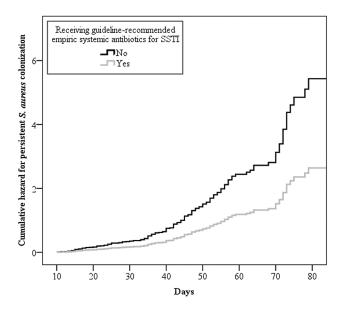


Figure 2. Cox proportional hazards regression analysis for remaining colonized with *Staphylococcus aureus* at follow-up sampling between participants receiving and not receiving guideline-recommended empiric systemic antibiotics for skin and soft tissue infection (SSTI) in conjunction with incision and drainage. Colonized participants receiving guideline-recommended empiric systemic antibiotics for SSTI were less likely to remain colonized with *S. aureus* than those not prescribed guideline-recommended antibiotics (adjusted hazard ratio, 0.49; 95% confidence interval, .30–.79).

DISCUSSION

Prior studies have demonstrated a significant reduction in recurrent SSTI after antibiotic utilization for the index SSTI [8, 18, 21]. However, the mechanism driving this phenomenon is unknown. We tested the hypothesis that this observation may be a result of the impact of systemic antibiotics on the burden of S. aureus colonization, which is a known risk factor for subsequent SSTI. In this study, we demonstrated a reduction in the prevalence of S. aureus colonization longitudinally, and confirmed a reduced incidence of recurrent infection, in children who received guideline-recommended empiric antibiotics in conjunction with I&D for their baseline S. aureus SSTI compared to those who did not receive guideline-recommended antibiotics. In looking for a link between these associations, we showed that children who remained colonized with S. aureus were more likely to report a recurrent infection than those in whom S. aureus colonization was eradicated. Hence, we have demonstrated that antibiotic administration at the time of acute SSTI affects S. aureus colonization, and that persistent colonization is associated with recurrent infection.

Children who were colonized with *S. aureus* at baseline screening and prescribed guideline-recommended antibiotics for their SSTI were less likely to remain colonized at their subsequent sampling (median of 38 days later) than children not prescribed antibiotics, suggesting that systemic antibiotics impact skin and nasal *S. aureus* colonization. Similar to the present

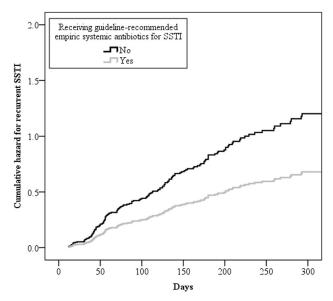


Figure 3. Cox proportional hazards regression analysis for recurrent skin and soft tissue infection (SSTI) for up to 1 year between participants receiving and not receiving guideline-recommended empiric systemic antibiotics for SSTI in conjunction with incision and drainage. Participants receiving guideline-recommended empiric systemic antibiotics for SSTI were less likely to report a recurrent SSTI than those not prescribed guideline-recommended antibiotics (adjusted hazard ratio, 0.57; 95% confidence interval, .34–.94).

study, in 144 children and adults with uncomplicated SSTI in Chicago, the prevalence of *S. aureus* colonization decreased significantly from baseline to 40-day follow-up after completion of a 10-day course of clindamycin or TMP-SMX [32]. Though not recommended for routine use as a decolonizing agent by the IDSA [17], systemic antibiotic administration for SSTI has benefits beyond resolution of the acute infection.

Topical decolonizing agents were prescribed for many of the participants in this study. To ensure that use of topical antimicrobials was not confounding the relationship between systemic antibiotics and S. aureus decolonization, use of such agents was included in the Cox proportional hazards regression analyses. Systemic antibiotics and topical antimicrobials were each independently associated with reduced S. aureus colonization at follow-up sampling. A randomized controlled trial of 146 MRSA-colonized hospitalized patients in Ontario, comparing a 7-day decolonization treatment consisting of chlorhexidine body washes, intranasal mupirocin, and oral rifampin and doxycycline vs no treatment, found this regimen to be efficacious in eradicating MRSA carriage 3 and 8 months following treatment [26]. This study, however, did not elucidate whether the topical or systemic components of the decolonization regimen were most effective in decolonizing patients of MRSA.

