Echocardiography Is Dispensable in Uncomplicated Staphylococcus aureus Bacteremia

Riad Khatib, MD, and Mamta Sharma, MD

Abstract: Current Staphylococcus aureus bacteremia (SAB) practice guidelines stratify treatment duration according to the likelihood of complications and recommend transesophageal echocardiography (TEE) in all cases. The benefit of TEE in uncomplicated SAB has not been validated. We performed a retrospective analysis of TEE and transthoracic echocardiography (TTE) among hospitalized adults with SAB in 3 prior observational studies (2002-2003, 2005-2006, and 2008-2009). Echocardiograms were ordered at the attending physician's discretion. SAB cases were stratified into the following types: complicated (persistent bacteremia [duration ≥ 3 d], relapse, and/or secondary foci); device-associated (intracardiac prosthetic devices); suspected endocarditis (the presence of murmurs or emboli); and uncomplicated (bacteremia duration ≤ 2 d, no device and/or secondary foci). We encountered 960 SAB cases; 83 were excluded (57 death/transfer/discharge within 48 h; 19 contaminants/ no treatment; 7 care withdrawn). TEE and TTE were performed within 0-28 days of SAB onset in 177 (20.2%) and 321 (36.6%) instances, respectively. TEE was positive (with signs of endocarditis) in 42/177 (23.7%) cases: 7/39 (17.9%) community associated and 35/138 (25.4%) health care associated. It was positive in 29/120 (24.2%) complicated, 3/11 (27.3%) device-associated, 9/15 (60.0%) suspected endocarditis, and 1/31 (3.2%) uncomplicated cases of SAB. TTE was positive in 25/321 (7.8%) cases of SAB, 1 was uncomplicated; it was negative in 20/30 (66.7%) TEE-positive cases. Follow-up of ≥100 days was possible in 282/361 (78.1%) uncomplicated SAB; many (46.8%) received ≤15 days of therapy. None of them had relapses or secondary foci.

These findings suggest that echocardiography is dispensable in cases of uncomplicated community-associated and health care-associated SAB. It should be limited to subsets with clinical findings of endocarditis, persistence, intracardiac devices, secondary foci, and relapse. The cost effectiveness of TTE prior to TEE among these patients is unknown.

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Abbreviations: BC = blood culture, SAB = Staphylococcus aureus bacteremia, TEE = transesophageal echocardiogram, TTE = transthoracic echocardiogram.

INTRODUCTION

Staphylococcus aureus bacteremia (SAB) remains a common and serious infection. It is associated with a high risk for secondary foci and infective endocarditis.^{1,2,4,11,17,25} Therefore, extended treatment duration was often used.11,17 With the widespread use of echocardiography, transesophageal echocardiogram

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ISSN: 0025-7974 DOI: 10.1097/MD.0b013e318294a710 (TEE) was advocated to optimize treatment duration. Subsequently, the need for extended treatment duration in uncomplicated intravenous catheter-associated SAB was challenged.^{15,18} In an attempt to support short-term treatment for uncomplicated intravascular catheter-associated bacteremia, Rosen et al²² proposed TEE as a cost-effective measure to avoid prolonged treatment. Since then, TEE was often advocated in SAB.^{3,7,24} Kern¹¹ recommended echocardiography in patients with communityonset disease, intravenous drug users, those with persistent bacteremia, patients with vascular implants, and hemodialysis-dependent patients. Mortara et al¹⁶ proposed limiting echocardiography to community-associated disease. The most recent Infectious Diseases Society of America practice guidelines for the treatment of SAB recommend echocardiography, preferably TEE, in all patients.14 The benefit of this approach in uncomplicated SAB has not been clearly documented. The risk for endocarditis and distant complications is known to be higher in patients with prolonged bacteremia.^{2,8,17,25} Whether echocardiography is dispensable in patients without these risk factors is unclear. Kaasch et al¹⁰ recently recommended the use of criteria to select cases for echocardiography among patients with nosocomial SAB. An accompanying editorial supported this approach.²³ We conducted the current study to review the results of echocardiography in community- and health care-associated SAB and to define clinical criteria that may optimize TEE predictive values.

