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Better Operating Room Ventilation as Determined by a Novel Ventilation Index is Associated with Lower Rates of Surgical Site Infections

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Associated with Lower Rates of Surgical Site Infections

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Running Head: Operating room: Ventilation index and SSI

ABSTRACT

Objective

To assess the impact of operating room (OR) ventilation quality on surgical site infections (SSI) using a novel ventilation index.

Summary Background Data

Previous studies compared laminar air flow with conventional ventilation, thereby ignoring many parameters that influence air flow properties. Freiburgstrasse 16p

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Methods

In this cohort study, we surveyed hospitals participating in the Swiss SSI surveillance and calculated a ventilation index for their ORs, with higher values reflecting less turbulent air displacement. For procedures captured between 01/2017–12/2019, we studied the association between ventilation index and SSI rates using linear regression (hospital-level analysis) and

with the individual SSI risk using generalized linear mixed-effects models (patient-level analysis).

Results

We included 47 hospitals (182 ORs). Among the 163'740 included procedures, 6791 SSIs were identified. In hospital-level analyses, a 5-unit increase in the ventilation index was associated with lower SSI rates for knee and hip arthroplasty (-0.41 infections per 100 procedures, CI -0.69 to -0.13), cardiac (-0.89, -1.91 to 0.12), and spine surgeries (-1.15, -2.56 to 0.26). Similarly, patient-level analyses showed a lower SSI risk with each 5-unit increase in ventilation index (adjusted odds ratio 0.71, CI 0.58 to 0.87 for knee and hip; 0.72, 0.49 to 1.06 for spine; 0.82, 0.69 to 0.98 for cardiac surgery). Higher index values were mainly associated with a lower risk for superficial and deep incisional SSIs. were identified. In hospital-level analyses, a 5-unit increase in the ventilation index was
associated with lower SSI rates for knee and hip arthroplasty (-0.41 infections per 100
procedures, CI -0.69 to -0.13), cardiac (-

Conclusions

Better ventilation properties, assessed with our ventilation index, are associated with lower rates of superficial and deep incisional SSIs in orthopedic and cardiac procedures. OR ventilation quality appeared to be less relevant for other surgery types.

Key Words: Operating room ventilation; surgical site infections; laminar air flow; prevention;

INTRODUCTION

Surgical site infection (SSI) is the most common preventable complication among patients who undergo surgery and is associated with substantial morbidity and healthcare $cost^{1-4}$. Prevention of SSIs requires a multimodal bundle approach: While efforts such as antimicrobial prophylaxis and surgical hand hygiene have been shown to be effective measures to prevent $SSIs^{5-7}$, and multiple other measures are being advocated in prevention guidelines, the role of operating room ventilation during the index surgery proved to be controversial $^{8-10}$.

Laminar air flow systems are designed to move filtered air uniformly with little or no turbulence into the operating field to minimize microbial wound contamination through $\ar{1}$. Although studies showed that laminar air flow systems reduce the bacterial load in an operating field¹², these findings did not translate into decreasing SSI rates in most clinical studies, as summarized in a large meta-analysis of observational studies¹³. However, all included studies relied on an oversimplified distinction between laminar air flow and conventional ventilation systems. These studies provide insufficient data on technical characteristics such as ceiling panel size, air flow, and presence or absence of objects above the operation field, which play an important role in the flow dynamics and hence in the capacity of ventilation systems to reduce microbial contamination $14-16$. antimicrobial prophylaxis and surgical hand hygiene have been shown to be effective
measures to prevent SSIs⁵⁻⁷, and multiple other measures are being advocated in preve
guidelines, the role of operating room ventilatio

To fill this research gap, we aimed to characterize the ventilation quality of Swiss operating rooms using a novel ventilation index encompassing a range of ventilation characteristics, and assessed the impact of ventilation quality on SSI rates using data from the national SSI surveillance database.

