

Diagnostic stewardship

**Does optimization of the blood culture pathway
have a role to play?**

Dr Michael Weinbren
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BSMT MAY 2018

Optimising the

A digital landscape with a glowing road leading to a bright light at the horizon, with binary code floating in the air.

Dublin 2014 IBMS ANNUAL CONFERENCE.

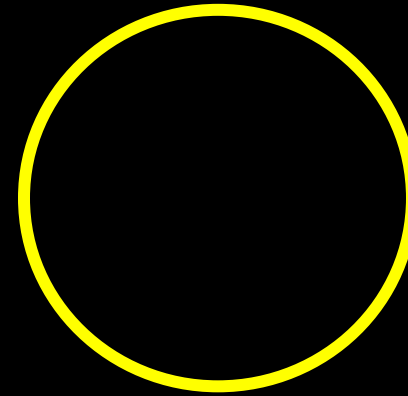
‘Blood cultures-
bacteria or microbiologists-
who is the
smart money on ?’

R. E. Holliman. The therapeutic impact of blood culture results. Journal of Hospital Infection (1986) 7, 185-188

- Summary: The influence on therapy of blood culture results was monitored over a 3-month period.
- Approximately half the patients yielding significant cultures commenced initial or altered antibiotic treatment on the basis of laboratory results.
- Therapy based on a defined antibiotic policy was found to be satisfactory in most instances.
- Whilst an antibiotic policy allows effective treatment of many patients, there remains a need for an early microbiological diagnosis.

GENTAMICIN

MEROPENEM




AMIKACIN

**PIPERACILLIN/
TAZOBACTAM**

CEFTAZIDIME

CIPROFLOXACIN



But is that all there is to blood culture microbiology?

Blood cultures are the gold standard for diagnosis of blood stream infections

Rapid results associated with improved outcome, shorter length of stay

Is there a role for improved antimicrobial stewardship?



