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Clinical Paper

The incidence and significance of bacteremia in out of hospital cardiac arrest^{a,b}

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Introduction

- The most common etiology of cardiac arrest is presumed of myocardial origin.
- Recent retrospective studies indicate that preexisting pneumonia is frequent in patients with cardiac arrest without preceding signs of septic shock, respiratory failure or severe metabolic disorders

- Carr et al examined the characteristics of IHCA diagnosed with pre-existing pneumonia
 - The incidence of pre-existing pneumonia was 12.1% (5367 of 44,416) from a multicenter cardiac arrest registry. S. Carr GE, Yuen TC, McConville JF, et al. Early cardiac arrest in patients hospitalized with pneumonia: a report from the American Heart Association's Get With The Guidelines-Resuscitation Program. Chest 2012;141:1528–36.
- Abrupt cardiac arrest without signs of hypotension, overt shock, respiratory failure or severe metabolic derangements.

- Little is known the prevalence of bacteremia in cardiac arrest patients arriving to the ED.
- ✓ To examine the incidence of bacteremia in out of hospital cardiac arrest (OHCA) and identify the characteristics associated with bacteremia.

Methods

- This was a **prospective observational convenient sampling study** of OHCA adult patients seen at an urban academic teaching institution
- August 2007 to August 2009
- The study was designed to obtain blood cultures near time of cardiac arrest event during ACLS therapy or return of spontaneous circulation (ROSC)

- The excluded were all trauma cardiac arrest victims, pregnant patients, and patients found younger than 18 years.
- Bacteremic OHCA group
 - one blood culture tube with non-skin flora pathogens
- Non-bacteremic OHCA group
 - no bacterial growth after at least 5 days

- The **primary objective** of the study was to identify the incidence of bacteremia in OHCA adults.
- **Secondly**, *patient characteristics* and *hospital mortality* of the bacteremic OHCA compared to the non-bacteremic OHCA group

- 65 (38%) were found to be bacteremic with **asystole** and **PEA** as the most common presenting rhythms.
- **Mortality in the ED** was significantly higher in bacteremic OHCA (75.4%) compared to non-bacteremic OHCA (60.2%, $p < 0.05$).
- Predictive factors associated with bacteremic OHCA were lower initial arterial pH, higher lactate, WBC, BUN and creatinine.

Table 2
Resuscitation Profile of the Bacteremic and Non-bacteremic OHCA groups.

	Bacteremic OHCA (n=65)	Non-bacteremic OHCA (n=108)	P
Resuscitation profile			
Cardiac arrest location			
Home	49% (32)	51% (55)	0.94
Nursing home	8% (5)	9% (10)	
Public place	18% (12)	16% (17)	
Emergency Department	6% (4)	8% (9)	
Unknown	18% (12)	16% (17)	
Presenting rhythm			
Asystole	65% (42)	62% (67)	0.45
PEA	26% (17)	22% (24)	
VF	9% (6)	16% (17)	
CPR time			
Pre-hospital CPR time (m)	24.49 ± 12.4	20.6 ± 12.6	0.06
ED CPR time (m)	16.9 ± 15.7	17.2 ± 14.2	0.35
Time to ROSC (m)	13.12 ± 9.8	14.65 ± 12.8	0.66
Total CPR time (m)	40.9 ± 21.7	35.9 ± 20.4	0.15

PEA indicates pulseless electrical activity; VF, ventricular fibrillation; CPR, cardiopulmonary resuscitation; ED, emergency medicine; ROSC, return of spontaneous circulation; m, minutes. Continuous variables are presented as mean ± SD unless stated otherwise. Categorical variables are presented as percentages and absolute numbers (n).

Results

- From 2007 to 2009, 250 OHCA cardiac arrest patients had 2 sets of blood cultures obtained.
- 77 (31%) met exclusion criteria
- There was 173 patients, 71 had positive blood cultures of which cultures were defined as 6 contaminants (skin flora) and included for analysis in the non-bacteremic OHCA group.

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Table 1
Demographics characteristics of patients with OHCA presenting to the emergency department and disposition from the emergency department.

