UCDAVIS HEALTH

Introduction

The rapid multiplex PCR (rmPCR)-based FilmArray® blood culture identification (BCID) assay reduces time from positive blood culture to organism identification. Polymicrobial bacteremia is a known area of reduced diagnostic fidelity for BCID and remains incompletely characterized.

Methods

All cases of clinically confirmed polymicrobial bacteremia at a large academic single center from a 23-month period were evaluated in a retrospective cohort analysis (figure 1). Samples were assorted into BCID/blood culture concordant and BCID/blood culture discordant groups. Clinical characteristics of the two groups were compared, missed organisms were characterized, and changes in antimicrobial regimen in response to BCID results were characterized.



Figure 1. Screening and exclusion process. 207 cultures were included in final analysis from a number screened of 2750 (constituting all positive blood cultures over a 23-month period from February 2019 to January 2021).

Blood Culture Identification (BCID) Performance in Polymicrobial Bacteremia

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Results

A total of 207 samples were identified and studied. Overall, 49.3% (N=102) of polymicrobial cultures were incompletely identified by FilmArray® result. There were no significant group differences in comorbidity status, length of stay, mortality, or source between patients with polymicrobial bacteremia who had complete versus incomplete BCID identification (see table 1). In the 102 BCID discordant samples, 127 individual organisms identified on phenotyping but not on BCID were found. Many were commensal or low virulence organisms, but a total of 38 (29.9%) were identified as organisms potentially requiring prompt treatment, while 49 (38.6%) of organisms were on panel for the BCID assay (list of organisms is shown in Table 2). Of note, there were no instances of false negative results for Pseudomonas aeruginosa, Neisseria meningitidis, or Listeria monocytogenes, organisms requiring prompt therapy sometimes missing in empiric antibiotic regimens.

	BCID/BCx Concordant (%)	BCID/BCx Discordant (%)	p value
Number of samples	105 (50.7%)	102 (49.3%)	
Average age of patient (mean,	58.6 (18.4)	54.1 (18.6)	0.08
SD)			
Length of stay (median, IQR)	13.0 (18.3)	12.0 (21.9)	0.12
Death within 100 days	35 (33.3%)	30 (29.4%)	0.54
Male Sex	61 (58.1%)	62 (60.8%)	0.69
Venue collected	40 (45 00/)	40 (0.00/)	
	16 (15.2%)	10 (9.8%)	
FIOOR	19 (18.1%)	27 (26.5%)	0.22
ER	68 (64.8%)	60 (58.8%)	
Comorbidition	2(1.9%)	5 (4.9%)	
Burn	8 (7 6%)	1 (2 0%)	0.26
Solid maliananay	0 (7.070) 25 (23.8%)	4 (3.970)	0.20
Liquid malignancy	ZJ (ZJ.070) Z (6 7%)	10(13.7.70) 13(13.80/)	$\begin{array}{c} 0.14 \\ 0.14 \end{array}$
On chamatharany	1 (0.7 / 0) 12 (11 / 0/)	15(12.070) 15(1770)	0.14
Noutropopia	12 (11.470)	13(14.770) 11(10.8%)	0.40
Abdominal surgery within 30	12 (12.470)	8 (7 8%)	0.72
Abuominal Surgery within 50	12 (11.470)	0 (7.070)	0.00
Solid organ transplant	2 (1 9%)	2 (2 0%)	<u>_0 00</u>
Hematonoietic stem cell	2 (1.970) 5 (1.8%)	2 (2.0%)	20.99 0 45
transplant	5 (4.070)	2 (2.070)	0.40
Current TPN use	9 (8 6%)	6 (5 9%)	0.46
Presence of central venous	38 (36 2%)	0(3.370) 11(10.2%)	0.40
catheter	30 (30.270)	+1 (+0.270)	0.00
Current hemodialysis	8 (7.6%)	7 (6.9%)	0.83
Cirrhosis	6 (5.7%)	8 (7.8%)	0.54
Current immunosuppressant	17 (16.2%)	9 (8.8%)	0.11
medication use	· · ·	· · ·	
Other comorbidity	10 (9.5%)	7 (6.9%)	0.49
None of the above	33 (31.4%)	38 (37.3%)	0.38
Presumed source			
Central line	17 (16.2%)	17 (16.7%)	
Gastrointestinal	21 (20.0%)	31 (30.4%)	
Genitourinary	11 (10.5%)	9 (8.8%)	
Respiratory	2 (1.9%)	6 (5.9%)	
Endovascular	3 (2.9%)	4 (3.9%)	
Bone	3 (2.9%)	4 (3.9%)	
Skin and soft tissue	17 (16.2%)	12 (11.8%)	
Other	1 (1.0%)	1 (1.0%)	
Multiple sources	19 (18.1%)	11 (10.8%)	
Unknown	11 (10.5%)	7 (6.9%)	