PATIENTS AND METHODS

Patients

Adult (aged ≥18 yr) inpatients with SAB identified during 3 prior observational studies (2002-2003, 2005-2006, and 2008-2009) were included.¹² Selection methods were similar in all studies. All patients with ≥ 1 positive blood culture (BC) for S. aureus were screened. All patients with SAB (positive BC and clinical signs of bacteremia) were included. Patients who died or were discharged within 48 hours and who did not receive any treatment (care withdrawn or considered contaminant) were excluded. SAB cases separated by ≥ 100 days were counted as a new bacteremia.

The following information was gathered prospectively during each of the prior observational studies: demographics, underlying conditions, presence of cardiac devices (prosthetic valve, pacemakers/defibrillators), source of infection, onset mode, duration of bacteremia, TTE and TEE performance with the date and the results, treatment, and outcome measures that included the development of secondary foci (anatomically unrelated to the primary focus and became apparent after the onset of bacteremia), relapse, and 100-day all-cause mortality. Comorbidity score was retrospectively calculated according to the modified Charlson weighted index.19

Definitions

The onset of SAB was classified into 3 mutually exclusive categories according to the Centers for Disease Control definition.¹⁸ Hospital onset: SAB after 3 days of inpatient hospitalization. Health care-associated community-onset: SAB in patients

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with invasive devices or exposure to a health care setting within the preceding 12 months. *Community-associated*: SAB within 3 days of admission without invasive devices or health care exposure during the preceding 12 months.

The duration of bacteremia was defined as the number of days between the first and last positive BC or resolution of symptoms in patients without follow-up BC as previously described.¹² Persistent bacteremia was defined as bacteremia for \geq 3 days. Positive echocardiogram was defined as echocardiographic evidence of infective endocarditis such as an intracardiac oscillating mass, abscess, new valvular regurgitation, or new dehiscence of a prosthetic valve, according to the criteria proposed by Fowler et al.⁷

Echocardiograms were ordered according to the discretion of the attending physician or the consultant. TEE was performed and evaluated by an experienced echocardiographer. TTE was performed by an experienced technician and evaluated by an experienced echocardiographer. For echocardiogram utility assessment we included all patients with TTE or TEE within 0-28 days of the first positive BC. Echocardiograms performed after 28 days were excluded because they were often ordered for reasons other than the SAB. SAB was classified into 4 exclusive categories: complicated (bacteremia duration ≥ 3 d, relapse, and/or secondary foci); the presence of cardiac devices (prosthetic valves, pacemakers, defibrillators) without complicated bacteremia, suspected endocarditis (the presence of murmurs or emboli) without complicated bacteremia, and uncomplicated bacteremia (bacteremia duration ≤ 2 d and without devices, secondary foci, or relapse within 100 d). TEE and TTE results were then stratified according to SAB type. Follow-up for 100 days or more was attempted by reviewing subsequent medical encounters in all patients with uncomplicated bacteremia who survived. Information sought in follow-up included recurrence of bacteremia and the development of distant foci of infection such as endocarditis or osteomyelitis.

The study was approved by the St John Hospital institutional review board.

Statistical Methods

The chi-square test and student t test were used to assess the significance of differences in categorical and continuous variables, respectively. Sensitivity, specificity, positive and negative predictive value for TEE evidence of infective endocarditis in SAB were calculated for each selected clinical criteria as follows: sensitivity (positive TEE in patients with the selected criteria/all TEE performed in patients with the selected criteria), specificity (negative TEE without the selected criteria/negative TEE without the selected criteria + positive TEE in patients with the selected criteria), positive predictive value (positive TEE in patients with the selected criteria/all positive TEE), and negative predictive value (negative TEE in patients without the selected criteria/all negative TEE).

Statistical analyses were performed using SPSS v 12 and a web-based open source epidemiologic statistics software for public health (openepi.com). P value < 0.05 was considered to indicate statistical significance.

RESULTS

We encountered 960 episodes of SAB in patients during the 3 study periods; 83 were excluded because of death within 48 h (n = 45), discharge within 48 h (n = 12), care withdrawn within 48 h (n = 7), and suspected contamination (n = 19). The remaining 877 cases of SAB occurred among 805 patients. Thirty-three patients had 2 cases of bacteremia, 33 had 3 cases, and 6 patients had 4 cases of bacteremia. The majority of SAB cases (87.2%) were health care associated; 30.7% were hospital-onset and 56.5% were health care-associated community-onset cases.