METHODS

Study setting and participants

This cohort study is based on the Swiss national SSI surveillance program (www.swissnoso.ch) including 168 of the 276 hospitals in Switzerland¹⁷, in which data on at least three procedure types per participating hospital are routinely recorded for all patients who undergo surgery. In addition to in-hospital surveillance, post-discharge surveillance is performed at 30 days after all procedures, and again at one year for individuals undergoing implant surgery, with complete one-year follow-up data available for more than 90% of operations. Trained infection control nurses perform regular systematic reviews of patient charts and standardized post-discharge phone interviews, supervised by infectious disease specialists. Here, we included all adult patients with complete follow-up between January 2017 and December 2019. Due to the high proportion of children undergoing appendectomy, we excluded this procedure from our analyses. who undergo surgery. In addition to in-hospital surveillance, post-discharge surveillan
performed at 30 days after all procedures, and again at one year for individuals underg
implant surgery, with complete one-year follow

For the study, all hospitals participating in the SSI surveillance program were contacted and asked to provide detailed technical information for all operating rooms in which procedures captured by the national surveillance were performed, including two standardized photographs to confirm the plausibility of the data entry (Figure S1, Supplemental Digital Content 1, http://links.lww.com/SLA/E164). All hospitals that provided measurements on at least one operating room were included in this analysis.

The Cantonal Ethics Committee of Bern, Switzerland (Project ID 2019-00294) approved the study. Patients were informed about their inclusion in the SSI surveillance on admission and given the opportunity to opt out. This study follows the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) reporting guideline¹⁸.

Ventilation Index

Previous laboratory experiments by members of our team examined the influence of specific ventilation system characteristics on air quality in operating rooms. Those experiments were performed in a real-life replication of an operating room (7.2 x 6.5 x 3.2m), in which specific changes both to the ventilation system and the operating field could be applied and tested. Under varying controlled simulations, optical particle measurements were obtained from within the operating field (in accordance with the Swiss and German industry recommendations), and a ratio was determined by comparing the measurements to a reference value. The resulting level of protection describes the ventilation induced air displacement and particle dilution, which correlate directly with the reduction in microbial burden¹⁹. Based on those results, we created a ventilation index that is influenced by the size of the supply air unit, the delivered air flow into the room, and affected negatively for factors that potentially cause turbulence (such as design and position of operating room lamps, location of return air outlets, and table position). The index was calculated differently for operating rooms with and without laminar air flow units, with higher values reflecting less turbulent air displacement properties (Table 1). Since only hospital- and procedure-specific data on SSI rates were available, a procedure specific ventilation index for each hospital and was calculated using the mean of indexes of all its operating rooms in which the respective procedures were performed. Under varying controlled simulations, optical particle measurements were obtained fre
within the operating field (in accordance with the Swiss and German industry
recommendations), and a ratio was determined by comparing t

Outcomes

The primary outcome was the hospital-specific rate of any SSI (including superficial incisional, deep incisional, or organ/space infection) according to National Healthcare Safety Network definitions from the Centers for Disease Control and Prevention²⁰. All SSIs at the 30-day or 1-year time points were included, and SSI rates were standardized using the National Nosocomial Infections Surveillance (NNIS) risk index, which accounts for differences in the American Society of Anesthesiology (ASA) score, the wound

contamination class (clean/clean-contaminated or contaminated/infected), and whether the procedure duration was above the 75th percentile²¹. Standardized SSI rates were calculated for each hospital, and individually for each type of surgical procedure within a given hospital. In addition to hospital-specific SSI rates, we also calculated the individual patient-level risk for any SSI in a secondary analysis.