BD

Pre analytical

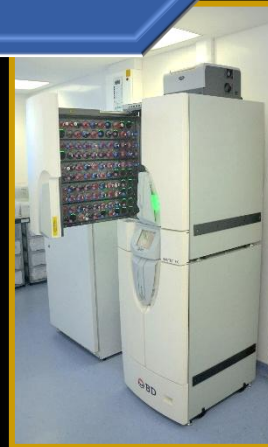
1



Right patient / volume
of blood to analyser
with minimum delay

Analytical

2



**SHORTEST TIME
TO KEY RESULT**

Post analytical

3

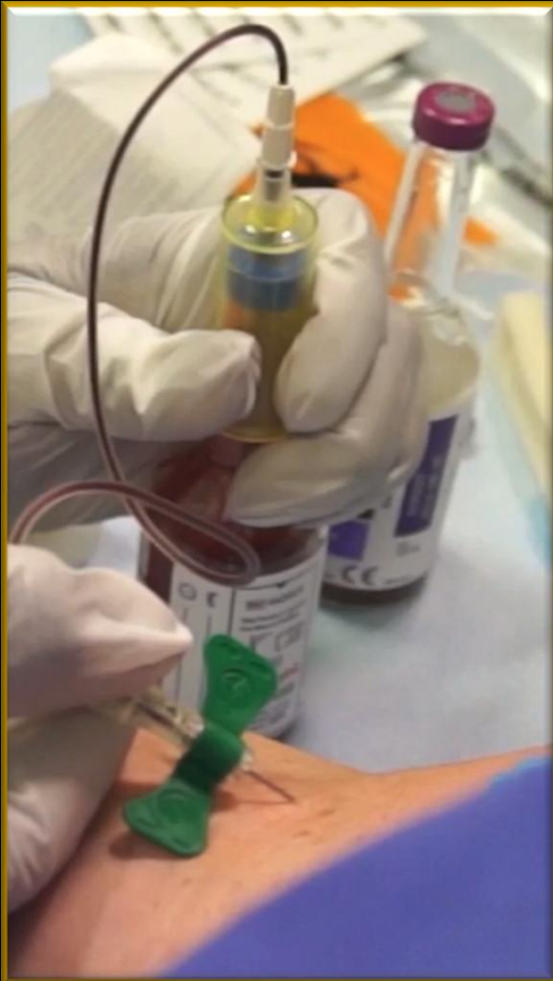


**USE KEY RESULT
TO MANAGE
PATIENT WITH
MINIMUM DELAY**

FINISH

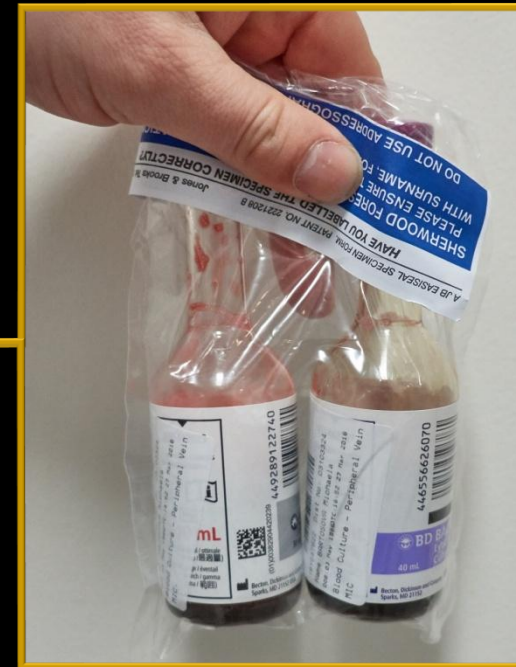
Pre analytical

Blood culture collection

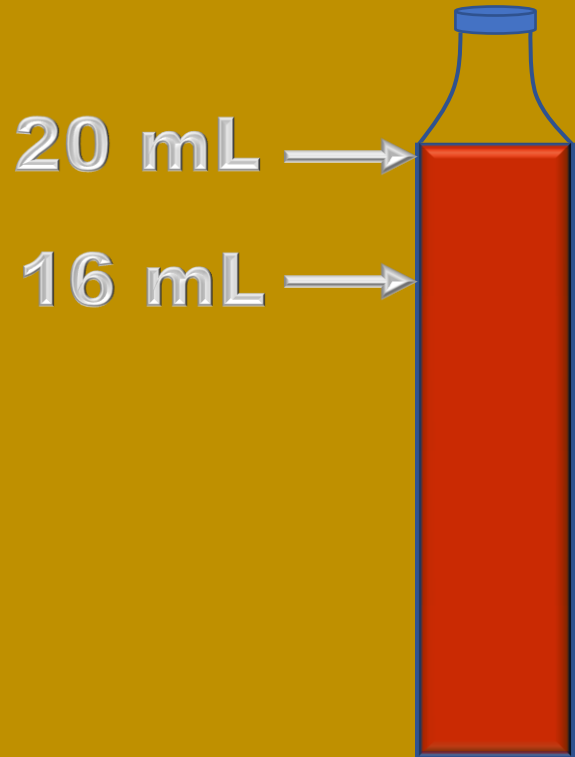


Right patient

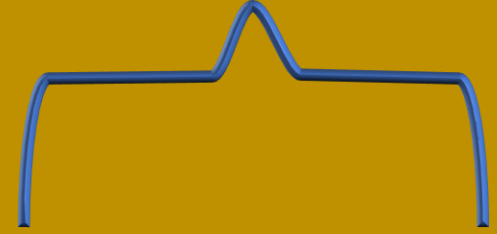
Right volume

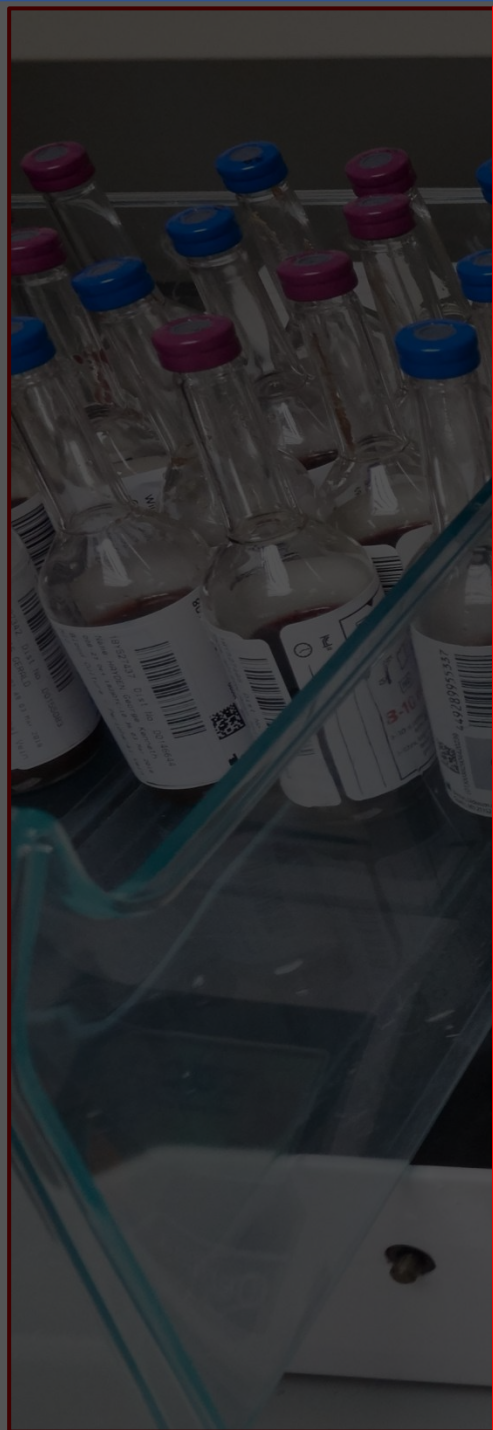


Volume of blood received in one set of blood cultures



25% adequately filled





Bottle signalling positive first

80%

