# HOSPITAL TOOLKIT

for Adult Sepsis Surveillance







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### **LIST OF ABBREVIATIONS**

ASE	adult sepsis event	ICU	intensive care unit
BSE	bacteremia/fungemia shock event	IV	intravenous
CDC	Centers for Disease Control and Prevention	NHSN	National Healthcare Safety Network
СО	community-onset	QAD	qualifying antimicrobial days
CPT	current procedural terminology code	РО	oral administration
eGFR	estimated glomerular filtration rate	RIT	repeat infection timeframe
ED	emergency department	SEP-1	Centers for Medicare & Medicaid Services
EHR	electronic health record		Early Management Bundle, Severe Sepsis/Septic Shock
НО	hospital-onset	SOFA	Sequential Organ Failure Assessment
ICD-10	10th Revision of the International Statistical Classification of Diseases and Related Health Problems		



### **BACKGROUND**

Sepsis is a clinical syndrome defined as life-threatening organ dysfunction caused by a dysregulated host response to infection.<sup>1</sup> A study using clinical data estimated 1.7 million cases of sepsis in 2014 in the U.S., which likely contribute to over 270,000 deaths.<sup>2</sup> Patients who survive sepsis often suffer long-term physical, psychological, and cognitive disabilities.

Because there is no confirmatory diagnostic test, the diagnosis of sepsis requires clinical judgment based on evidence of infection and organ dysfunction. A 1991 consensus conference established a clinical definition based on the patient's systemic inflammatory response syndrome (SIRS) to infection (later referred to as Sepsis-1)³, and these clinical criteria were expanded in 2001 in the Sepsis-2 criteria.⁴ In response to increasing understanding of sepsis pathobiology, a task force updated the clinical definitions in 2016¹, and in Sepsis-3, defined sepsis as "life-threatening organ dysfunction caused by a dysregulated host response to infection", with clinical guidelines defining organ dysfunction as acute change in total Sequential Organ Failure Assessment (SOFA) score ≥2 points consequent to the infection.¹

The updated clinical definitions are critical for identifying patients at risk for sepsis and further complications, but there is also a need to have a definition that can be used to track sepsis. Different definitions for sepsis are needed for different purposes, which could include clinical care, research, surveillance, and quality improvement and audit.<sup>5,6</sup> For example, a sepsis definition optimized for public health surveillance would prioritize reliability and validity across healthcare facilities, and low measurement burden. However timeliness may be less of a priority because such definitions would be used retrospectively and not be intended for clinical management of individual patients.<sup>5,6</sup>

Prior to 2017, U.S. national estimates of sepsis burden primarily relied on the use of administrative codes, which have consistently demonstrated increasing incidence and decreasing mortality. Administrative codes are also frequently used in sepsis quality initiatives, including Centers for Medicare & Medicaid Services Early Management Bundle, Severe Sepsis/Septic Shock (SEP-1). However, studies have demonstrated that coding for sepsis has steadily increased over the past decade, yet coding for the most common underlying infections has been stable or decreasing. These analyses demonstrate that coding practices are likely vulnerable to biases from increasing sepsis awareness and financial incentives (higher reimbursement for sepsis coding), and therefore unreliable for surveillance purposes.

In 2017, a CDC Prevention Epicenters-funded consortium published results of a study which used a new definition for sepsis based on objective clinical data elements conceptually analogous to Sepsis-3. This definition was optimized for surveillance directly from electronic health records across over 400 facilities, and displayed superior sensitivity and similar specificity compared to administrative codes when using Sepsis-3 criteria determined by medical record reviews as a gold standard. Furthermore, this definition demonstrated that national sepsis incidence and outcomes (combination of death and discharge to hospice) were stable from 2009-2014, in contrast to administrative codes which showed increasing incidence and decreasing mortality, but are fundamentally confounded by increasing sepsis awareness, coding bias, and financial incentives.

### **OBJECTIVES**

This toolkit allows healthcare professionals who are interested in using the sepsis surveillance methodology from the national burden study to track healthcare facility-level sepsis incidence and outcomes using an objective definition based on clinical data. Necessary data may be obtained and processed directly from an electronic health record, but could also be obtained using manual chart review. These data may be useful for understanding the effectiveness of local sepsis prevention, early recognition, and treatment programs.

### **SURVEILLANCE SETTINGS AND APPROPRIATE USE**

These sepsis definitions were developed for surveillance of adult patents in acute care hospitals where denominator data (total admissions or patient days) can be collected. Because this toolkit features sepsis definitions not subject to coding bias or variation in provider documentation, it may be helpful for objectively tracking sepsis incidence and outcomes within a facility for the purpose of quality improvement initiatives or public health and epidemiology.

It is not recommended to use these definitions for sepsis surveillance among pediatric patients, since these surveillance definitions were developed to mirror adult sepsis definitions and have not been validated in pediatric populations.

The following sepsis definitions are not intended to guide clinical diagnostic or therapeutic decision making. The antibiotic prescribing requirements to meet a sepsis case imply that patient can only meet one of the sepsis surveillance definitions after at least 4 days of antibiotic therapy, and therefore the determination of sepsis with these definitions is retrospective.

Hospitals serve a variety of different patient populations that vary by factors that influence patient outcomes, including sepsis risk factors, comorbidities, and acuity of illness. Therefore direct comparison of sepsis outcomes among hospitals may not reflect quality of sepsis care. Rigorous comparisons of sepsis outcomes for regional, state, or national ranking purposes would require additional research to create validated risk adjustment models that account for many patient and hospital-level factors. However, despite these limitations, these standardized surveillance definitions could be useful for comparing sepsis outcomes within the same hospital over time or to a peer institution serving a similar population.

### **DEFINITIONS**

### Two sepsis events definitions are presented.

The **Adult Sepsis Event (ASE)** captures the widest variety of sepsis patients. This definition was developed and tested in a study by Rhee, *et al.* which included more than 400 U.S. hospitals using a wide variety of electronic health records (EHR).<sup>2</sup> ASE surveillance requires access to microbiology, medication administration, laboratory, and administrative (ICD) coding data. Clinical records of mechanical ventilation may also be used in lieu of procedure codes.

This definition was validated by Rhee, *et al.* and shown to be present in 6% of hospital admissions, with a sensitivity of 69.7% (95% confidence interval [CI] 52.9% to 92.0%), 98.1% specificity (95% CI 97.7% to 98.5%), 70.4% positive predictive value (95% CI 64.0% to 78.8%), and 98.0% negative predictive value (95% CI 95.9% to 99.6%) when using Sepsis-3 criteria as the reference standard.<sup>2</sup>

There is no single diagnostic test or comprehensive clinical criteria for sepsis, and thus it is important to note the following limitations of this surveillance definition, which has been optimized for objectivity and standardization:

- While ASE is not subject to provider variation in documentation and coding of sepsis, it is still dependent on provider-initiated interventions (ordering blood cultures, antimicrobials, laboratory tests, vasopressors or mechanical ventilation) to measure presumed infection and organ dysfunction. This is also a limitation of the Sepsis-3 definition.
- It is not possible to determine whether organ dysfunction was a result of infection or other causes when using EHR-based automated determination of ASE. This can lead to misclassification.
- ASE criteria will miss organ dysfunction due to mild or moderate hypoxia that does not result in mechanical ventilation but would fulfill organ dysfunction criteria in the Sepsis-3 clinical criteria.

The **Bacteremia/Fungemia Shock Event (BSE)** is a simplified definition that tracks a narrower sub-population of patients with a much higher mortality. This definition is based on positive blood cultures (excluding common contaminants), which almost always reflect true infections, and vasopressors, which almost always reflect clinically important hypotension. BSE requires the fewest data elements, and can be determined using only microbiology and medication administration records.