Table 1. Comparison of the characteristics of the
 BCID/blood culture (BCx) concordant and BCID/BCx discordant groups.

De-escalation from adequate empiric to inadequate step-down antibiotic coverage following incomplete BCID result occurred in only 8.8% (N=9) of cases (shown in table 3). No statistically significant association with mortality was seen among patients' empiric coverage (p=0.07), although this analysis was limited by the small sample size.

Organism	No.		No.
On-panel organisms	49	Other off-panel organisms	50
Enterococcus spp.	11	Achromobacter spp	2
Viridans group streptococci	8	Acinetobacter spp	3
Coagulase negative Staphylococci	7	Actinomyces spp	1
Staphylococcus aureus	6	Actinotignum schaelii	1
Klebsiella pneumoniae	3	Bacteroides spp.	6
Acinetobacter baumanii	2	Chryseobacterium spp.	5
Candida albicans	2	Corynebacterium spp.	2
Enterobacter cloacae complex	2	Eikenella spp	1
Klebsiella oxytoca	2	Gemella spp.	7
Proteus mirabilis	2	Granulicatella spp	2
Candida glabrata	1	Lactobacillus spp	3
Candida tropicalis	1	Micrococcus luteus	2
Escherichia coli	1	Mycobacterium abscessus	1
Streptococcus agalectiae	1	Neisseria subflavia	1
Off-panel organisms of high		Parabacteroides spp.	1
pathogenic potential	28	Pluralibacter gergoviae	1
Clostridium perfringens	6	Pseudomonas putida grp	2
Bacillus spp.	4	Rothia mucilaginosa	4
Morganella morganii	4	Veilonella spp.	3
Corynebacterium striatum	3	Wohlfahrtimonas spp.	1
Aerococcus viridans	2		
Candida dublinensis	2		
Clostridium ramosum	2		
Stenotrophomonas maltophilia	2		
Fusobacterium nucleatum	1		
Fusobacterium necrophorum	1		
Providencia stuartii	1		

 Table 2. Organisms not identified by BCID but later identified
 on blood culture phenotyping.



Empiric therapy	De- escalation therapy	No. (%)	30-day mortality, no. (%)	Pooled Inpatient time p-value to discharge, median [IQR]	Pooled p- value
Effective	Effective	58 (56.9%)	10 (17.2%)	11.6 [22.0]	
Effective	Did not occur	14 (13.7%)	6 (42.9%)	25.0 [27.9] 0.07	0.14
Effective	Inadequate	9 (8.8%)	3 (33.3%)	8.3 [4.2]	
Ineffective	N/A	21 (20.6%)	2 (9.5%)	17.1 [19.5]	

Table 3. Among patients with inaccurate BCIDs, antimicrobial changes and outcomes following return of BCID result are shown below. P-values comparing the four groups are shown. There was no significant association that could be seen in mortality and time to discharge and patient's empiric therapy and de-escalation therapy status.

Conclusions

BCID frequently results in incomplete identification of blood culture results in patients with polymicrobial bacteremia, but clinical characteristics and outcomes were similar to those of patients with accurate BCID identification. Clinical team de-escalation to inappropriate antibiotic coverage following return the BCID assay was uncommon and was not clearly associated with inferior outcomes.

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