70%

60%

50%

40%

30%

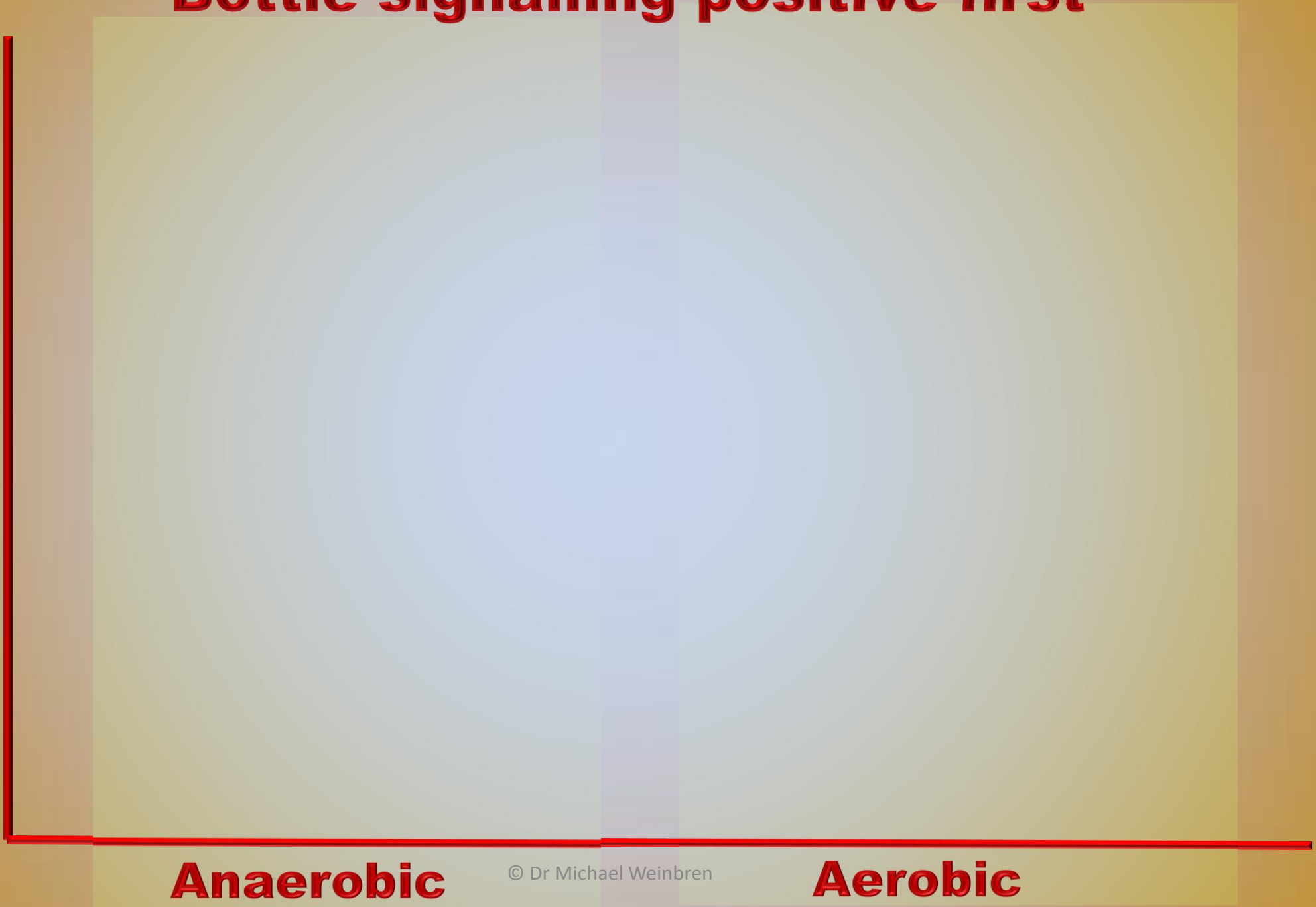
20%

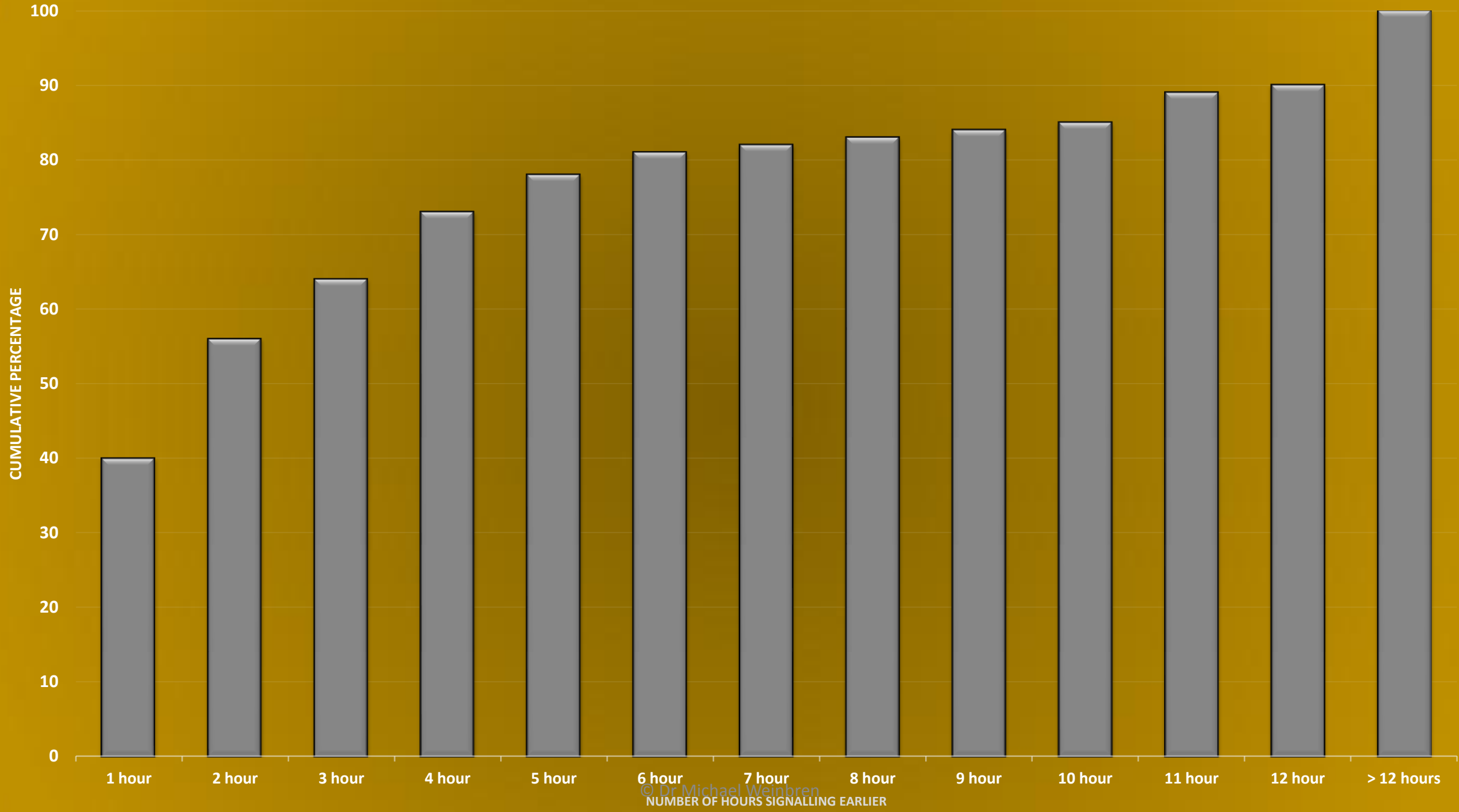
10%

Anaerobic

© Dr Michael Weinbren

Aerobic





Pre analytical

Blood culture collection



Right patient

Right volume

Right time

2 sets- 20% extra E.coli
Vivienne Weston

Two Sets
in adults



SEPSIS- 40ml blood-
50% extra significant
positive blood cultures
Shabnam Iyer

© Dr Michael Weinbren

≤4 hours



Analytical

Blood culture processing

TIME TO KEY INFORMATION

36 HOUR NEGATIVE
BLOOD CULTURE

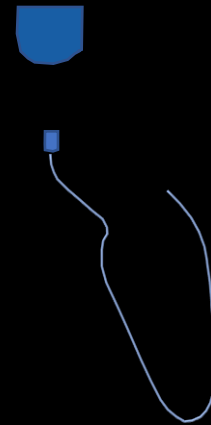
GRAM STAIN

ORGANISM ID

ANTIBIOTIC
SENSITIVITIES



Post
analytical

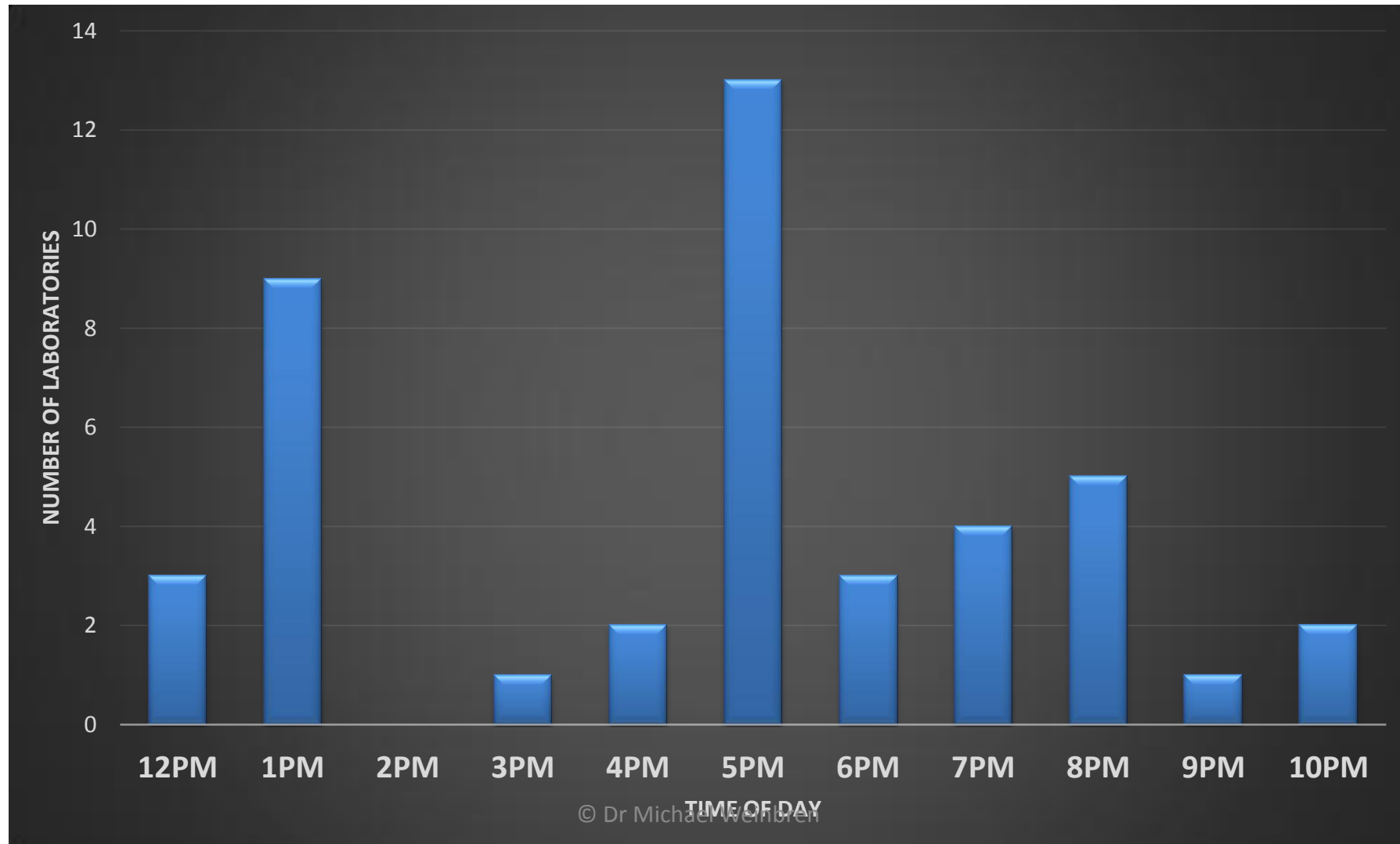