Statistical analyses

To explore the association between the ventilation index and hospital-specific SSI rates, we fitted hospital-level multivariable linear regression models, adjusted for procedure type (cesarean section, cardiac, colorectal, hernia, hysterectomy, knee and hip, spine, and upper gastrointestinal surgery), including weights for the number of procedures performed during the observation period. As each hospital could contribute data for more than one procedure type, we calculated sandwich-type "robust" standard errors to account for intra-hospital correlation. In addition, separate weighted univariable linear regression models were performed individually for each procedure type. Adjusted odds ratios (aOR) were calculated using patient-level data by fitting multivariable generalized linear mixed effect models with a logit link function. The procedure-specific mean ventilation index for each hospital was used as exposure in these models, fitted with three distinct adjustment sets: The first model included the procedure type and all components from the NNIS risk index; the second model added whether the procedure was elective vs. urgent, use of adequate antibiotic prophylaxis (within 120 minutes prior to the incision for fluoroquinolones and vancomycin, and within 60 minutes for other antibiotics), and the patients age and sex; and the third model additionally comprised hospital-level information such as hospital size (by number of beds) and setting (public, private or university hospital). To account for correlation within hospitals, all patientlevel models included a random intercept for each hospital. We repeated the overall patientfor any SSI in a secondary analysis.

Statistical analyses

To explore the association between the ventilation index and hospital-specific SSI rate

fitted hospital-level multivariable linear regression models, adjusted fo

level analysis for each type of infection (superficial, deep incisional, and organ space) as outcome. As the proportion of missing data was low (below 0.5% for all variables included), analyses were performed on complete cases. Statistical analyses were performed using R, version $4.1.1²²$.

Sensitivity Analyses

We explored the robustness of our results using several sensitivity analyses on the hospital level. As our data did not allow matching patients and operating rooms directly, we relied on aggregating ventilation indexes within hospitals. To explore the impact of the aggregation method, we repeated the hospital-level analyses using the minimum and maximum ventilation index of all operating rooms, respectively. In addition, we performed an analysis including only hospitals with homogeneous ventilation indexes (defined as having a standard deviation of ventilation indexes below the 75th percentile) to limit the influence of operating rooms with very good or very poor ventilation properties. We also performed a hospital-level analysis restricted to public hospitals to limit the potential bias of differences in case mix and Sensitivity Analyses
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RESULTS

Description of participating hospitals

Out of 168 hospitals contacted, 51 hospitals from all parts of Switzerland completed our survey, and 48 (94%) provided enough details for us to calculate the ventilation index and had SSI data available. Another hospital was excluded since only data on appendectomies were available. Most of the 47 included hospitals were public (26, 55.3%), followed by private (17, 36.2%) and university hospitals (4, 8.5%). Thirty-one hospitals (66.0%) had less than 200 beds, 10 hospitals had 200 to 499 beds (21.3%), and 6 hospitals (12.8%) had 500 or more beds. Compared to hospitals participating in the SSI surveillance program that were excluded from our analyses $(n=121)$, those included were more likely to be university hospitals (8.5 % vs. 1.7%), and less likely to be private (36.2 % vs. 40.2%). The number of beds did not differ substantially between included and excluded hospitals, although the proportion of larger hospitals (500 beds and more) was higher in the included set of hospitals (12.5% vs. 5.1%). had SSI data available. Another hospital was excluded since only data on appendecton
were available. Most of the 47 included hospitals were public (26, 55.3%), followed by
private (17, 36.2%) and university hospitals (4,

Ventilation indexes were calculated for 182 operating rooms in the 47 included hospitals. The mean ventilation index was 8.3 (95% CI 7.7 to 9.0) and ranged from -5 to 18. Figure 1 shows a detailed description of ventilation indexes for all operating rooms and the hospital-specific mean ventilation index. Ventilation indexes disaggregated by procedure types are presented