In subsequent analysis of the dataset used by Rhee *et al.*, BSE was present in 0.48% of hospital admissions (compared to 6.0% with ASE), and 35.4% of BSE events resulted in death during hospitalization (compared to 15% of ASE). It had a sensitivity of 6.0% and positive predictive value of 100% when using Sepsis-3 criteria as the reference standard (unpublished).

### **ASE: Adult Sepsis Event**

(Must include the 2 components of criteria A AND include one or more organ dysfunction listed among B criteria)

- **A. Presumed Infection** (presence of both 1 and 2):
- 1. Blood culture obtained (irrespective of the result), AND
- 2. At least 4 Qualifying Antimicrobial Days (QAD) starting within the time period 2 calendar days before and after the collection date of a blood culture. See below.

### **AND**

- **B.** <u>Organ Dysfunction</u> (at least 1 of following criteria met within the time period 2 calendar days before and after the collection date of a blood culture.):
- 1. Initiation of a new vasopressor infusion (norepinephrine, dopamine, epinephrine, phenylephrine, OR vasopressin). To count as a new vasopressor, that specific vasopressor cannot have been administered in the prior calendar day. See Appendix B.
- 2. Initiation of invasive mechanical ventilation (must be greater than 1 calendar day between mechanical ventilation episodes). Invasive mechanical ventilation can be identified by ICD-10 Procedure Codes (5A1935Z, 5A1945Z, 5A1955Z) or CPT codes (94002, 94003, 94004, 94656, 94657), or by other clinical records.
- 3. Doubling of serum creatinine OR decrease by ≥50% of estimated glomerular filtration rate (eGFR) relative to baseline (see below), excluding patients with ICD-10 code for end-stage renal disease (N18.6). (If eGFR values are not readily available, creatinine alone can be used to determine renal dysfunction).
- 4. Total bilirubin ≥ 2.0 mg/dL and increase by 100% from baseline (see below).
- Platelet count <100 cells/µL AND ≥ 50% decline from baseline (see below) baseline must be ≥100 cells/µL.</li>
- 6. Optional: Serum lactate ≥ 2.0 mmol/L, note that serum lactate has become an increasingly common test to measure tissue perfusion. When serum lactate is included in the surveillance definition, the likely effect will be to slightly increase the number of sepsis cases identified. However, if serum lactate ordering practices are not stable over time in a particular hospital this will bias the incidence of sepsis. For this reason, serum lactate was not used in the primary analysis of sepsis trends over time in the original study by Rhee et al.

### **BSE: Bacteremia/Fungemia Shock Event**

(Must meet BOTH criteria A AND B)

### A. Bacteremia OR Fungemia

Patient has a recognized pathogen identified (i.e., an organism which is not a common commensal – see CDC National Healthcare Safety Network list for guidance) from one or more blood specimens by a culture or non-culture based microbiologic testing method.

### **AND**

### **B.** New Vasopressor

Initiation of a new vasopressor (norepinephrine, dopamine, epinephrine, phenylephrine, vasopressin) within the time period 2 calendar days before and after the collection date of a blood culture demonstrating a recognized pathogen (see A). To count as a new vasopressor, that specific vasopressor cannot have been administered in the prior calendar day.

### **Key Terms and Concepts**

### **Blood Culture**

Qualifying cultures include those drawn for bacterial (aerobic and/or anaerobic), acid-fast bacilli (AFB), and fungal cultures. Blood cultures for specific viruses (e.g., cytomegalovirus) are excluded. For ASE, blood cultures merely need to have been drawn, regardless of result. For BSE, blood cultures or other bacteremia or fungemia testing of blood must have yielded a recognized pathogen.

### Window Period

The date the blood culture is obtained is the center of a window period extending both 2 days before and 2 days after the blood culture for both ASE and BSE. Multiple window periods during a hospitalization are possible. If multiple blood cultures are obtained in a short period of time, window periods may overlap.

Figure 1: Examples Illustrating Window Period

Hospital Day No.	1	2	3	4	5	6	7	8	9
Blood Culture				Х					
Window Period			Wi	ndow Per	iod				
<b>Hospital Day No.</b>	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>	Α						Α		
Window Period	Win	dow Peri	od A			Win	dow Peri	od B	
Hospital Day No.	1	2	3	4	5	6	7	8	9
Blood Culture	Α			В					
Window Period	Overlapping Window Periods								

If blood cultures, antimicrobial use, OR any organ dysfunction criteria are obtained on hospital day -1 or day 0 (i.e., up to 2 days prior to admission), these criteria can be used to qualify for events in order to account for medical care in the emergency department prior to admission.

### Qualifying Antimicrobial Days (QAD)

For ASE events, the first QAD is the first day in window period extending both 2 days before and 2 days after the patient receives a new antimicrobial (see Appendix A for list of qualifying antimicrobials). A new antimicrobial is defined as an antimicrobial not previously administered in the prior 2 calendar days. Oral and intravenous formulations of the same antimicrobial are counted as the same antimicrobial EXCEPT for vancomycin.

Subsequent QADs can be the same antimicrobial, or a different antimicrobial as long as the first dose of each antimicrobial in the sequence is new (not previously administered in the prior 2 calendar days). A new antimicrobial does not have to be started within the window period to be counted as a QAD.

**Figure 2: Examples Illustrating QAD** 

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Χ					
Window Period			Wi	ndow Per	iod				
Levofloxacin IV administration				X	Х	Х	Х		
QAD				Х	Х	Х	Х	Х	

This scenario includes at least 4 QAD that begin within the window period, and therefore meets criteria for presumed infection.

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Х					
Window Period			Wi	ndow Per	iod				
Vancomycin IV administration				X	Х	Х	Х	Х	
Piperacillin/Tazobactam IV administration	Х	Х	х	Х	Х				
QAD				1	2	3	4	5	

This scenario includes at least 4 QAD from Vancomycin that begin within the window period, and therefore meets criteria for presumed infection. Because Piperacillin/Tazobactam administration was administered prior to the window period, it does not qualify towards QADs.

There must be at least one new parenteral (intravenous or intramuscular) antimicrobial administered within the window period for the QADs to satisfy the definition.

Figure 3: Examples Illustrating QAD with Multiple Antimicrobials

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Х					
Window Period			Wii	ndow Per	iod				
Vancomycin IV administration				Х	Х	Х			
Piperacillin/Tazobactam IV administration				Х	Х	Х			
Levofloxacin Oral administration						х	Х	Х	
QAD				1	2	3	4	5	

In this scenario, the first 3 QADs were met by vancomycin IV administration and piperacillin/tazobactam IV administration. QAD 4 and QAD 5 were met by levofloxacin oral administration. Subsequent QADs may be oral as long as at least one QAD in the window period is parenteral. This scenario qualifies for presumed infection.

A gap of a single calendar day between administrations of the same antibiotic (oral or intravenous) count as QADs as long as the gap is not greater than 1 day.

Figure 4: Examples Illustrating QAD with Gaps in Antimicrobial Dosing

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>	Χ								
Window Period	Wi	ndow Peri	iod						
Levofloxacin IV administration	х		х		Х				
QAD	1	2	3	4	5				

Even though levofloxacin is administered every other day, because the gap between doses is only 1 day, this levofloxacin administration counts as consecutive treatment doses, for a total of at least 4 QAD that begin within the window period, and therefore meets criteria for presumed infection.