NATIONAL TELEPHONE SURVEY IN 2012 – 43 LABOARTORIES

- **NO** LABORATORY LOADS BLOOD CULTURE MACHINE DURING NIGHT
- **21 LABORATORIES** PRE-INCUBATE BLOOD CULTURES OVERNIGHT
- **22 LABORATORIES** LEAVE BLOOD CULTURES AT ROOM TEMPERATURE OVERNIGHT
- 24 HOUR SHIFT SYSTEM IN BLOOD SCIENCES IN SAME LABORATORY **31 LABORATORIES**

WEEKEND

TIME LAST POSITIVE BLOOD CULTURE PROCESSED



AUDIT OF BLOOD CULTURE TURNAROUND TIME

- REGULARLY- 2
- OCCASIONALLY – 11
- DO NOT KNOW- 1
- **NEVER- 29**

UK Standards for Microbiology Investigations

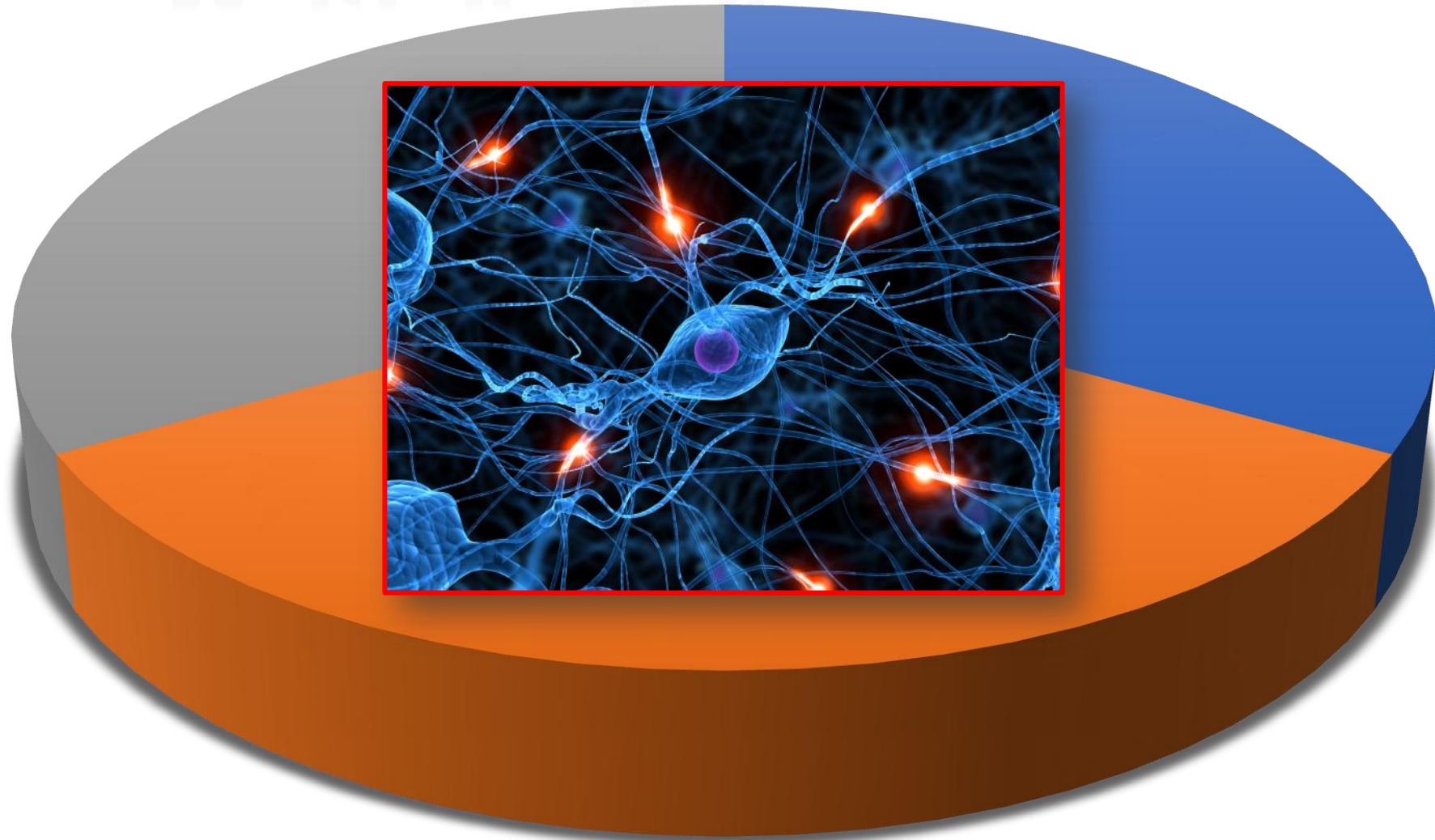
Investigation of Blood Cultures (for Organisms other than *Mycobacterium* species)