Patient characteristics	Bacteremic OHCA (n=65)	Non-bacteremic OHCA (n=108)	P
Age, years	65.5 ± 15.1	64.8 ± 18.5	0.74
Male	55.4% (36)	59.3% (64)	0.42
Race			
African American	88% (57)	83% (90)	0.86
Caucasian	9% (6)	13% (14)	
Other	3% (2)	4% (4)	
Preexisting comorbid conditions			
Diabetes mellitus	34% (22)	31% (31)	0.81
Hypertension	22% (14)	48% (52)	0.63
Congestive heart failure	23% (15)	23% (25)	0.55
Coronary artery disease	20% (13)	31% (33)	0.38
Chronic renal failure on hemodialysis	8% (5)	6% (6)	0.78
Chronic renal failure without hemodialysis	5% (3)	13% (14)	0.07
Chronic obstructive pulmonary disease	22% (14)	29% (31)	0.28
ED survival	25% (16)	40% (43)	0.042
28 days mortality	94% (61)	93% (100)	0.75
Overall mortality	97% (63)	95% (103)	0.66

ED indicates emergency department. Continuous variables are presented as mean ± SD. Categorical variables are presented as percentages and absolute numbers (n).

Table 3
Bacterial species isolated from the Bacteremic OHCA group.

Bacterial species	n (%)
Other Streptococcus species	12 (18.5%)
Staphylococcus epidermidis	11 (17%)
Other Staphylococcus species	9 (14%)
Corynebacterium	5 (7.7%)
Staphylococcus aureus	4 (6%)
Streptococcus pneumoniae	4 (6%)
Klebsiella pneumoniae	3 (4.6%)
Enterococcus faecalis	3 (4.6%)
Propionibacterium	3 (4.6%)
Escherichia coli	3 (4.6%)
Clostridium species	3 (4.6%)
Gram positive bacilli NOS	2 (3%)
Enterococcus species	2 (3%)
Streptococcus agalactiae	2 (3%)
Clostridium cadaveris	2 (3%)
Proteus mirabilis	2 (3%)
Enterobacterium	1 (1.5%)
Enterococcus faecium	1 (1.5%)
Actinomyces	1 (1.5%)
Bacteriodes (Prevotella)	1 (1.5%)
Aerococcus viridans	1 (1.5%)

NOS indicates not otherwise specified. All bacterial species isolated were reported and on occasion more than one bacterial species were isolated noting percent summation greater than 100%. Categorical values are shown as absolute numbers and percentages (%).

Table 4
Baseline physiologic and laboratory parameters of the Bacteremic and Non-bacteremic OHCA ED survivors as well as ED disposition.

	Bacteremic ED survivors (n=16)	Non-bacteremic ED survivors (n=43)	P
Vital signs			
Temperature (°C)	35.0 ± 1.5	35.5 ± 1.6	0.48
HR, beats per min	104 ± 27	105 ± 28	0.90
RR, breath per min	14 ± 4	18 ± 8	0.25
Systolic BP (mmHg)	125 ± 51	136 ± 52	0.58
Diastolic BP (mmHg)	70 ± 34	79 ± 32	0.50
MAP (mmHg)	89 ± 37	98 ± 38	0.51
O ₂ Saturation (%)	90% ± 6.8	90% ± 14	0.54
Shock index	0.89 ± 0.47	0.89 ± 0.34	0.83
Vasopressors dependent (%)	93.8% (15)	74.4% (32)	0.055
Laboratory values			
White blood cells (Kp/L)	14.08 ± 7.8	12.5 ± 7.7	0.40
Hemoglobin (g/dL)	11.3 ± 1.8	11.5 ± 2.4	0.69
Hematocrit (%)	34.9 ± 6.5	34.7 ± 6.7	0.91
Lactate (mmol/L)	13.4 ± 7.3	8.8 ± 5.5	0.01
Arterial pH	7.03 ± 0.2	7.17 ± 0.17	0.01
Arterial pCO ₂ (mmHg)	67.3 ± 28.2	52.6 ± 23.1	0.05
Arterial pO ₂ (mmHg)	132.2 ± 118	132.2 ± 114	0.16
Base excess (mmol/L)	14.8 ± 8.3	10.5 ± 6.0	0.04
Sodium (mmol/L)	144.6 ± 10.1	142.4 ± 8.4	0.41
Potassium (mmol/L)	5.2 ± 1.2	4.1 ± 2.4	0.005
Chloride (mmol/L)	104.0 ± 7.7	107 ± 8.5	0.39
Bicarbonate (mmol/L)	18.5 ± 5.3	20.1 ± 8.6	0.40
BUN (mg/dL)	48.8 ± 31.1	27.8 ± 19.6	0.02
Creatinine (mg/dL)	3.7 ± 2.2	2.4 ± 1.7	0.02
Glucose (mg/dL)	253 ± 102	240.7 ± 121	0.93
Magnesium (mg/dL)	2.2 ± 0.6	2.1 ± 0.6	0.61
Phosphorus (mg/dL)	10.3 ± 4.6	7.4 ± 3.7	0.01
Troponin I (ng/mL)	1.23 ± 1.08	1.34 ± 0.17	0.15
CK-MB (ng/mL)	7.8 ± 0.14	30.1 ± 42.36	0.13
PT (s)	22.3 ± 5.0	22.7 ± 28.8	0.93
INR	1.94 ± 0.57	2.21 ± 4.5	0.71
Hospital disposition			
ICU LOS, d	3.50 ± 5.34	5.49 ± 7.99	0.36
Hospital LOS, d	4.25 ± 7.15	6.63 ± 8.88	0.34
28 day mortality	81% (13)	81% (35)	0.65
Overall mortality	88% (14)	84% (36)	0.25
Antibiotics initiated in the ED	69% (11)	30% (13)	0.01