If a patient's care transitions to comfort measures only, or the patient dies, is discharged to another hospital, or discharged to hospice before 4 QADs have elapsed, then the presumed infection criteria can be met with less than 4 QADs as long as they have consecutive QADs until day of, or 1 day prior to, death or discharge. For a patient only seen in the ED or ED observation unit, a QAD is required each day until day of, or 1 day prior to, death.

QAD does not apply to BSE events since administration of antibiotics is not required to meet the definition.

Figure 5: Example Illustrating QAD with Patient Death

Hospital Day No.	1	2	3	4	5	6	7
<b>Blood Culture</b>				Χ	Χ		
Window Period				Windov	v Period		
Levofloxacin IV administration					Х	х	
QAD					1	2	
Death							X

Even though only 2 QADs had elapsed before patient death, this scenario counts as an event since there were consecutive QADs until the day prior to death.

### **Onset Date**

For ASE, onset date is defined as the earliest day in the window period extending both 2 days before and 2 days after the blood culture when EITHER the blood culture, first QAD, OR organ dysfunction criteria is met. For BSE, onset date is defined as the earliest day in the window period extending both 2 days before and 2 days after the blood culture when the vasopressor infusion was started.

Figure 6: Examples Illustrating Onset Date

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Х					
Window Period			Wi	ndow Peri	od				
QAD				1	2	3	4	5	
Organ Dysfunction			Х						
Onset Date			Х						

Organ dysfunction takes place on hospital day 3, while blood cultures and QADs begin on hospital-day 4. Because onset date is the date of the first criteria met (in this case organ dysfunction), onset date is hospital day 3.

### Community-Onset vs. Hospital-Onset Events

Hospital-Onset Events require onset date to be on hospital day 3 or later, counting the date of admission as hospital day 1. Community-Onset Events require onset date to be on hospital day 2 or earlier, when the date of admission counts as hospital day 1.

Figure 7: Examples Illustrating Community-Onset vs. Hospital-Onset Events

Hospital Day No.	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Χ					
Window Period			Wi	ndow Peri	iod				
QAD				1	2	3	4	5	
Organ Disfunction			Х						
Onset Date			Х						

Because Onset Date is on hospital day 3, this example qualifies as a Hospital-Onset Event

<b>Hospital Day No.</b>	1	2	3	4	5	6	7	8	9
<b>Blood Culture</b>				Χ					
Window Period			Wi	ndow Peri	iod				
QAD				1	2	3	4	5	
Organ Disfunction			Х						
Onset Date		Х							

In this similar scenario, since the Organ Dysfunction was on hospital day 2, the Onset Date is also hospital day 2, and therefore this example qualifies as a Community-Onset Event.

### **Baseline Organ Function**

### **Community-Onset Events**

- Creatinine baseline is the lowest value during hospitalization.
- Estimated Glomerular Filtration Rate (eGFR) baseline is the highest value during the hospitalization.
   If eGFR value is documented as "≥60", eGFR is treated as 60.
- Bilirubin baseline is the lowest value during hospitalization.
- Platelet baseline is the highest value during hospitalization. If all platelet counts during hospitalization are <100 cells/µL, then platelet count should not be used to satisfy organ dysfunction criteria.

### Hospital-Onset Infection Events

- Creatinine baseline is the lowest value within the window period extending both 2 days before and 2 days after the date of blood culture collection.
- Estimated Glomerular Filtration Rate (eGFR) baseline is the highest value within the window period extending both 2 days before and 2 days after the date of blood culture collection. If eGFR value is documented as "≥60", eGFR is treated as 60.
- Bilirubin baseline is the lowest value within the window period extending both 2 days before and 2 days after the date of blood culture collection.
- Platelet baseline is the highest value within the window period extending both 2 days before and 2 days after the date of blood culture collection, AND must be ≥100 cells/µL. If all platelet counts within window period are <100 cells/µL, then platelet count should not be used to satisfy organ dysfunction criteria.</p>

### Repeat Infection Timeframe Considerations

The repeat infection timeframe (RIT) is a timeframe after an ASE or BSE onset date when no new events are counted, in order to minimize the chance a single, prolonged episode of ASE or BSE is counted twice. RIT therefore only applies to determination of hospital-onset events.

Note that in Rhee et al., only one sepsis episode was counted per hospitalization, and thus no RIT was used.

Among healthcare associated infections reported to the CDC's National Healthcare Safety Network, an RIT of 14 days is used, and may be reasonable for hospitals looking to track hospital-onset events. However, other RIT timeframes (or simply counting a single sepsis episode per admission) may be more appropriate depending on the hospital's surveillance needs.

Use of an RIT is probably most useful in surveillance efforts where sepsis incidence is of primary interest. However, if mortality outcomes are the primary interest, then it may be prudent to eliminate the RIT since a patient with multiple sepsis episodes and a single hospitalization outcome (survival, death, or discharge to hospice) would be counted multiple times and thus skew results.

### **SURVEILLANCE METHODS**

### Pre-assessment of Sepsis Identification, Care and Surveillance Practices (Optional)

Prior to beginning sepsis surveillance efforts, it would be prudent to investigate sepsis diagnostic and treatment practices across the hospital, as well as administrative programs. This is especially important if this toolkit will be used to track quality improvement efforts. A pre-assessment of broad sepsis practices would be helpful for interpreting surveillance results and creating recommendations for improving patient care.

Existing sepsis practices to review may include:

### Administrative and Programmatic Practices

- Sepsis CMS SEP-1 compliance efforts
- Emergency department sepsis teams
- Hospital ward or intensive care unit sepsis teams
- Presence of "Code Sepsis" teams
- Electronic health record sepsis prediction, alerts, or protocols
- Trends in sepsis ICD coding

### Clinical Practices

- Emergency department sepsis identification and triage practices
- Hospital ward or intensive care unit sepsis screening practices
- Presence and compliance with sepsis diagnostic and management protocols
- Concurrent clinical trials or quality improvement initiatives
- Referral/Transfer of patients with sepsis to other hospitals



### **Electronic Case Finding**

Electronic data collection will need to be planned and implemented with the hospital's information technology department, or the clinical data warehouse, if available. Detailed data specifications are provided in Appendix C. A summary of the data elements is below:

### The following categories of data are needed to determine ASE:

- Microbiology data- blood and fungal culture collection date
- Medication administration records- antimicrobial and vasopressor administration
- Laboratory data- serum lactate, serum creatinine, total bilirubin, platelet count
- ICD-10 codes- end-stage renal disease (N18.6), mechanical ventilation (5A1935Z, 5A1945Z, 5A1955Z, OR may use other clinical documentation of mechanical ventilation, see below)
- CPT codes- indicating mechanical ventilation: (94002, 94003, 94004, 94656, 94657)
- Patient outcome- death during hospitalization, discharge to hospice, transfer to another acute care hospital
- (Optional) Clinical documentation of receipt of mechanical ventilation- (in lieu of mechanical ventilation procedure codes above)

### The following categories of data are needed to determine BSE:

- Microbiology data-blood and fungal culture collection date
- Medication administration records-vasopressor administration by infusion

**Optional:** In addition to the data necessary to determine the presence of sepsis events, additional information could be collected to aid in the epidemiologic analysis and understanding of patients with sepsis. This include but are not limited to the following data categories:

- Patient demographics
- Comorbidities-from either patient problem lists or from ICD-10 codes
- Patient location within the hospital
- Additional microbiological culture data: including pathogen identification from blood culture, non-blood culture testing and diagnostic results
- Record of surgeries, procedures, and indwelling devices such as central venous catheters
- Additional patient outcomes: discharge to home, discharge to nursing home or rehabilitation
- Cost/Billing

### **Manual Case Finding**

An Adult Sepsis Event worksheet (Appendix D) is provided to aid collection of appropriate data for each ASE case that is identified. A Bacteremia/Fungemia Shock Event Worksheet (Appendix E) is also provided to aid collection of appropriate data for each BSE case that is identified. Both templates are provided in a modifiable format and can be customized to include additional data of interest. Of note, these forms include the option of specifying the causative organism and antimicrobial sensitivities similar to healthcare-associated infections reported to the CDC's National Healthcare Safety Network, however these fields are optional.