1. Audit of blood culture pathway is key

2. Set time standards for critical control points

Multi-disciplinary group



■ BLOOD SCIENCE ■ MICROBIOLOGY ■ CLINICAL

© Dr. Michael Welborn

SAMPLE COLLECTION

PORTER

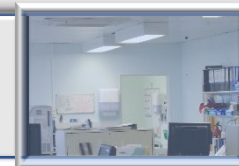
SPECIMEN RECEPTION

MICROBIOLOGY

**FLAGS
POSITIVE**

**BLOOD
CULTURE
MACHINE**

CHESTERFIELD PATHOLOGY BUILDING







SAMPLE COLLECTION

SPECIMEN RECEPTION

FLAGS
POSITIVE

FLAGS
POSITIVE

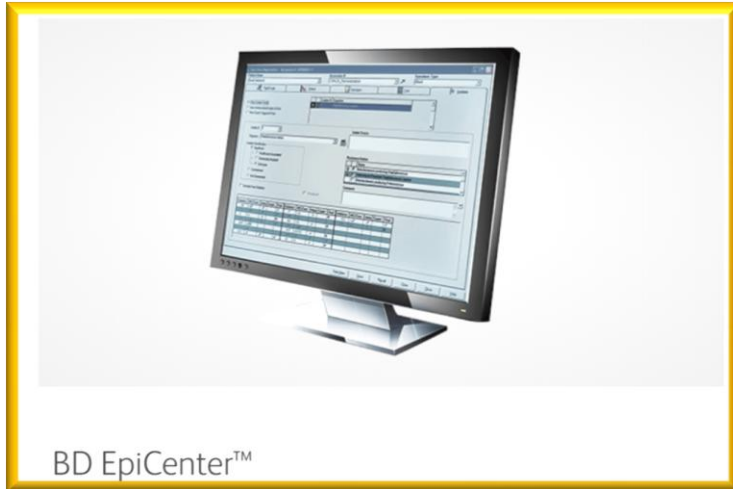
MICROBIOLOGY



Sensitivity plates



Rapid sensitivities



BD EpiCenter™

Table 1 The average time to positivity

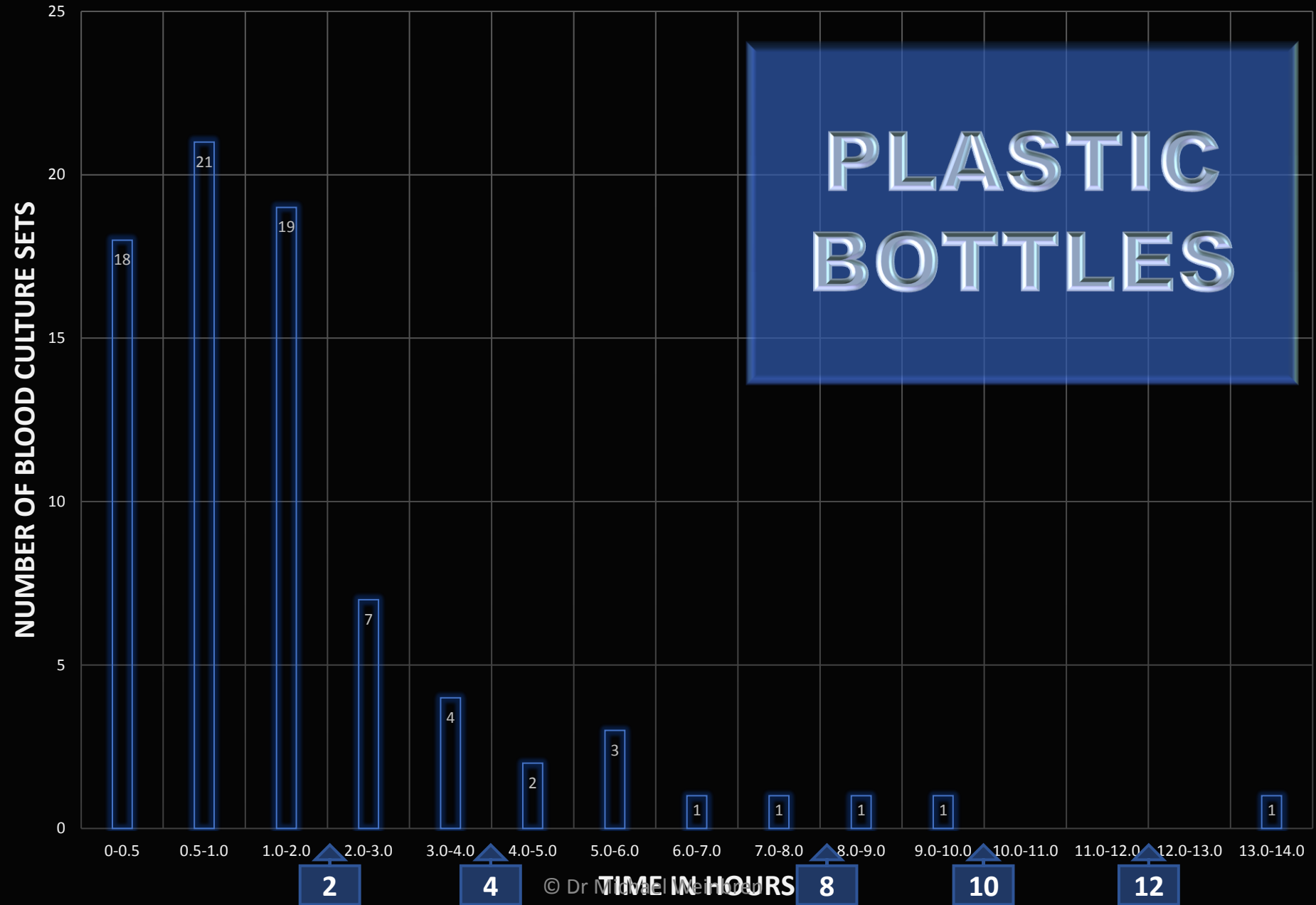
pre-move	17 hours and 49 minutes
post move	18 hours and five minutes