Hospital-level analysis

In hospital-level analyses, an increase in ventilation index was not associated with an overall decrease in NNIS-adjusted SSI rates (a change in SSI rate per 5 steps increase in ventilation index: -0.38 infections per 100 interventions, 95% confidence interval [CI] -1.04 to 0.28, Figure 2). In procedure-stratified analyses, an increase of 5 units in ventilation index was associated with significant decreases in SSI rates for knee and hip arthroplasty (-0.41 infections per 100 interventions, 95% CI -0.69 to -0.13). Similarly, increases in ventilation indexes were associated with lower SSI rates in cardiac (-0.89 infections per 100 interventions, 95% CI -1.91 to 0.12) and in spine surgeries (-1.15, 95% CI -2.56 to 0.26), albeit confidence intervals included a null effect for the latter two procedure types. No changes in SSI rates were observed in upper gastrointestinal and colorectal surgeries, cesarean sections, hysterectomies, and hernia repairs (Table 2). Exploring the role of implants, ventilation index was associated with decreases in infections among spine surgeries without the use of implants $(-1.13$ infections per 100 interventions, 95% CI-2.16 to -0.11) and in those with implants (-1.01 infections per 100 interventions, 95% CI -4.08 to 2.06) although the latter was not statistically significant. Figure 2). In procedure-stratified analyses, an increase of 5 units in ventilation index w
associated with significant decreases in SSI rates for knee and hip arthroplasty (-0.41
infections per 100 interventions, 95% CI -

Patient-level analysis

Between January 2017 and December 2019, 163'740 procedures were included in our analyses. Patient and hospital characteristics are summarized in Table S1, Supplemental Digital Content 1, http://links.lww.com/SLA/E164. Within this three-year period, 6971 SSIs (4.3%) were identified: 2399 superficial incisional infections (34.4%), 1109 deep incisional infections (15.9%), and 3172 organ space infections (45.5%).

In models adjusting for variables from the NNIS risk index (minimal model), a 5-unit increase in ventilation index was associated with an overall lower risk for SSIs (aOR 0.90,

95% CI 0.80 to 1.00). In subgroup analyses, the largest reductions were again observed in knee and hip, spine, and cardiac surgery, with no clear associations observed in other surgery types. After additionally adjusting for whether the procedure was elective or not, the timing of antibiotic prophylaxis, patient age and sex (extended model), and after taking hospital size and type into account (full model), reductions in SSI rates associated with higher ventilation indexes remained significant for knee and hip, spine and cardiac surgery, but confidence intervals of overall estimates included a null effect (Figure 3). In patient-level analyses adjusted for the intervention type, the components of the NNIS risk index, emergent indication, use of antibiotic prophylaxis, age, and sex, a 5 unit increase in the ventilation index was associated with overall lower rates of superficial (aOR for all procedures 0.82, 95% CI 0.71 to 0.95), but not with deep incisional (aOR for all procedures 0.97, 95% CI 0.81 to 1.17), or organ/space infections (aOR 0.94, 0.82 to 1.08). Subgroup analyses showed lower rates of superficial incisional infections for knee and hip, cardiac and spine surgeries, and deep incisional infections for knee and hip surgeries with higher ventilation indexes. There were no significant associations with organ/space infections (Figure 4). and type into account (full model), reductions in SSI rates associated with higher venti-
indexes remained significant for knee and hip, spine and cardiac surgery, but confiden
intervals of overall estimates included a nu

Sensitivity analyses

Repeating the hospital-level analyses using minimum and maximum ventilation indexes instead of the mean ventilation index per hospital did not alter our findings substantially (Table S2, Supplemental Digital Content 1, http://links.lww.com/SLA/E164). In addition, results remained similar when only hospitals with homogeneous ventilation indexes were considered (knee and hip: -0.39 infections per 100 interventions, 95% CI -0.71 to -0.06; spine: -1.15 per 100 interventions, 95% CI -2.56 to 0.26; cardiac: -1.53, 95% CI -4.8 to 1.7). As ventilation indexes were higher in private (median 10.8, IQR 7.3 to 11.5) and university hospitals (11.0, IQR 8.0 to 11.0) compared with public hospitals (7.7, IQR 3.0 to 9.2, $p <$

0.001), we repeated the hospital-level analyses restricted to public hospitals. Similar to the full hospital-level analysis, ventilation indexes were not associated with an overall change in SSI rate (-0.20 infections per 100 interventions, 95% CI -0.75 to 0.36), however, the association remained statistically significant for knee and hip surgeries (-0.47 infections per 100 interventions, 95% CI -0.80 to -0.14).