### **Denominator Data Collection**

For community-onset sepsis, several denominator options are available depending on the outcome measure of interest.

**Table 1: Suggested Community-onset (CO) Sepsis Outcome Measures** 

Outcome Measure	Numerator	Denominator Data
Mortality Rate	Patients with CO Sepsis events (ASE or BSE) who either died during hospitalization or were discharged to hospice	Total CO sepsis events (ASE or BSE)
Sepsis Rate	Total CO sepsis events (ASE or BSE)	Total hospital admissions (inpatient AND observation/24 hour stays)

For hospital-onset sepsis, several denominator options are available depending on the outcome measure of interest.

**Table 2: Suggested Hospital-onset Sepsis Outcome Measures** 

Outcome Measure	Numerator	Denominator Data			
Mortality Rate	Patients with HO Sepsis events (ASE or BSE) who either died during hospitalization or were discharged to hospice	Total HO sepsis events (ASE or BSE)			
Sepsis Rate	Total HO sepsis events (ASE or BSE)	Total hospital admissions (inpatient AND observation/24 hour stays)			

Data regarding the total number of hospital admissions is often obtained from hospital administrative databases and vary by institution.

### **Data Analysis and Quality Assessment**

A timeframe should be chosen for each sepsis event calculation. Months, quarters, and years are common time units for analysis of many healthcare surveillance and quality measures. The optimal choice of time-unit will vary by facility and volume of cases.

Onset	Outcome	Calculation	Notes
Community-Onset (CO)	CO event rate per 100 admissions	(Total CO events)  ————— × 100 Total Admissions	
	CO Mortality Rate per 100 CO admissions	(CO events with in-hospital deaths + CO events with discharge to hospice)  ———————————————————————————————————	May reflect results of care in multiple settings (emergency room, intensive care unit), thus may not bea reliable measure of care in a single unit
Hospital-Onset (HO)	HO event rate per 1000 patient days	(CO events)  ———————————————————————————————————	
	HO Mortality Rate per 100 events	(HO events with in-hospital deaths + HO events with discharge to hospice)  ———————————————————————————————————	

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### **APPENDICES**

### Appendix A. Antimicrobials Qualifying for Adult Sepsis Event

For purposes of case finding criteria, the antimicrobials were divided into parenteral (IV) and oral (PO) antimicrobials (where antimicrobial refers to antibacterial, antifungal, or antiviral agents). All PO and IV antimicrobials were considered identical for purposes of determining whether an antimicrobial is "new" or not (meaning that a switch from IV to PO or vice versa does not count as a "new antimicrobial.") The one exception is IV vs PO Vancomycin (meaning that a switch from IV to PO vancomycin, or initiation of PO vancomycin while still on IV vancomycin, and vice versa, all count as "new" antimicrobials). Intramuscular (IM) antimicrobials were treated equivalently as IV antimicrobials. The generic names of the antimicrobials included are listed below. This list represents antibmicrobials commercially available in the U.S. in 2017, and users should update the list as new antimicrobials are introduced into clinical care.

Antimicrobial									
	IV Antibacterials								
amikacin ampicillin ampicillin/ sulbactam azithromycin aztreonam cefamandole cefazolin cefepime cefmetazole cefonicid cefoperazone cefotaxime cefotetan	cefotetan cefoxitin ceftaroline ceftazidime ceftazidime/ avibactam ceftizoxime ceftolozane/ tazobactam ceftriaxone cefuroxime cephalothin cephapirin chloramphenicol	ciprofloxacin clindamycin cloxacillin colistin dalbavancin daptomycin doripenem doxycycline ertapenem gatifloxacin gentamicin mipenem kanamycin	evofloxacin lincomycin linezolid, meropenem/ vaborbactam linezolid meropenem/ vaborbactam methicillin metronidazole mezlocillin minocycline	moxifloxacin minocycline moxifloxacin nafcillin oritavancin oxacillin penicillin piperacillin piperacillin/ tazobactam polymyxin B quinupristin/ dalfopristin streptomycin	tedizolid telavancin ticarcillin ticarcillin/ clavulanate tigecycline tobramycin trimethoprim/ sulfamethoxazole vancomycin				
		PO Antil	bacterials						
amoxicillin/ clavulanate amoxicillin ampicillin azithromycin cefaclor cefadroxil cefdinir cefditoren cefixime	cefpodoxime cefprozil ceftibuten cefuroxime cephalexin cephradine cefixime cefpodoxime cefprozil ceftibuten	cefuroxime cephalexin cephradine chloramphenicol cinoxacin ciprofloxacin clarithromycin clindamycin cloxacillin dicloxacillin	doxycycline fidaxomicin fosfomycin gatifloxacin levofloxacin lincomycin linezolid metronidazole lincomycin linezolid	metronidazole minocycline moxifloxacin nitrofurantoin norfloxacin ofloxacin penicillin pivampicillin rifampin sulfadiazine	sulfadiazine- trimethoprim sulfamethoxazole sulfisoxazole tedizolid telithromycin tetracycline trimethoprim trimethoprim/ sulfamethoxazole vancomycin				
		IV Ant	ifungals						
amphotericin B anidulafungin	caspofungin fluconazole	isavuconazonium itraconazole	micafungin	posaconazole	voriconazole				
		PO Ant	ifungals						
fluconazole	isavuconazoniu	itraconazole	posaconazole	voriconazole					
			tivirals						
acyclovir	ganciclovir	cidofovir	foscarnet	peramivir					
L		PO An	ntivirals						
oseltamivir									

### Appendix B: Vasopressors Included in Adult Sepsis Event Definition

Eligible vasopressors must have been administered via continuous intravenous infusion. Vasopressors administered in an operating room are excluded as these are frequently needed to counteract hypotension induced by sedative medication administration. Since the location of administration may be challenging to identify in an EHR, single bolus injections of vasopressors (a frequent method of delivering perioperative vasopressors) are generally excluded. The following medications were included:

Norepinephrine

Vasopressin

Epinephrine

Phenylephrine

Dopamine

### Appendix C: Data Specifications for Direct Sepsis Determination from Electronic Health Records

The following data specifications were adopted from the study by Rhee et al. to assist hospitals with extracting, cleaning, and organizing data during the original study. Some updates have been made to reflect updates in ICD coding, streamlining, or other improvements in the surveillance definition.

Use of a common specification minimized variability in data across hospitals and facilitated analysis across multiple hospitals using a common analytic code. Use of this data specification is optional, but may be useful if your hospital wishes to compare unadjusted data to other hospitals using this toolkit.

### **Population:**

All adult patients (≥18 years old) who were hospitalized, or died in the Emergency Department (ED).

The "denominator" of hospitalized patients includes:

- 1. Inpatient hospitalizations
- 2. "Observation" or "24 Hour" stays (we are also counting this functionally the same as "Inpatient")

Different EHR systems may have different ways of identifying hospitalizations. Possible strategies include looking for patients with inpatient encounter flags or encounters with valid DRG codes. Some datasets include encounters without clinical data (shell encounters) – for this protocol each encounter should have some minimum clinical data to be an eligible encounter (i.e. at least one laboratory procedure, at least one medication, and possibly at least one diagnosis code).