**Table 3 The average time to positivity for
E. coli**

pre-move	15 hours 34 minutes
post move	12 hours 56 minutes

E. coli common pathogen in blood cultures
Rarely a contaminant
Allows coarse comparison between laboratories

TIME FROM COLLECTION TO PLACEMENT ON ANALYSER



	1 HOUR	2 HOURS	3 HOURS	4 HOURS	>4 HOURS	NUMBER OF SETS
GLASS	46%	69%	77%	82%	18%	84
PLASTIC						
PLASTIC 2						
JAN 1						
JAN 2						
SEPT						

The 'clinical' perspective

- Urgent requirement for blood culture transportation not common knowledge
- Not taught at medical school or afterwards
- Conduct survey of hospital staff across grades and staff groups

MICROBIOLOGY QUESTIONNAIRE

- The purpose of this questionnaire is not to test your own knowledge
- It is to find out if pathology has provided you with adequate information.
- Therefore please answer honestly , write what you think– there are no rights or wrongs in this setting.
- Only by learning what you know can we discover if there are any issues that can be improved upon.

Question 1. The microbiology laboratory opening times are;

- (a) Routine service 24 hours / day
- (b) 08.30 to 20.00 hours Monday to Friday, 09.00 to 17.00 Saturday / Sunday with an on call service outside of these times
- (c) None of the above

Question 2. How quickly are initial results routinely available for the following specimens;

	CSF	BLOOD CULTURE	URINE
1-2 HOURS			
24 HOURS			
48 HOURS			
72 HOURS			
OTHER PLEASE STATE			

Question 3. During routine hours how quickly should the following specimens be sent to the Laboratory;

	CSF	BLOOD CULTURE	URINE
IMMEDIATELY			
WITHIN 1 HOUR			
UP TO 2 HOURS			
NO URGENCY AS LONG AS ARRIVES SAME DAY			
OTHER PLEASE STATE			

91% of staff thought a CSF sample should be sent immediately to the lab

68% would send a blood culture immediately

Out of hours only 52% of staff would send a blood culture immediately

28% of staff thought it took ≥ 48 hours for a blood culture to flag positive

UNIVERSITY HOSPITAL PATHOLOGY REPORT

Patient- J. Brown NHS No. 456785943

Specimen- blood culture Collected 14/04/2016

Result-

NEGATIVE AFTER 48 HOURS INCUBATION

Authorised BSF 17/04/16

**‘there was a box
in ED for blood cultures
which was emptied
once or twice / day’**





BLOOD CULTURES

Use the Pod -save a life

This bottle may contain a killer.
Help us stop them before they
succeed.

Pod it- save time, save lives

The quicker a blood culture reaches the laboratory the sooner
the correct antibiotic can be found for the patient.

For the sick patient minutes count.

By podding the blood culture at the earliest opportunity you could save a life.

Use the Pod -save a life



Blood cultures sitting around on a ward could mean a patient lying down forever.

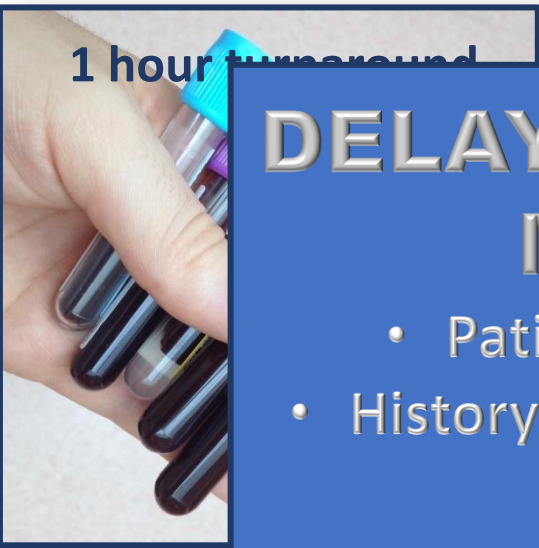
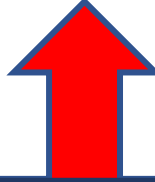
Help us protect patients.

The sooner a blood culture reaches the laboratory the quicker the invading organism can be identified, so the correct antibiotic can be found to stop it in its tracks.

Blood cultures– Pod them and you could save a life

© Dr Michael Weinbren

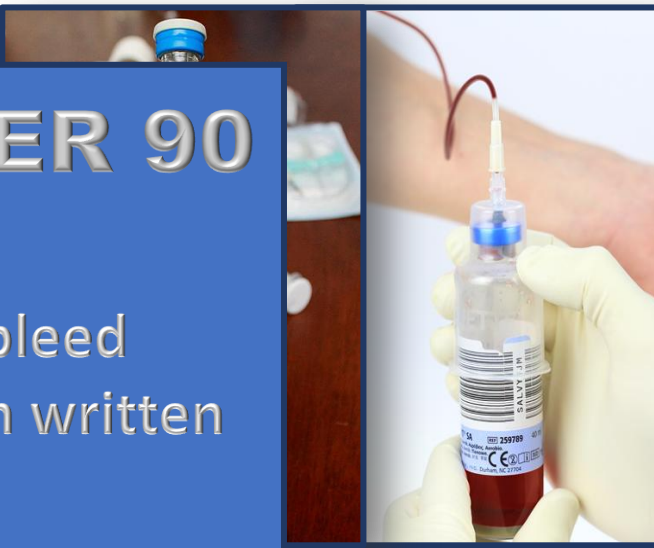
	1 HOUR	2 HOURS	3 HOURS	4 HOURS	>4 HOURS	NUMBER OF SETS
GLASS	46%	69%	77%	82%	18%	84
PLASTIC	71%	83%	91%	95%	5%	65
PLASTIC 2	79%	88%	94%	95%	5%	82
JAN 1						
JAN 2						
SEPT						