DISCUSSION

In this nationwide study, operating room ventilation properties assessed using a novel ventilation index varied markedly between participating hospitals and between individual operating rooms within these hospitals. We observed lower rates of SSIs when orthopedic and cardiac procedures were performed in hospitals with better operating room ventilation properties and higher ventilation indexes. This association was observed in hospital- and patient-level analyses and was mainly driven by higher rates of superficial and deep incisional infections associated with lower ventilation indexes. Importantly, the strongest associations were present for knee and hip arthroplasty, spine, and cardiac surgeries, whereas no clear signal was present for gynecologic or abdominal surgeries, indicating that laminar air flow might be less important in these procedure types. 100 interventions, 95% CI-0.80 to -0.14).

DISCUSSION

In this nationwide sudy, operating room ventilation properties assessed using a novel

ventilation index varied markedly between participating hospitals and between i

The finding that ventilation quality is associated with lower SSI rates is supported by microbiological studies which demonstrated substantial reductions in bacterial counts within the operating field when laminar air flow was used^{12,23,24}. However, clinical studies evaluating the role of laminar air flow in orthopedic surgery did not show a beneficial impact on SSI rates^{13,25–27}. Since no standardized metric of operating room ventilation quality was available to date, previous studies mainly classified ventilation types into laminar air flow and conventional ventilation. However, this distinction does not capture the complexity of operating room ventilation, given that properties such as ceiling panel size¹⁴ or air flow play a crucial role in generating a truly laminar air flow¹⁴. For instance, lower SSI rates were

observed with high-volume but not with low-volume laminar air flow ventilation in an observational study performed in Norway, illustrating the importance of incorporating all components of operating room ventilation when determining the impact of ventilation on SSI rates 28 . To overcome this problem, we developed a novel and easy-to-calculate ventilation index, which encompasses a range of ventilation system characteristics. From an infection prevention perspective, such a tool may help to assess and compare operating room ventilation systems and provide a basis for allocating selected procedures to operating rooms with better ventilation characteristics.

While several previous studies explored the role of ventilation on the occurrence of deep incisional or organ/space infections^{13,25,26,29,30}, very little is known about the impact of ventilation quality on superficial incisional infections. One older study from Turkey observed a higher rate of sternal wound infections when operations were performed in rooms with outof-date ventilation technology³¹. Confirming and extending these findings, our study indicates that the lower rate of SSIs associated with improved laminar air flow properties was mainly driven by changes in the rate of superficial incisional infections followed by deep incisional, but not by organ/space nor implant-associated infections. index, which encompasses a range of ventilation system characteristics. From an infect
prevention perspective, such a tool may help to assess and compare operating room
ventilation systems and provide a basis for allocatin

SSI rates for procedures other than orthopedic and cardiac surgeries did not differ in function of the ventilation index in our study, indicating that ventilation may play a lesser role in these procedure types. These findings suggest that priorities of improving ventilation could be given to operating rooms where orthopedic and cardiac surgeries are performed, while maintaining conventional ventilation in other operating rooms to limit costs. However, whether laminar air flow ventilation actually leads to increases in operational costs remains a matter of debate $32-34$.

The present study included a large number of operations from a nationwide, wellcharacterized surveillance cohort of SSIs. Ascertainment of infections by trained infectious

disease physicians with outcome assessment extending to one year after the intervention further strengthens the validity of our results. In addition, accounting for antibiotic prophylaxis in our analyses avoids one of the major limitations of previous studies which did not include this information^{25,26}. Finally, findings were robust across a wide range of sensitivity analyses, including an analysis restricted to public hospitals, thereby reducing the potential bias due to differences in resources and case mix between hospitals. Some limitations should be noted. The surveillance database did not record the individual operating room in which the procedure was performed. Therefore, we had to rely on aggregated ventilation indexes per hospital as the exposure, which could have led to exposure misclassification. However, the results remained robust when using different aggregation methods, including when limiting our analyses to hospitals with homogeneous ventilation indexes across their operating rooms. Further, the need for aggregating the ventilation quality across operating rooms precluded an individual evaluation of each index component's influence on the occurrence of SSIs. In addition, subgroup analyses in the hospital-level analyses resulted in small numbers, which limited our power to detect differences. In consequence, patient-level analyses with increased statistical power (but with the limitation that no direct link between patient and operating room can be made) were performed in a second step, which largely confirmed the hospital-level findings. Finally, we cannot exclude the possibility that some hospitals reported only data on their best operating rooms. However, relying on an overly optimistic exposure is more likely to bias our results towards the null, and therefore would imply that the true association between ventilation index and SSI rates would be even stronger. sensitivity analyses, including an analysis restricted to public hospitals, thereby reduci
potential bias due to differences in resources and case mix between hospitals.
Some limitations should be noted. The surveillance d