While the impact of systemic antibiotics on clinical cure of SSTI has varied among studies, several studies have demonstrated that patients receiving antibiotics experience significantly fewer recurrent SSTIs [8, 18, 21, 33]. In the present longitudinal study, risk of recurrent infection was decreased by half when antibiotics were prescribed after I&D for SSTI management, consistent with the results of several placebo-controlled trials. Among 1247 patients ≥12 years of age enrolled at 5 US emergency departments, TMP-SMX was superior to placebo in preventing subsequent I&D, skin infections at new sites, and skin infections in household members 14 days posttreatment [18]. In patients recruited from a pediatric emergency department in St Louis, TMP-SMX was superior to placebo in preventing new lesions at 10-day follow-up in 149 children with SSTI and I&D [8]. Additionally, among 190 adult patients with uncomplicated SSTI at 4 military emergency departments, fewer recurrent SSTIs were reported over 1 month following treatment in patients receiving TMP-SMX compared to placebo [21]. Length of therapy also impacts the incidence of recurrent infection. Of 249 children presenting to an emergency department in Buffalo with MRSA SSTI, recurrent infection was significantly higher 1 month following treatment among patients randomized to receive a 3-day course of TMP-SMX compared to those receiving a 10-day course of TMP-SMX [33].

Choice of antibiotic (eg, clindamycin vs TMP-SMX) may also be important in the management of SSTI. In the present study, within the subgroup of subjects receiving guideline-recommended antibiotics, clindamycin was superior to TMP-SMX in lowering the prevalence of persistent S. aureus colonization and in reducing the incidence of recurrent SSTI. There are conflicting reports in the literature regarding the superiority of clindamycin in the treatment of SSTI. Two studies of SSTI in Texas found no differences between clindamycin and TMP-SMX in terms of treatment failure or recurrence of infection [34, 35]. However, a retrospective comparative effectiveness study of nearly 50 000 children with SSTI in Tennessee revealed an increased risk of treatment failure and recurrent infection in those prescribed TMP-SMX compared to clindamycin in both patients with and without a drainage procedure [36]. Additionally, 3 randomized clinical trials of patients prescribed clindamycin or TMP-SMX found that clindamycin recipients were less likely to experience recurrent infection than TMP-SMX recipients [22, 37, 38]. Last, a Pennsylvania study of serial MRSA colonization cultures in patients with MRSA SSTI found that those receiving clindamycin for their SSTI showed earlier clearance of colonization [39] and were less likely to have recurrence of MRSA colonization after clearance than those prescribed other antibiotics [40]. While some have proposed a differential impact on the skin microbiome, the mechanism driving these observed effects requires further investigation [34, 39].

The present study has several limitations that should be noted. Participants were drawn from multiple *S. aureus* research studies, thus introducing some variation in both the interval between baseline and follow-up colonization swabs and longitudinal documentation of recurrent SSTI; however, this limitation was in part mitigated by use of Cox proportional hazards regression models. Additionally, antibiotic prescription and decolonization regimens varied. As most participants were seen as outpatients, antibiotic use was recorded as prescribed; proof of consumption was not feasible. Possibly reflecting current clinical practice and despite IDSA guidelines (based largely on expert opinion) suggesting that I&D alone is likely adequate for management of uncomplicated SSTI [17], the cohort of patients that did not receive antibiotics as part of treatment for their SSTI was small. Finally, all study participants presenting with *S. aureus* SSTI included in this cohort were also colonized with *S. aureus*; thus, the results of this study and the longitudinal impact of systemic antibiotics may not be generalizable to patients presenting with *S. aureus*.

In summary, we have demonstrated that systemic antibiotics prescribed at the time of I&D for acute SSTI reduces *S. aureus* colonization, which is protective against recurrent infection, establishing benefit of systemic antibiotics beyond the resolution of acute infection. Clindamycin was more effective than TMP-SMX at eradicating *S. aureus* colonization and preventing recurrent SSTI. These data will inform continuing conversations about optimal clinical management of SSTI in the contemporary era.

Notes

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