1 hour turnaround

DELAYS OF OVER 90 MINUTES

- Patient difficult to bleed
- History of fever so form written for next spike
- Endocarditis



Impact of optimising the pathway

NEONATAL 36 HOUR NEGATIVE BLOOD CULTURE REPORTS	
TIME FROM COLLECTION TO NEGATIVE REPORT	CUMULATIVE % OF NEGATIVE BLOOD CULTURE REPORTS ISSUED
36- 36.5 HOURS	43%
36-37 HOURS	81%
36-38 HOURS	100%

NEONATAL 36 HOUR NEGATIVE BLOOD CULTURE REPORTS

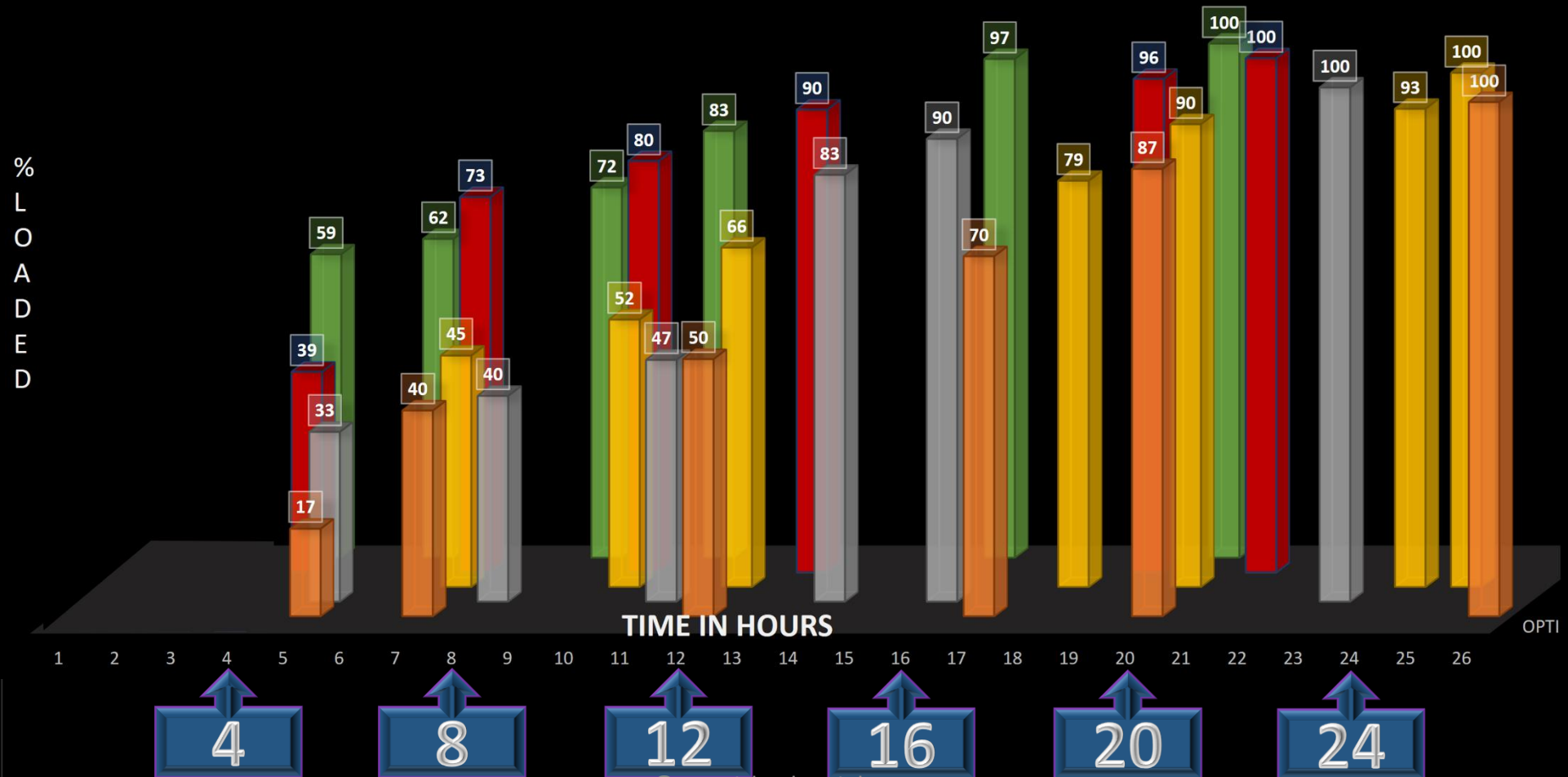
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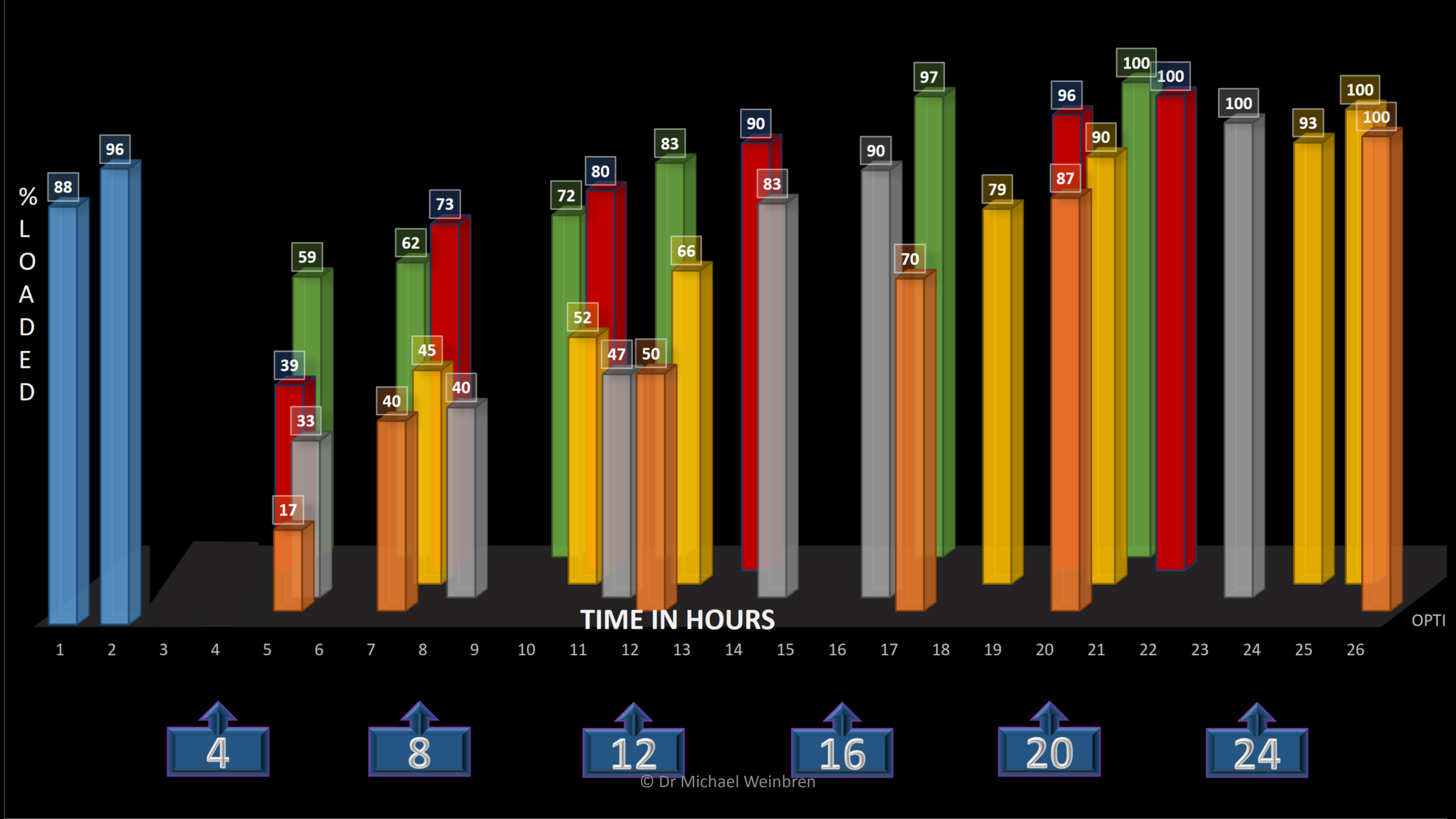
TIME FROM COLLECTION TO LOAD