In conclusion, our results indicate that performing orthopedic and cardiac interventions in operating rooms with good operating room ventilation properties (as assessed using a novel ventilation index proposed here) is associated with lower rates of superficial and deep

incisional SSIs. In contrast, ventilation might play a minor role in other surgical procedures. The beneficial impact of operating room ventilation quality on the subset of superficial and deep incisional infections needs confirmation in other cohorts. Further studies should prospectively include data on specific aspects of a given operating room that allow examining the influence of each component of the ventilation index on SSI detected in patients operated on in this particular room. Finally, cost-benefit analyses taking our findings into account are needed to further delineate the role of operating room ventilation on the occurrence of SSIs.

FOOTNOTES

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REFERENCES

- 1. Astagneau P, Rioux C, Golliot F, et al. Morbidity and mortality associated with surgical site infections: results from the 1997-1999 INCISO surveillance. J Hosp Infect. 2001;48:267–274.
- 2. Magill SS, Edwards JR, Bamberg W, et al. Multistate point-prevalence survey of health care-associated infections. N Engl J Med. 2014;370:1198–1208.
- 3. Shepard J, Ward W, Milstone A, et al. Financial impact of surgical site infections on hospitals: the hospital management perspective. JAMA Surg. 2013;148:907–914. 2. Magill SS, Edwards JR, Bamberg W, et al. Multistate point-prevalence survey of I
care-associated infections. N Engl J Med. 2014;370:1198-1208.
3. Shepard J, Ward W, Milstone A, et al. Financial impact of surgical site i
	- 4. Swissnoso. Swiss Point Prevalence Survey about Healthcare-associated Infections and Antibiotic Use in acute-care hospitals. March 2, 2020.
	- 5. Weber WP, Marti WR, Zwahlen M, et al. The timing of surgical antimicrobial prophylaxis. Ann Surg. 2008;247:918–926.
	- 6. Classen DC, Evans RS, Pestotnik SL, et al. The timing of prophylactic administration of antibiotics and the risk of surgical-wound infection. N Engl J Med. 1992;326:281–286.
	- 7. Tanner J, Dumville JC, Norman G, et al. Surgical hand antisepsis to reduce surgical site infection. Cochrane Database Syst Rev. 2016;CD004288.
	- 8. Kümin M, Scarborough M. Laminar flow ventilation during surgery. Lancet Infect Dis. 2017;17:581.
	- 9. Jutte PC, Traversari RA, Walenkamp GH. Laminar flow: the better choice in orthopaedic implants. The Lancet infectious diseases. 2017;17:695–696.
	- 10. Weinstein RA, Bonten MJM. Laminar airflow and surgical site infections: the evidence is blowing in the wind. The Lancet infectious diseases. 2017;17:472–473.
- 11. Hirsch T, Hubert H, Fischer S, et al. Bacterial burden in the operating room: impact of airflow systems. Am J Infect Control. 2012;40:e228–32.
- 12. Knudsen RJ, Knudsen SMN, Nymark T, et al. Laminar airflow decreases microbial air contamination compared with turbulent ventilated operating theatres during live total joint arthroplasty: a nationwide survey. J Hosp Infect. 2021;113:65–70.
- 13. Bischoff P, Kubilay NZ, Allegranzi B, et al. Effect of laminar airflow ventilation on surgical site infections: a systematic review and meta-analysis. Lancet Infect Dis. 2017;17:553–561.
- 14. Diab-Elschahawi M, Berger J, Blacky A, et al. Impact of different-sized laminar air flow versus no laminar air flow on bacterial counts in the operating room during orthopedic surgery. Am J Infect Control. 2011;39:e25–9. iont arthroplasty: a nationwide survey. J Hosp Infect. 2021;113:65–70.