The "numerator" definitions for counts of septic patients per different criteria are summarized in the toolkit (page 12). In terms of which patients are eligible for inclusion in the numerator please note the following:

1. Include patients with evidence of sepsis who died in the ED. (If a patient otherwise meets criteria for this EHR sepsis definition, i.e. IV antibiotic, blood culture, and organ dysfunction) and dies in the ED, the patient would count as a case. (These will not be included in the denominator ("per 100 hospitalizations"); however, it is expected that the number of sepsis patients who die in the ED will be extremely low.

For the purposes of generating denominator counts, use a dataset that include all inpatient encounters. For numerators (i.e. cases), you can limit the size of the patient level analytic dataset in order to decrease file size and increase efficiency. In particular, the analytic dataset for numerators can be limited to the following way:

### 1. EHR Clinical Data Starting Points (use either a. or b.):

a. All inpatients who received a **systemic antibiotic**, **antifungal**, or **antiviral** (PO or IV) for ≥1 calendar day. If a patient was only seen in the ED and was never admitted, but received antibiotics, they should be included if they died in the ED. Please see appendix for a list of antibiotics, antifungals, and antivirals included in this definition.

### OR

**b.** All inpatients who had **at least one blood cultures obtained.** If a patient was only seen in the ED and was never admitted, but had a blood culture obtained, they should be included if they died in the ED.

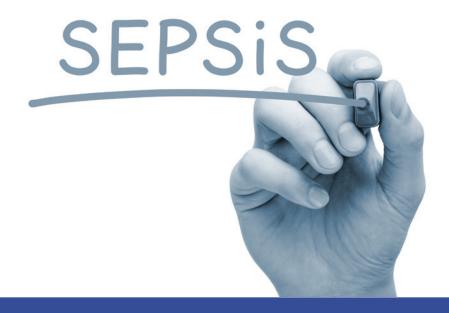
For all patients, data should be provided for their ED stay (if applicable) as well as their inpatient hospitalization. This means that some patients may have date for a "day 0" or "day 1", since day 1 will refer to the inpatient admission date.

### **Notes:**

- For convention, label the day of admission as "day 1".
- Change all dates from actual dates to relative dates. Relative dates are defined relative to date of admission. For example, if a patient is admitted on Monday, and discharged on Wednesday, discharge day would = 3.

### **Overview of Tables (Files)**

Table Name	Description of Contents
Basic	Demographics and basic hospital summary characteristics (i.e. admission source, LOS, discharge disposition)
Diagnosis_Codes	All diagnosis codes and DRGs
Procedure_Codes	All procedure codes
Medications	Medications (Emergency Department and Inpatient meds only)
Laboratory	Laboratory values
Blood_culture	Blood culture dates (and results, if available)
Vent	Dates of mechanical ventilation
Location	Location of patient on each hospitalization day (e.g. ER, ward, ICU)
Hospital	Summary statistics on each hospital



### I. Basic

Data Field	Description	Output		
Patient_ID	A unique dummy ID that can identify the same patient across different hospitalizations	Number		
Admission_ID	A unique identifier for every unique hospitalization  *Note –this should be unique for each hospitalization, i.e. even if the same patient is hospitalized again later in the year, they should have another unique_ID	Number		
Hospital_ID	Unique Hospital Identifier	Number		
Age	Age in years on Day 0	Whole Number		
Age_months	Age in months if age in years is 2 or less	Whole Number		
Gender		M or F		
Race	*Note, Hispanic is often listed as a race. If listed separately as Ethnicity, this takes precedence over any other race. i.e. White Race + Hispanic Ethnicity = Hispanic	White Black Asian Hispanic Other .=Not Identified / Missing		
Admit_source	Admission Source (if available)	1=ER from community 2=Direct Admit to ward or ICU 3=OSH Transfer to ER 4=OSH Transfer to ward or ICU 5=Newborn 6=Other		
Prior_location	Residence prior to hospitalization (if available)	1= Home 2= Long-term facility		
Discharge_date	Date of hospital discharge, relative to admission	Integer Number		
Discharge dispo	Discharge disposition	1=Home 2=Transfer to Another Acute Care Hospital 3=DC to Subacute Facility 4=Death 5=Hospice		
Year	Calendar Year of Hospitalization (Based on Date of Admission)	4-digit year		
Season	Season of Hospitalization (Based on Date of Admission)	Winter (December, January, February) Spring (March, April, May) Summer (June, July, August) Fall (September, October, November)		
ICU_admitdate	Date of first admission to ICU (relative to admission date)	Whole Number		
ICU_LOS	Total number of calendar days in the ICU	Whole number (partial days counted as whole day)		
Service_line	A Primary Service Line for Admission *There may be multiple ways to impute this. Specialty of admitting physician is probably best; if not available, specialty of discharging physician.	Medical Surgical Cardiac Oncology Obstetrics Neuro Other *See next page for more details about classification		

- Obstetrics includes maternal and fetal medicine
- Surgery includes: Surgical critical care, Anesthesiology, Cardiac Surgery, Neurosurgery, gynecology, gynecologic oncology, Surgical oncology, Ophthalmology, Otolaryngology, Urology, Podiatry, Oral Surgery
- Cardiac includes Cardiology and Cardiac Critical Care
- Neurology includes Neuro critical care
- All other types of critical care (Non-surgical/Neuro/Cardiac) is Medical
- Other includes Emergency Medicine, Radiology, Pathology, Dermatology, and anything else

### **Example Format for Basic File:**

Patient_ID	Admission _ID	Hospital _ID	Age	Gender	Race	Admit_ Source	Discharge _Date	ETC.
12345678	1234567890	100	65	М	White	1	7	
22221234	1234567891	105	53	F	Asian	2	10	

### II. Diagnosis\_Codes

This file should include admitting diagnosis, primary diagnosis, all discharge diagnoses, the sequence number (i.e. numerical order of the code) and DRG codes. If present-on-admission (POA) flags are available for each diagnosis, please include those as well.

Data Field	Description
Diagnosis_Codes	Diagnosis code – either ICD-10 or DRG. No decimal points
CodeType	DRG or ICD10
Sequence No	Sequence of codes at discharge (for ICD-10 codes).  1=Primary diagnosis, 2 and above = all secondary diagnosis
POA	Present on admission flag (for ICD-10 codes) $-1 = yes$ , $0 = No$
Primary_Dx	Primary (or Principal) Diagnosis at discharge.
Admit_Dx	Admitting Diagnosis (if available)

### The "Long" Format (one diagnosis per line) is preferred:

Patient_ID	Admission _ID	Diagnosis Code	Code Type	Sequence No	POA	Primary Dx	Admit Dx
12345678	1234567890	R6520	ICD10	1	1	1	1
22221234	1234567891	020	DRG				
22221234	1234567891	R6521	ICD10	1	1	1	1
22221234	1234567891	1509	ICD10	2	0	0	0

### III. Procedure\_Codes

This file should include all procedure codes (CPT and ICD10), the day relative to admission on which the procedure occurred, and the procedure specialty code (if available; this refers to Medicare provider/supplier specialty codes). The "Long" format is preferred (one line per procedure).