Follow-up BC was obtained in 86.4% of cases. The first follow-up culture was obtained within a median of 2 days. Additional follow-up BC were obtained within a median of 3 days. The time between last positive and first negative BC, available in 677 patients, was 1-17 days with a median = 3 days. A summary of all patients and the frequency of endocarditis based on clinical and/or TEE findings is shown in Table 1. An echocardiogram was performed in 379 (43.2%) episodes (Fig. 1).

The frequency of TEE and TTE performance was stratified according to the type of bacteremia (Fig. 2). The characteristics of patients with and without echocardiograms were compared (Table 2). Clinicians were more likely to resort to echocardiography in community-associated infections (p < 0.001), the presence of intracardiac devices (p < 0.001), secondary foci (p < 0.001), and with bacteremia duration ≥ 3 days (p < 0.001).

TTE was performed within 0-28 days (median, 4.2 d) in 321 (36.6%) cases. TEE was performed within 1-28 d (median, 7.2 d) in 177 (20.2%) instances.

TTE and TEE showed findings consistent with infective endocarditis in 25/321 (7.8%) and 42/177 (23.7%) instances, respectively. Echocardiogram results were stratified according to

TABLE 1. Characteristics of Patients With Staphylococcus aureus Bacteremia and Frequency of Infective Endocarditis Stratified According to the Characteristics

Characteristic (N)	N With IE (%; 95% CI)
Diabetes (364)	24 (6.6; 4.4–9.5)
Hemodialysis (222)	13 (5.9; 3.3–9.6)
Intravenous drug use (86)	18 (20.9; 13.3–30.5)*
Type of onset	. (, ,
Community-associated (112)	26 (23.2; 16.1-31.7)*
Health care-onset (269)	7 (2.6; 1.1–5.1)
Health care-related community-onset (496)	31 (6.3; 4.4–8.7)
Intracardiac device (104)	17 (16.3; 10.2–24.4) [†]
Oxacillin-resistant (505)	31 (6.1; 4.3-8.5)
Uncomplicated SAB [‡] (466)	2 (0.4; 0.07–1.4)*
Source [§]	
IVC (244)	41 (16.8; 12.5–21.9)
Other endovascular foci (76)	21 (27.6; 18.5–38.5)
Tissue source (344)	53 (15.4; 11.9–19.5)
Unknown (149)	17 (4.7; 2.1–9.1)
Secondary foci (111)	38 (34.2; 25.9–43.4) [∥]
SAB duration	
<3 d (561)	19 (3.4; 2.1–5.1)*
≥3 d (316)	45 (14.2; 10.7–18.4)
N of SAB episodes	
1 (805)	57 (7.1; 5.5–9.0)
2-4 (72)	7 (9.7; 4.4–18.3)
All patients (877)	64 (7.3; 5.2–9.2)

Abbreviations: CI = confidence interval, IE = infective endocarditis, IVC = intravascular catheter.

*P < 0.001.

 $\mbox{$^{\rm C}$}$ Bacteremia duration ${}^{\rm <3}$ d without intracardiac device or secondary foci.

§Excluding cases with endocarditis.

||P < 0.0001.

 $[\]dagger P = 0.001.$

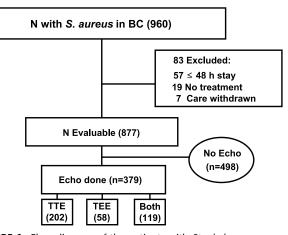


FIGURE 1. Flow diagram of the patients with *Staphylococcus aureus* in 1 or more blood culture (BC) included in echocardiography (Echo) assessment. Abbreviations: TTE = transthoracic echocardiogram; TEE = transesophageal echocardiogram.

SAB category (Fig. 3). TEE was positive in 1 (3.2%) case of uncomplicated methicillin-susceptible *S. aureus* (MSSA) bacteremia in a hemodialysis patient with a hemodialysis graft infection. TTE was positive in a hemodialysis patient with methicillin-resistant *S. aureus* (MRSA) bacteremia suspected to be due to access infection. TEE performed 2 days later did not verify TTE findings.