13. Bischoff P, Kubilay NZ, Allegranzi B, et al. Effect of laminar airllow ventilation of

surgical site infections: a systematic review and meta-anal
	- 15. Cao G, Storås MCA, Aganovic A, et al. Do surgeons and surgical facilities disturb the clean air distribution close to a surgical patient in an orthopedic operating room with laminar airflow? Am J Infect Control. 2018;46:1115–1122.
	- 16. Erichsen Andersson A, Petzold M, Bergh I, et al. Comparison between mixed and laminar airflow systems in operating rooms and the influence of human factors: experiences from a Swedish orthopedic center. Am J Infect Control. 2014;42:665–669.
	- 17. Troillet N, Aghayev E, Eisenring M-C, et al. First Results of the Swiss National Surgical Site Infection Surveillance Program: Who Seeks Shall Find. Infect Control Hosp Epidemiol. 2017;38:697–704.
	- 18. von Elm E, Altman DG, Egger M, et al. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. Ann Intern Med. 2007;147:573–577.
- 19. Külpmann R, Hildebrand K. OP-Lüftungssysteme im Vergleich. GI-Gebäude Techn Innenraum Klima. 2013;134:12–29.
- 20. Horan TC, Andrus M, Dudeck MA. CDC/NHSN surveillance definition of health careassociated infection and criteria for specific types of infections in the acute care setting. Am J Infect Control. 2008;36:309–332.
- 21. Culver DH, Horan TC, Gaynes RP, et al. Surgical wound infection rates by wound class, operative procedure, and patient risk index. National Nosocomial Infections Surveillance System. Am J Med. 1991;91:152S–157S.
- 22. R Core Team. R: A Language and Environment for Statistical Computing Available from: https://www.R-project.org/. 2020.
- 23. Marsault LV, Ravn C, Overgaard A, et al. Laminar airflow versus turbulent airflow in simulated total hip arthroplasty: measurements of colony-forming units, particles, and energy consumption. J Hosp Infect. 2021;115:117–123.
- 24. Kirschbaum S, Hommel H, Strache P, et al. Laminar air flow reduces particle load in TKA-even outside the LAF panel: a prospective, randomized cohort study. Knee Surg Sports Traumatol Arthrosc. 2020;29:3641–3647. Am J Infect Control. 2008;36:309–332.

21. Culver DH, Horan TC, Gaynes RP, et al. Surgical wound infection rates by wound

operative procedure, and patient risk index. National Nosocomial Infections Surve

System. Am J Med
	- 25. Brandt C, Hott U, Sohr D, et al. Operating room ventilation with laminar airflow shows no protective effect on the surgical site infection rate in orthopedic and abdominal surgery. Ann Surg. 2008;248:695–700.
	- 26. Breier A-C, Brandt C, Sohr D, et al. Laminar airflow ceiling size: no impact on infection rates following hip and knee prosthesis. Infect Control Hosp Epidemiol. 2011;32:1097– 1102.
- 27. Bao J, Li J. The effect of type of ventilation used in the operating room and surgical site infection: A meta-analysis. Infect Control Hosp Epidemiol. 2020;42:931–936.
- 28. Langvatn H, Schrama JC, Cao G, et al. Operating room ventilation and the risk of revision due to infection after total hip arthroplasty: assessment of validated data in the Norwegian Arthroplasty Register. J Hosp Infect. 2020;105:216–224.
- 29. Teo BJX, Woo YL, Phua JKS, et al. Laminar flow does not affect risk of prosthetic joint infection after primary total knee replacement in Asian patients. J Hosp Infect. 2020;104:305–308.
- 30. Pinder EM, Bottle A, Aylin P, et al. Does laminar flow ventilation reduce the rate of infection? an observational study of trauma in England. Bone Joint J. 2016;98-B:1262– 1269.
- 31. Simsek Yavuz S, Bicer Y, Yapici N, et al. Analysis of risk factors for sternal surgical site infection: emphasizing the appropriate ventilation of the operating theaters. Infect Control Hosp Epidemiol. 2006;27:958–963. Norwegian Arthroplasty Register. J Hosp Infect. 2020;105:216–224.