### **Example Format for Procedure\_Codes:**

Patient_ID	Admission _ID	Procedure _Code	ICD10 or CPT	Day	Proc_Specialty _Code
12345678	1234567890	5A1935Z	ICD10	1	99
22221234	1234567891	5A1935Z	ICD10	3	A3
22221234	1234567891	00100	CPT	3	A3

### **IV. Medications**

The medications we are interested in are:

- 1. Vasopressors = IV norepinephrine, epinephrine, phenylephrine, dopamine, and vasopressin
- $\textbf{2. Systemic Antibiotics} \ (\textbf{see Appendix C for list}) \ \textbf{including Antibacterials, Antifungals,} \ \textbf{and Antivirals}$

(see below for preferred format for cleaned Medications file)

Note: "PO" refers to any enteral route - PO, per NG Tube, per G tube, etc.

### **Format for Cleaned Med File:**

Patient_ID	Admission _ID	Day	Med	Route	Med Type
11111111	1234567890	0	Norepinephrine	IV	Vasopressors
11111111	1234567890	0	Vancomycin	IV	Antibacterial
11111111	1234567890	0	Cefepime	IV	Antibacterial
11111111	1234567890	1	Vasopressin	IV	Vasopressors
11111111	1234567890	1	Vancomycin	IV	Antibacterial
11111111	1234567890	2	Cefepime	IV	Antibacterial
2222222	1234567891	0	Ceftriaxone	IV	Antibacterial
2222222	1234567891	1	Ceftriaxone	IV	Antibacterial
2222222	1234567891	2	Ceftriaxone	PO	Antibacterial
2222222	1234567891	2	Oseltamivir	PO	Antiviral
2222222	1234567891	3	Caspofungin	IV	Antifungal

### V. Laboratory

We are interested in daily values of each of the laboratory values, in the units specified.

- Cr\_max and Cr\_min = daily min and max Creatinine (mg/dL)
- eGFR\_max and eGFR\_min = daily min and max estimated glomerular filtration rate (mL/min/1.73m2). This generally reported as part of the Basic Metabolic Panel along with the creatinine. Note that a lab reporting "Creatinine Clearance" can be used interchangeably with eGFR.
- Tbili\_max and Tbili\_min = daily min and max Total Bilirubin (mg/dL)
- Plt\_max and Plt\_min = daily min and max Platelets (109/L)
- Lactate\_max = daily max Lactate (mmol/L)

### Format for cleaned lab file:

Patient_ID	Admission _ID	Day	Cr _max	Cr _min	Tbili _max	Tbili _min	Pit _max	Plt _min	Lactate _max
12345678	1234567890	1	1.2	1.2	2.1	2.1	125	125	2.1
12345678	1234567890	2	1.4	1.3	2.5	2.5	110	90	1.9
12345678	1234567890	3	1.5	1.4			70	70	
12345678	1234567890	4	1.5	1.5	3.1	3.1	60	60	
22221234	1234567891	1	2.5	2.5			400	400	4.2
22221234	1234567891	2	2.7	2.7	3.6	3.6	248	200	3.6

### **VI. Blood Culture**

Provide list of all days when a blood culture was ordered, dates of results, and final microbiologic results. There are several ways of identifying a blood culture order, depending on what your EHR system captures, which may have different date stamps if they occur close to midnight:

- a. Date and Time the blood culture was ordered
- **b.** Date and Time the blood culture was drawn (same as time collected)
- c. Date and Time the blood culture was logged/received by micro lab

### Format for cleaned lab file:

Patient_ID	Admission _ID	Specimen _ID	Bcx _order _Day	Bcx _order _time	Bcx _drawn _day	Bcx _drawn _time	Bcx _log _day	Bcx _log _time	Result
22221234	1234567890	555555	1	01:00	0	01:15	0	01:45	No growth
22221234	1234567890	555556	3	05:15	3	05:20	3	05:35	Staphylococcus aureus
22221234	1234567890	555557	3	05:15	3	05:21	3	05:35	Staphylococcus aureus
12345678	1234567890	3333333	1	12:15	1	12:20	1	12:40	E.coli
12345678	1234567891	3333334	7	03:30	7	03:35	7	03:45	No growth

### VII. Vent

Create a clean table with yes/no vent exposure per day, where 1 = yes patient received mechanical ventilation and 0 = no, patient did not receive mechanical ventilation that day

Method of identifying mechanical ventilation may differ depending on your EHR data.

Although actual respiratory therapy data is optimal, identifying mechanical ventilation by ICD-10 and CPT procedure codes is our default option.

**ICD-10 Vent Codes:** 5A1935Z, 5A1945Z, 5A1955Z **CPT Vent Codes:** 94002, 94003, 94004, 94656, 94657

Patient_ID	Admission_ID	Day	Mech _vent**
12345678	1234567890	1	0
12345678	1234567890	2	0
12345678	1234567890	3	0
12345678	1234567890	4	1
12345678	1234567890	5	1
12345678	1234567890	6	0
22221234	1234567891	1	0
22221234	1234567891	2	1
22221234	1234567891	3	1
22221234	1234567891	4	1
22221234	1234567891	5	0

<sup>\*\*</sup>Mech\_vent refers to invasive mechanical ventilation.

Option 2. If only the date of initiation of mechanical ventilation is available, please include this.

Patient_ID	Admission_ID	Vent_Initiation_Day
12345678	1234567890	3
78123456	1234567890	2
78123456	1234567890	6

### **VIII. Location**

### Option 1. Create a day-by-day location guide for the patient.

Please list the highest level of care on a given day: ex: if a patient is admitted to an ICU on a given day from the hospital ward, please list ICU rather than ward. **Hierarchy = ICU > Stepdown Unit > Ward > ER.** (An alternative is to use a midnight census to define the patient location for the day.)

Patient_ID	Admission_ID	Day	Location
12345678	1234567890	0	ER
12345678	1234567890	1	Ward
12345678	1234567890	2	Ward
12345678	1234567890	3	ICU
12345678	1234567890	4	ICU
12345678	1234567890	5	ICU
12345678	1234567890	6	Ward
12345678	1234567890	7	Ward
12345678	1234567890	8	ICU
22221234	1234567891	1	Ward
22221234	1234567891	2	Ward
22221234	1234567891	3	Ward

Option 2. If day-by-day location is not available, please provide date of first ICU admission and total ICU length of stay in the Basic Info File (File #1).

### IX. Hospital

Please provide the following information about each of your hospitals:

Variable	Description	Description 2
Hospital_ID	Hospital identifier (can be linked to each patient via the Basic file)	
State	State	2-digit state code
	Urban vs Suburban vs Rural	1=Urban
Urban		2=Suburban
		3=Rural—
Beds	Total number of inpatient beds	
ICU_beds	Total number of ICU beds	
Teaching	Teaching status	1=Teaching hospital
reaching		2=Non-teaching hospital
Annual_discharges	Total number of adult discharges per year	