TEE was negative in 6/15 patients with clinically suspected endocarditis without meeting the criteria for complicated bacteremia. One with a stroke and SAB of unknown source, 1 with meningitis plus septic knee and a murmur, 1 with multiple traumatic soft tissue injuries and pulmonary contusions plus a murmur, 1 intravenous drug user with multiple pulmonary infiltrates, 1 with a known valvular heart disease, and 1 hemodialysis patient with access infection and a murmur. The remaining patients with suspected endocarditis who did not have TEE included 3 patients who died within 3–7 days, 2 who left against medical advice, 1 with a negative TEE performed for unrelated reasons before SAB was diagnosed, and 1 patient who refused the procedure.

Comparison of uncomplicated SAB cases with and without echocardiography is shown in Table 3.

The sensitivity, specificity, and positive and negative predictive values of selected clinical criteria for TEE evidence of infective endocarditis are shown in Table 4. TEE was more often positive among hemodialysis-dependent patients, intravenous drug users, and those with clinically suspected endocarditis. It was significantly less likely to be positive in patients with uncomplicated bacteremia. No difference was noted in mean comorbidity score between patients with and without endocarditis (2.8 ± 2.2 [with endocarditis] vs. 2.6 ± 2.1 [without endocarditis]; p = 0.5).

Among hemodialysis cases, TEE was positive in 11/34 (32.4%) complicated, 1/2 (50.0%) device-associated, 6/7 (85.7%) suspected endocarditis, and 1/8 uncomplicated bacteremia cases. Among intravenous drug users, TEE was positive in 10/15 (66.7%) complicated, 0/2 device-associated, 3/4 (75.0%) suspected endocarditis, and 0/2 uncomplicated bacteremia cases.

We applied the criteria of Kaasch et al¹⁰ to our patients who had TEE. Both TTE and TEE were performed in 119 cases. TEE was done within 1–2, 3–5, and \geq 6 days of TTE in 34 (28.6%), 41 (34.4%), and 44 (37.0%) cases, respectively. TTE was negative in 20/30 TEE-positive cases, and was positive in 2 cases with negative TEE. The 2 cases with TTE-positive/TEE-negative results were a patient with positive TTE on the first day of bacteremia and negative TEE 2 days after the TTE, and a second patient with a negative TEE 4 days after the onset of bacteremia and a positive TTE 7 days after TEE.

Follow-up for 100 days or more was possible in 507/669 patients who survived. Follow-up was accomplished in 282/361 patients with uncomplicated bacteremia including 81/92 (88.0%) with echocardiography. Duration of therapy was >15 days in 71.6% and 46.0% of patients with and without follow-up, respectively. None had a relapse or secondary complications.

DISCUSSION

These findings illustrate that endocarditis is rare in cases of uncomplicated community-associated, hospital-onset, and health care-associated community-onset SAB, and question the need for echocardiography in these cases. Current Infectious Diseases Society of America practice guidelines advocates echocardiography, preferably TEE, in all cases of SAB.14 These guidelines were probably generated based on the concerns for SAB complications and the positive predictive value of TEE.^{10,24} Going back to the history of SAB, a high incidence of endocarditis among patients with SAB without an identifiable source was often noted.¹⁷ Because of these concerns, and the acknowledged insensitivity of clinical findings for the diagnosis of endocarditis,7,24 treatment was often extended in all patients with SAB. Subsequently, treatment for 10-14 days was recommended for uncomplicated intravascular catheter-associated SAB,15,18 although this short-term therapy was not universally accepted.9,20

Then, additional studies attempted to identify patients at high risk for complications who warrant extended therapy.⁵ Fowler et al⁸ proposed that patients with positive surveillance BC, 3 days or more after the first positive culture, have higher incidence of complications. The risk for complications among patients with positive follow-up BC on day 3 after the initial set was also noted in other studies.² Around the same time, the Duke criteria⁶ for the diagnosis of infective endocarditis, which use echocardiography, were proposed. Consequently, echocardiography was advocated in SAB, and treatment stratification was suggested based on echocardiography results.^{14,22} Most of these

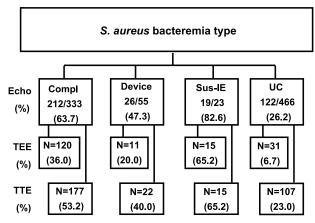


FIGURE 2. Echocardiography in *Staphylococcus aureus* bacteremia stratified according to bacteremia category. Abbreviations: Compl = complicated; Sus-IE = suspected infective endocarditis; UC = uncomplicated.