29. Teo BJX, Woo YL, Phua JKS, et al. Laminar flow does not affect risk of prosihetion

infection after primary total knee replacement in Asian patients. J
	- 32. Cacciari P, Giannoni R, Marcelli E, et al. [Cost evaluation of a ventilation system for operating theatre: an ultraclean design versus a conventional one]. Ann Ig. 2004;16:803– 809.
	- 33. Seipp HM, Sprengel M, Wagner KH. [Differentiation of allotments for occupationally necessary and hygiene relevant costs of air technology designs for operating rooms]. Zentralbl Hyg Umweltmed. 1994;195:398–410.
	- 34. Külpmann R, Kramer A. Raumlufttechnische Anlagen. Krankenhaus- und Praxishygiene. München: Elsevier, Urban & Fischer Verlag; 2016.

FIGURE LEGENDS

Figure 1 Ventilation indexes of all 182 operating rooms from 47 participating hospitals Legend: Each square indicates a distinct operating room (OR) within a hospital. The ventilation index summarizes properties of laminar air flow quality, with higher ventilations indexes implying less turbulent air displacement. For each hospital, ventilation quality was

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Figure 2 Association between ventilation index and NNIS-adjusted surgical site infection rate on the hospital level

Legend: Each point represents one hospital, and its size correlates with the number of procedures performed within that hospital between 2017 and 2019. The line and 95% confidence interval ribbon show the association between ventilation index and the surgical site infection rates, adjusted for the number of procedures and the type of surgery performed. confidence interval ribbon show the association between ventilation index and the surge
site infection rates, adjusted for the number of procedures and the type of surgery perfection.
NNIS = National Nosocomial Infections

Figure 3 Patient-level analysis assessing the odds ratios of surgical site infections, overall and stratified by surgery type, per 5 units increase in ventilation index

Legend: The minimal model is adjusted for all components of the NNIS risk index, and the intervention type. The extended model includes the same covariates as the minimal model, and additionally elective vs. urgent surgery, adequate timing of antibiotic prophylaxis (within 120 minutes for fluoroquinolones and vancomycin, and within 60 minutes for other antibiotics), age, and sex. The full model includes all covariates, including hospital size and type (public, private, and university). and additionally elective vs. urgent surgery, adequate timing of antibiotic prophylaxis

120 minutes for fluoroquinolones and vancomycin, and within 60 minutes for other

antibiotics), age, and sex. The full model includes

 $GI =$ gastrointestinal, C-section = cesarean section, $OR =$ odds ratio, $CI =$ confidence interval.

Figure 4 Patient-level analysis assessing the odds ratios of different types of surgical site infections, overall and stratified by surgery type, per 5 units increase in ventilation index. Legend: Patient-level analysis for each type of surgical site infection. The models are adjusted for all components of the NNIS risk index, elective vs. urgent surgery, adequate timing of antibiotic prophylaxis (within 120 minutes for fluoroquinolones and vancomycin, and within 60 minutes for other antibiotics), age, and sex. timing of antibiotic prophylaxis (within 120 minutes for fluoroquinolones and vancom
and within 60 minutes for other antibiotics), age, and sex.
GH = gastrointestinal, C-section = cesarean section, OR = odds ratio, CH = co

 $GI =$ gastrointestinal, C-section = cesarean section, $OR =$ odds ratio, $CI =$ confidence interval,

Table 1 Calculation of the ventilation index

Table 2 Hospital-level analysis of ventilation index and SSI rate, by surgery type

*Changes in SSI rates, per 5 units in the ventilation index.