# Appendix D: Adult Sepsis Event (ASE) Manual Case Report Template (\* = optional)

Hospital ID:	Event #:	
Patient ID:	Social Security #:	
Secondary ID:	Medicare #:	
Patient Name, Last:	First:	_ Middle:
Gender: F M Other	Date of Birth:	
Ethnicity (Specify):	Race (Specify):	
Date Admitted to Hospital:		
Sepsis Onset Date:	*Location at Onset Date: _	
<b>Event Details</b> (must meet Criteria #1 AND #2)		
CRITERIA #1: Presumed Infection		
☐ Blood culture obtained AND		
$\square \ge 4$ Qualifying Antimicrobial Days (starting $\pm 2$ days of blood	culture day)	
CRITERIA #2: Organ Dysfunction: (Any one of the following	± 2 days of blood culture)	
☐ Initiation of a new vasopressor infusion <sup>1</sup>		
☐ Initiation of invasive mechanical ventilation <sup>2</sup>		
☐ Acute renal failure (only for patients without end-stage renal of	disease), defined as <b>EITHER</b> :	
a. Doubling of serum creatinine compared to baseline <sup>3</sup> ,	DR .	
b. Decrease in estimated glomerular filtration rate (eGFR)	by ≥ 50% compared to baselin	$e^4$
☐ Hyperbilirubinemia, defined as <b>BOTH</b> :		
<ul><li>a. Total bilirubin ≥ 2 mg/dL, AND</li></ul>		
<b>b.</b> Total bilirubin increase of $\geq$ 50% compared to baseline	5	
$\square$ Thrombocytopenia (only for patients with baseline platelet co	$punt > 100 cells/\mu L$ ) defined as <b>I</b>	BOTH:
a. Platelet count $< 100 \text{ cells/}\mu\text{L}$ , AND		
<b>b.</b> Decrease in platelet count $\geq$ 50% compared to baselin	e <sup>6</sup>	
$\square$ *Serum lactate ≥ 2 mg/dL <sup>7</sup>		
<sup>1</sup> Vasopressor must not have been administered in prior calendar day. Qualifyin dopamine, epinephrine, phenylephrine, vasopressin. Single vasopressor doses not quality.	or those given in an operating room o	
<ul> <li>Must be &gt; 1 calendar days after previous invasive mechanical ventilation epis</li> <li>Baseline creatinine defined as: 1) Community-onset: lowest value during hosp</li> </ul>		
value in $\pm 2$ days of blood culture	·	
$^4$ Baseline eGFR is defined as 1) Community-onset: highest value during hospit value in $\pm$ 2 days of blood culture	alization, 2) Hospital Onset- highest	
$^5$ Baseline total bilirubin defined as 1) Community-onset: lowest value during h value in $\pm2$ days of blood culture	ospitalization, 2) Hospital Onset- lowe	st
<sup>6</sup> If all platelet counts are < 100, then platelet count should not be used to satis platelets defined as 1) Community-onset: highest value during hospitalization, days of blood culture AND must be ≥ 100.		2

<sup>7</sup> Optional criteria

Pied: Yes □ No □ *Sepsis Contributed to Death: Yes □ No □					
Discharge or Death Date:					
*Discharge Location:					
☐ Private residence	☐ Acute care hospital				
☐ Long-term acute care hospital (LTACH)	☐ Long-term care/Skilled nursing facility (SNF)				
☐ Homeless	☐ Incarcerated ☐ Other				
*Pathogens Identified: Yes □ No □ */	f Yes, option to specify on next pages				

Pathogen #			Gram-po	sitive Org	anisms			
	Staphylococcus coagulase- negative (specify species if available):	VANC SIRN						
	Enterococcus faecium	<b>DAPTO</b> S NS N	<b>GENTHL</b> § SRN	<b>LNZ</b> SI R N	<b>VANC</b> SIRN			
	Enterococcus faecalis							
	<b>Enterococcus</b> <b>spp.</b> (Only those not identified to the species level)							
	Staphylococcus	CIPRO/LEVO/ MOXI SIRN	<b>CLIND</b> SIRN	<b>DAPTO</b> S NS N	DOXY/ MINO SIRN	<b>ERYTH</b> SIRN	<b>GENT</b> SIRN	LNZ SRN
	aureus	OX/CEFOX/ METH SIRN	<b>RIF</b> SIRN	TETRA SIRN	TIG SIRN	TMZ SIRN	<b>VAC</b> SIRN	

Pathogen #			Gra	ım-negative	Organisms			
	Acinetobacter (specify species)	AMK SIRN GENT SIRN TMZ SIRN	AMPSUL SIRN IMI SIRN TOBRA SIRN	AZT SIRN MERO/DORI SIRN	<b>CEFEP</b> SIRN	CEFTAZ SIRN PIP/PIPTAZ SIRN	CIPRO/LEVO SIRN	COL/PB SIRN
	Escherichia coli	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	AZT SIRN CIPRO/ LEVO/MOXI SIRN MERO/DORI SIRN	CEFAZ SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB <sup>†</sup> S R N TETRA/ DOXY/ MINO S I R N	CEFOT/ CEFTRX SIRN
	Enterobacter (specify species)	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	ATZ SIRN CIPRO/ LEVO/MOXI SIRN MERO/DORI SIRN	GENT SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB <sup>†</sup> S R N TETRA/ DOXY/ MINO S I R N	CEFOT/ CEFTRX SIRN
	Klebsiella pneumonia Klebsiella oxytoca	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	AZT SIRN CIPRO/ LEVO/MOXI SIRN MERO/DORI SIRN	CEFAZ SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB <sup>†</sup> S R N	CEFOT/ CEFTRX SIRN CEFTAZ SIRN TETRA/ DOXY/ MINO
	Pseudomonas aeruginosa	AMK SIRN IMI SIRN	AZT SIRN MERO/DORI SIRN	CEFEP SIRN	CEFTAZ SIRN PIP/PIPTAZ SIRN	CIPRO/LEVO SIRN TOBRA SIRN	COL/PB SIRN	GENT SIRN

Pathogen #				Fungal Orç	janisms			
	Candida	AMK	<b>AZT</b>	<b>AZT</b>	<b>CEFEP</b>	<b>CEFTAZ</b>	CIPRO/LEVO	COL/PB
	(specify species)	SIRN	SIRN	SIRN	SIRN	SIRN	SIRN	SIRN

Pathogen #				Otl	her Organ	isms				
	Organism 1 (specify)	Drug 1 SIRN	Drug 2 SIRN	Drug 3	Drug 4 SIRN	Drug 5 SIRN	Drug 6	Drug 7 SIRN	Drug 8 SIRN	Drug 9 SIRN
	Organism 1 (specify)	Drug 1 SIRN	Drug 2 SIRN	Drug 3 SIRN	Drug 4 SIRN	Drug 5 SIRN	Drug 6 SIRN	Drug 7	Drug 8 SIRN	Drug 9 SIRN
	Organism 1 (specify)	Drug 1 SIRN	Drug 2 SIRN	Drug 3	Drug 4	Drug 5 SIRN	Drug 6	Drug 7	Drug 8 S I R N	Drug 9 SIRN

### **Results Codes**

S = Susceptible I = Intermediate R = Resistant NS = Non-susceptible S-DD = Susceptible-dose dependent N = Not tested

 $\S$  GENTHL results: S = Susceptible/Synergistic and <math>R = Resistant/Not Synergistic

† Clinical breakpoints have not been set by FDA or CLSI, Sensitive and Resistant designations should be based upon epidemiological cutoffs of Sensitive MIC  $\leq$  2 and Resistant MIC  $\geq$  4

### **Drug Codes**

AMK = amikacin	CEFTRX = ceftriaxone	FLUCY = flucytosine	OX = oxacillin
AMP = ampicillin	CEFUR= cefuroxime	GENT = gentamicin	PB = polymyxin B
AMPSUL = ampicillin/sulbactam	CETET= cefotetan	GENTHL = gentamicin –high level test	PIP = piperacillin
AMXCLV = amoxicillin/clavulanic acid	CIPRO = ciprofloxacin	IMI = imipenem	PIPTAZ = piperacillin/tazobactam
ANID = anidulafungin	CLIND = clindamycin	ITRA = itraconazole	RIF = rifampin
AZT = aztreonam	COL = colistin	LEVO = levofloxacin	TETRA = tetracycline
CASPO = caspofungin	DAPTO = daptomycin	LNZ = linezolid	TIG = tigecycline
CEFAZ= cefazolin	DORI = doripenem	MERO = meropenem	TMZ = trimethoprim/sulfamethox azole
CEFEP = cefepime	DOXY = doxycycline	METH = methicillin	TOBRA = tobramycin
CEFOT = cefotaxime	ERTA = ertapenem	MICA = micafungin	VANC = vancomycin
CEFOX= cefoxitin	ERYTH = erythromycin	MINO = minocycline	VORI = voriconazole
CEFTAZ = ceftazidime	FLUCO = fluconazole	MOXI = moxifloxacin	