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TABLE 2. Characteristics of Patients With SAB, by Echocardiogram Status

Characteristic (N)	Echocardiogram Status, No. (%)				
	None	TEE and TTE	TEE Only	TTE Only	
Diabetes (364)	212 (58.2)	42 (11.5)	26 (7.2)	84 (23.1)	
Hemodialysis (222)	118 (53.1)	30 (13.5)	21 (9.5)	53 (23.9)	
Intravenous drug use (86)	45 (52.3)	16 (18.6)	7 (8.2)	18 (20.9)	
Type of onset					
Community-associated (112)	47 (42.0)	30 (26.8)	9 (8.0)	26 (23.2)	
Health care-onset (269)	185 (68.8)	18 (6.7)	20 (7.4)	46 (17.1)	
Health care-related community-onset (496)	266 (53.6)	71 (14.3)	29 (5.9)	130 (26.2)	
Intracardiac device (104)	36 (34.6)	25 (24.0)	16 (15.4)	27 (26.0)	
Oxacillin-resistant (505)	281 (55.6)	61 (12.1)	36 (7.1)	127 (25.2)	
Uncomplicated SAB* (466)	344 (73.8)	16 (3.4)	15 (3.2)	91 (19.6)	
Source					
IVC (244)	152 (62.3)	24 (9.8)	17 (7.0)	51 (20.9)	
Endocarditis (64)	6 (9.4)	35 (54.7)	10 (15.6)	13 (20.3)	
Other endovascular foci (76)	30 (39.5)	13 (17.1)	8 (10.5)	25 (32.9)	
Tissue source (344)	208 (60.5)	35 (10.2)	18 (5.2)	83 (24.1)	
Unknown (149)	102 (68.5)	12 (8.0)	5 (3.4)	30 (20.1)	
Secondary foci (111)	14 (12.6)	49 (44.2)	18 (16.2)	30 (27.0)	
Blood cultures in 1–3 d (618)	327 (52.9)	97 (15.7)	48 (7.8)	146 (23.6)	
SAB duration					
<3 d					
≥3 d (316)	112 (35.4)	83 (26.3)	36 (11.4)	85 (26.9)	
100-d mortality (183)	94 (51.4)	20 (10.9)	11 (6.0)	58 (31.7)	
Follow-up ≥100 d (507)†	293 (57.8)	73 (14.4)	36 (7.1)	105 (20.7)	
All patients (877)	496 (56.8)	119 (13.6)	58 (6.6)	202 (23.0)	

Abbreviations: See Table 1.

*Bacteremia duration <3 d without intracardiac device, secondary foci, or relapse within 100 d.

[†]Percentage among surviving patients.

studies, however, were observational and may have selection bias favoring TEE performance to patients with high risk for complications.²³ Additionally, cost-effectiveness analysis was based on decreasing treatment duration in cases with negative TEE.²² This analysis did not assess the feasibility of 10-14 days of therapy without resorting to echocardiography in uncomplicated bacteremia. Nevertheless, the widespread use of echocardiography, the modified Duke criteria for the diagnosis of infective endocarditis,¹³ and the reported complications in SAB without identifiable risk factors probably influenced the formulation of treatment guidelines.

The utility of echocardiogram in SAB remains debatable. Kaasch et al¹⁰ recently recommended clinical/microbiologic criteria to aid in selecting cases for TEE. The criteria included bacteremia for >4 days' duration, the presence of a permanent intracardiac device, hemodialysis dependency, spinal infection, and nonvertebral osteomyelitis. Our criteria were slightly different. We chose bacteremia for 3 days or more because it has been shown to be associated with higher complications.⁸ We included all secondary foci, whereas Kaasch et al included only spinal infection and osteomyelitis. We did not include hemodialysis dependency as a criterion. Our proposed criteria had excellent predictive values.