ED indicates emergency department; °C indicates Celsius; HR, heart rate; RR, respiratory rate; BP, blood pressure; MAP, mean arterial pressure; O₂, oxygen. Continuous variables are presented as mean ± SD unless stated otherwise. BUN, indicated blood urea nitrogen; CKMB, creatine kinase myocardial band; PT, prothrombin time; INR, international normalized ratio and LOS, length of stay. Categorical variables are presented as percentages and absolute numbers (n). Continuous variables are presented as

Discussion

- OHCA patients presenting to the ED, we found a 38% incidence of bacteremia
- The presence of bacteremia was associated with a significant decrease in ED survival compared to non-bacteremic patients.

Bacteremia: incidence and significance

- Risk factors contributing significantly to mortality in bacteremic patients were
 - increasing age
 - underlying ultimately fatal disease
 - presence of severe sepsis, shock and gram-positive pathogen infections
- Our most common pathogens identified were not of *intestinal origin* rather **Staphylococcus and Streptococcus species**.
- The pathogens identified was likely related to the timing of blood culture sampling
- The incidence of bacteremic complications OHCA was first reported by Gaussorgues et al. at 8%.
 - They hypothesized that a low flow shock state post cardiac arrest along with mesenteric ischemia
 - With gut flora bacteremia in blood cultures obtained 12 h after ROSC
- In this study, the incidence of bacteremia in OHCA is **higher than** any previously reported
- In this study, the ED medical team initiated empiric antibiotic therapy in 69% of bacteremic OHCA ED survivors.

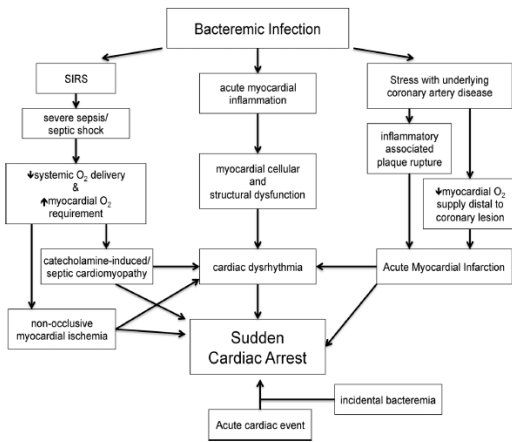


Fig. 1. Proposed association between bacteremic infection and sudden cardiac arrest.

How this paper can change management or current ACLS guideline

- Current 2012 AHA/ACLS note the importance of post-CPR for
 - therapeutic hypothermia
 - hemodynamic and ventilation optimization
 - coronary reperfusion
 - glycemic control
 - neurologic care

Limitations

- This study suggests that one major precipitating factor or comorbidity associated with OHCA is unrecognized **bacteremia**.
- To initiate **prompt empiric antibiotic therapy** during the post cardiac arrest resuscitation period for the bacteremic OHCA.
- The study was designed to report blood cultures results without requiring specimens from other potential sites of infection.
- The ED medical team initiated empiric antibiotic treatments for both groups.

Conclusion

- This study is the first to report over 38% incidence of bacteremia in OHCA adults.
- The bacteremic OHCA adults had severe metabolic derangements, higher lactates and lower ED survival
- Further study is needed to identify the cause and effect relationship between bacteremia and sudden cardiac arrest