## Appendix E: Adult Bacteremia/Fungemia Shock Event (BSE) Manual Case Report Template (\* = optional)

Hospital ID:	Event #:
Patient ID:	Social Security #:
Secondary ID:	Medicare #:
Patient Name, Last:	First: Middle:
Gender: F M Other	Date of Birth:
Ethnicity (Specify):	Race (Specify):
Date Admitted to Hospital:	
Sepsis Onset Date:	*Location at Onset Date:
Event Type:	
$\hfill\Box$ Community-Onset (onset date on hospital day 2 or	earlier, when date of admission is hospital day 1)
$\ \square$ Hospital-Onset (onset date on hospital day 3 or late	r, when date of admission is hospital day 1)
Event Details (must meet Criteria #1 AND #2)  CRITERIA #1: Presumed Infection  □ Blood or Fungal culture with non-commensal patho  AND  CRITERIA #2: Organ Dysfunction: (± 2 days of blood	
☐ Initiation of a new vasopressor infusion¹	<u>o carare</u>
Vasopressor must not have been administered in prior calendar Qualifying vasopressors include: norepinephrine, dopamine, epi Single vasopressor doses or those given in an operating room do	nephrine, phenylephrine, vasopressin.
Died: Yes $\square$ No $\square$ *Sepsis Contributed to D	Peath: Yes □ No □
Discharge or Death Date:	
*Discharge Location:	
☐ Private residence ☐ Lor	ng-term care/Skilled nursing facility (SNF)
☐ Acute care hospital ☐ Ho	meless
$\square$ Long-term acute care hospital (LTACH) $\square$ Inc	arcerated
*Pathogens Identified: Yes □ No □ *If Yes, option	on to specify on next pages

Pathogen #	Gram-positive Organisms								
	Staphylococcus coagulase-negative (specify species if available):	VANC SIRN							
	Enterococcus faecium	DAPTO S NS N	<b>GENTHL</b> § SRN	LNZ SIRN	VANC SIRN				
	Enterococcus faecalis								
	<b>Enterococcus</b> <b>spp.</b> (Only those not identified to the species level)								
	Staphylococcus aureus	CIPRO/LEVO/ MOXI SIRN	<b>CLIND</b> SIRN	<b>DAPTO</b> S NS N	DOXY/ MINO SIRN	<b>ERYTH</b> SIRN	<b>GENT</b> SIRN	LNZ SRN	
		OX/CEFOX/ METH SIRN	RIF SIRN	TETRA SIRN	TIG SIRN	TMZ SIRN	<b>VAC</b> SIRN		

Pathogen #	Gram-negative Organisms							
	Acinetobacter (specify species)	AMK SIRN GENT SIRN TMZ SIRN	AMPSUL SIRN IMI SIRN TOBRA SIRN	AZT SIRN MERO/DORI SIRN	CEFEP SIRN	CEFTAZ SIRN PIP/ PIPTAZ SIRN	CIPRO/LEVO SIRN	COL/PB SIRN TETRA/ DOXY/ MINO SIRN
	Escherichia coli	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	AZT SIRN CIPRO/ LEVO/ MOXI SIRN MERO/ DORI SIRN	CEFAZ SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB† S R N TETRA/ DOXY/MINO S I R N	CEFOT/ CEFTRX SIRN
	Enterobacter (specify species)	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	ATZ SIRN CIPRO/ LEVO/ MOXI SIRN MERO/ DORI SIRN	CEFAZ SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB <sup>†</sup> S R N TETRA/ DOXY/MINO S I R N	CEFOT/ CEFTRX SIRN
	Klebsiella pneumonia Klebsiella oxytoca	AMK SIRN CEFTAZ SIRN ERTA SIRN TIG SIRN	AMP SIRN CEFUR SIRN GENT SIRN TMZ SIRN	AMPSUL/ AMXCLV SIRN CEFOX/ CETET SIRN IMI SIRN TOBRA SIRN	AZT SIRN CIPRO/ LEVO/ MOXI SIRN MERO/ DORI SIRN	CEFAZ SIRN PIPTAZ SIRN	CEFEP S I/S-DD R N COL/PB <sup>†</sup> S R N	CEFOT/ CEFTRX SIRN CEFTAZ SIRN TETRA/ DOXY/ MINO
_	Pseudomonas aeruginosa	AMK SIRN IMI SIRN	AZT SIRN MERO/ DORI SIRN	<b>CEFEP</b> SIRN	CEFTAZ SIRN PIP/PIPTAZ SIRN	CIPRO/ LEVO SIRN TOBRA SIRN	COL/PB SIRN	<b>GENT</b> SIRN

Pathogen #	Fungal Organisms									
	Organism 1 (specify)	Drug 1 SIRN	Drug 2 SIRN	Drug 3 SIRN	Drug 4 SIRN	Drug 5 SIRN	Drug 6 SIRN	Drug 7 SIRN	Drug 8 SIRN	Drug 9 SIRN
	Organism 1	Drug 1	Drug 2	Drug 3	Drug 4	Drug 5	Drug 6	Drug 7	Drug 8	Drug 9
	(specify)	SIRN								
	Organism 1	Drug 1	Drug 2	Drug 3	Drug 4	Drug 5	Drug 6	Drug 7	Drug 8	Drug 9
	(specify)	SIRN	S I R N							

### **Results Codes**

S = Susceptible I = Intermediate R = Resistant NS = Non-susceptible S-DD = Susceptible-dose dependent N = Not tested

§ GENTHL results: S = Susceptible/Synergistic and R = Resistant/Not Synergistic

† Clinical breakpoints have not been set by FDA or CLSI, Sensitive and Resistant designations should be based upon epidemiological cutoffs of Sensitive MIC  $\leq$  2 and Resistant MIC  $\geq$  4

### **Drug Codes**

AMK = amikacin	CEFTRX = ceftriaxone	FLUCY = flucytosine	OX = oxacillin
AMP = ampicillin	CEFUR= cefuroxime	GENT = gentamicin	PB = polymyxin B
AMPSUL = ampicillin/sulbactam	CETET= cefotetan	GENTHL = gentamicin – high level test	PIP = piperacillin
AMXCLV = amoxicillin/clavulanic acid	CIPRO = ciprofloxacin	IMI = imipenem	PIPTAZ = piperacillin/tazobactam
ANID = anidulafungin	CLIND = clindamycin	ITRA = itraconazole	RIF = rifampin
AZT = aztreonam	COL = colistin	LEVO = levofloxacin	TETRA = tetracycline
CASPO = caspofungin	DAPTO = daptomycin	LNZ = linezolid	TIG = tigecycline
CEFAZ= cefazolin	DORI = doripenem	MERO = meropenem	TMZ = trimethoprim/sulfamethoxazole
CEFEP = cefepime	DOXY = doxycycline	METH = methicillin	TOBRA = tobramycin
CEFOT = cefotaxime	ERTA = ertapenem	MICA = micafungin	VANC = vancomycin
CEFOX= cefoxitin	ERYTH = erythromycin	MINO = minocycline	VORI = voriconazole
CEFTAZ = ceftazidime	FLUCO = fluconazole	MOXI = moxifloxacin	

# NOTES



U.S. Department of Health and Human Services Centers for Disease Control and Prevention