Our findings support their recommendation, and we propose to extend it to all uncomplicated SAB, including community-associated cases. Kaasch et al¹⁰ limited their analysis to health-care related cases.

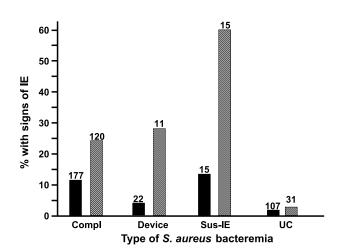


FIGURE 3. Echocardiogram with signs of infective endocarditis (IE) in *Staphylococcus aureus* bacteremia: Transthoracic (solid bars) and transesophageal (shaded bars) echocardiogram results, stratified according to the category of bacteremia. Abbreviations: Compl = complicated; Sus-IE = suspected infective endocarditis; UC = uncomplicated.

Characteristic (N)	Echocardiogram Status, No. (%)				
	None	TEE and TTE	TEE Only	TTE Only	
Diabetes (201)	149 (74.1)	9 (4.5)	4 (2.0)	39 (42.9)	
Hemodialysis (101)	73 (21.2)	13 (12.9)	5 (5.0)	20 (19.8)	
Intravenous drug use (39)	31 (9.0)	1 (2.6)	1 (2.6)	6 (15.4)	
Type of onset					
Community-associated (61)	40 (65.6)	50 (8.2)	0	16 (26.2)	
Health care-onset (146)	122 (83.6)	1 (0.7)	4 (2.7)	19 (13.0)	
Health care-related community-onset (259)	182 (70.3)	10 (3.9)	11 (1.2)	56 (21.6)	
Oxacillin-resistant (265)	198 (74.4)	7 (2.6)	9 (3.4)	51 (19.2)	
Source					
IVC (114)	88 (77.2)	1 (0.9)	7 (6.1)	18 (15.8)	
Endocarditis (2)	0	2 (100)	0	0	
Other endovascular foci (30)	18 (60.0)	3 (10.0)	2 (6.7)	7 (23.3)	
Tissue source (222)	160 (72.1)	6 (2.7)	4 (1.8)	52 (23.4)	
Unknown (98)	78 (79.6)	4 (4.1)	2 (2.0)	14 (14.3)	
SAB duration					
1 d (414)	311 (75.1)	14 (3.4)	10 (2.4)	79 (19.1)	
2 d (52)	33 (63.5)	2 (3.8)	5 (9.6)	12 (23.1)	
100-d mortality (86)	60 (69.8)	4 (4.6)	1 (1.2)	21 (24.4)	
N of SAB episodes					
1 (434)	318 (73.3)	15 (3.4)	13 (3.0)	88 (20.3)	
2-4 (32)	26 (81.2)	1 (3.1)	2 (6.3)	3 (9.4)	
Follow-up >100 d (282)†	201 (71.3)	11 (3.9)	13 (4.6)	57 (20.2)	

TABLE 3. Characteristics of Patients With Uncomplicated* SAB, by Echocardiogram Status

*Bacteremia duration <3 d without intracardiac device, secondary foci, or relapse within 100 d. †Percentage among surviving patients.

TABLE 4. Sensitivity, Specificity, Positive and Negative Predictive Value* for TEE Evidence of IE in SAB, Stratified by Selected Clinical Criteria

Selected criterion (N)	N With IE (p)†	Sensitivity (%; 95% CI)	Specificity (%; 95% CI)	Positive PV (%; 95% CI)	Negative PV (%; 95% CI)
Uncomplicated SAB [‡] (31)	1 (0.003)	NA	NA	NA	NA
Hemodialysis (51)§	19 (0.02)	37.3 (24.9–51.1)	81.7 (74.3-87.8)	45.2 (30.8-60.4)	76.3 (68.6-82.9)
Intravenous drug use (23)	13 (<0.001)	56.5 (36.1-75.4)	81.2 (74.4-86.8)	31.0 (18.4-46.0)	92.6 (87.2–96.2)
Diabetes (68)	18 (0.5)	26.5 (17.0-37.9)	78.0 (69.5-85.0)	42.9 (28.6-58.1)	63.0 (54.6–70.8)
Community-associated (39)	7 (0.3)	17.9 (8.2–32.3)	74.6 (66.9-81.4)	16.7 (7.6-30.2)	76.3 (68.6-82.9)
SAB duration ≥ 3 d (109)	25 (0.8)	22.9 (15.8–31.5)	74.1 (61.8-84.2)	64.3 (49.1–77.6)	68.1 (59.9–75.6)
Device-associated (41)**	12 (0.3)	29.3 (16.9-44.5)	77.9 (70.4–84.3)	28.6 (16.5-43.5)	78.5 (71.0-84.8)
Suspected endocarditis (45)††	23 (0.001)	51.1 (36.7-65.4)	85.6 (78.8–90.8)	54.8 (39.6-69.2)	83.7 (76.8-89.2)
Secondary foci (67)	27 (<0.001)	40.3 (29.1-52.3)	86.4 (79.0–91.9)	64.3 (49.1–77.6)	70.4 (62.3–77.6)
Proposed criteria, present report ^{‡‡} (146)	41 (0.003)	28.1 (21.3-35.8)	96.8 (85.1–99.8)	97.6 (88.8–99.9)	22.2 (15.8–29.8)
Kaasch criteria ¹⁰ (126)	35 (0.05)	27.8 (20.5-36.1)	86.3 (74.7–93.8)	83.3 (69.8–92.4)	32.6 (25.1-40.8)

Abbreviations: See previous tables. NA = not applicable, PV = predictive value.

*Calculated based on TEE results in the presence or absence of the selected criteria.

[†]P values represent chi-square test result in comparison to patients without the characteristic.

[‡]Bacteremia duration <3 d without intracardiac device or secondary foci.

§Forty-three met the criteria for complicated SAB.

¶Twenty-one met the criteria for uncomplicated SAB.

**Thirty met the criteria for complicated bacteremia.

††Thirty met the criteria for complicated bacteremia.

^{‡‡}Complicated bacteremia, intracardiac devices, and/or secondary foci.

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Our observations also suggest that most clinicians in our hospital do not follow current practice guidelines and do not routinely resort to echocardiography. Additionally, they often order TTE only. The approach of screening with TTE and preserving TEE for cases with negative TTE had been suggested, probably because of potential complications due to anesthesia or the endoscopies.^{3,16,26} The cost-effectiveness of this approach among patients who meet the criteria for echocardiography has not been thoroughly evaluated. With the known low TTE sensitivity,^{7,21} TEE is probably pursued in most cases. Our data are insufficient to assess the benefit of TTE because of the wide variation in the time-lag between TTE and TEE.

The current study has many limitations. First, although the data were obtained prospectively, the analysis was retrospective with potential bias. Second, most of our attending physicians and consultants appear to have a bias for selecting cases for echocardiography and do not follow current practice guidelines. However, this bias is likely to select patients with suspected complications and overestimates TEE utility. Additionally, the timing of echocardiography was not standardized. It is possible that some patients may develop vegetations several days after the onset of bacteremia. The numbers of patients with uncomplicated bacteremia among drug users and hemodialysis cases were too small to assess the TEE benefit in these patients. We believe that the high frequency of positive TEE in these patients could be explained by their propensity for having complications. Furthermore, we did not have follow-up in about 22% of patients with uncomplicated bacteremia who survived. Lastly, most patients received prolonged therapy that may have prevented the development of complications.

In summary, our findings suggest that TEE is dispensable in cases with uncomplicated community-associated, hospital-onset, and health care-associated community-onset SAB. We recommend limiting echocardiography to patients with positive follow-up BC 3 days after the initial set, prior history of endocarditis, clinical signs of endocarditis, device-associated cases, secondary foci, and relapse. Additional studies are needed to validate the use of TEE in other risk factors, such as community-associated disease, intravenous drug users, and hemodialysis dependence. Other remaining issues that need to be defined include the optimal time for echocardiography and the cost-benefit of TTE before TEE.

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