DELAY BETWEEN COLLECTION AND PLACEMENT ON BLOOD CULTURE ANALYSER

■ OPTIMISED ■ HOSP A ■ HOSP B ■ HOSP C ■ HOSP D ■ HOSP E



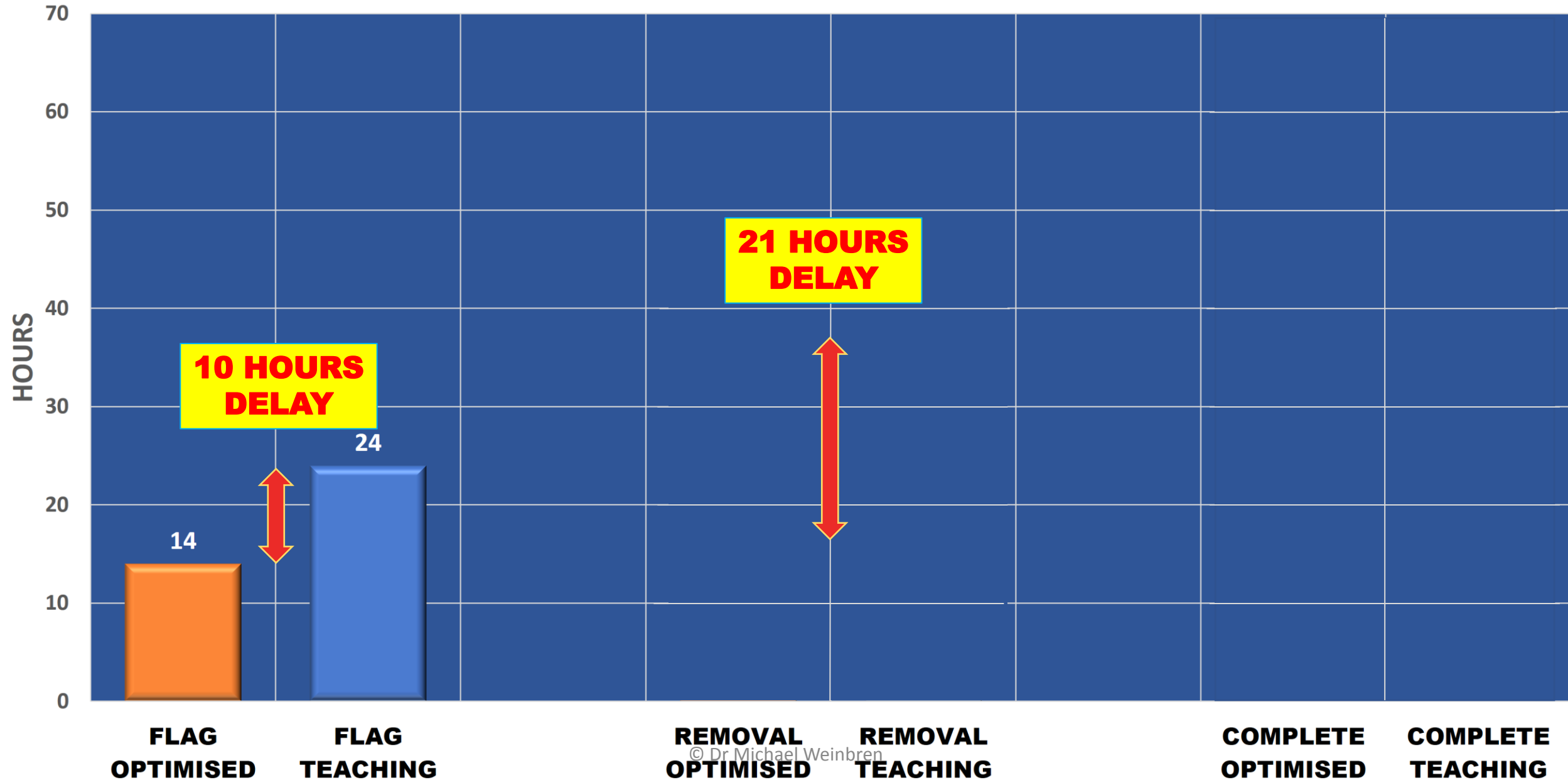


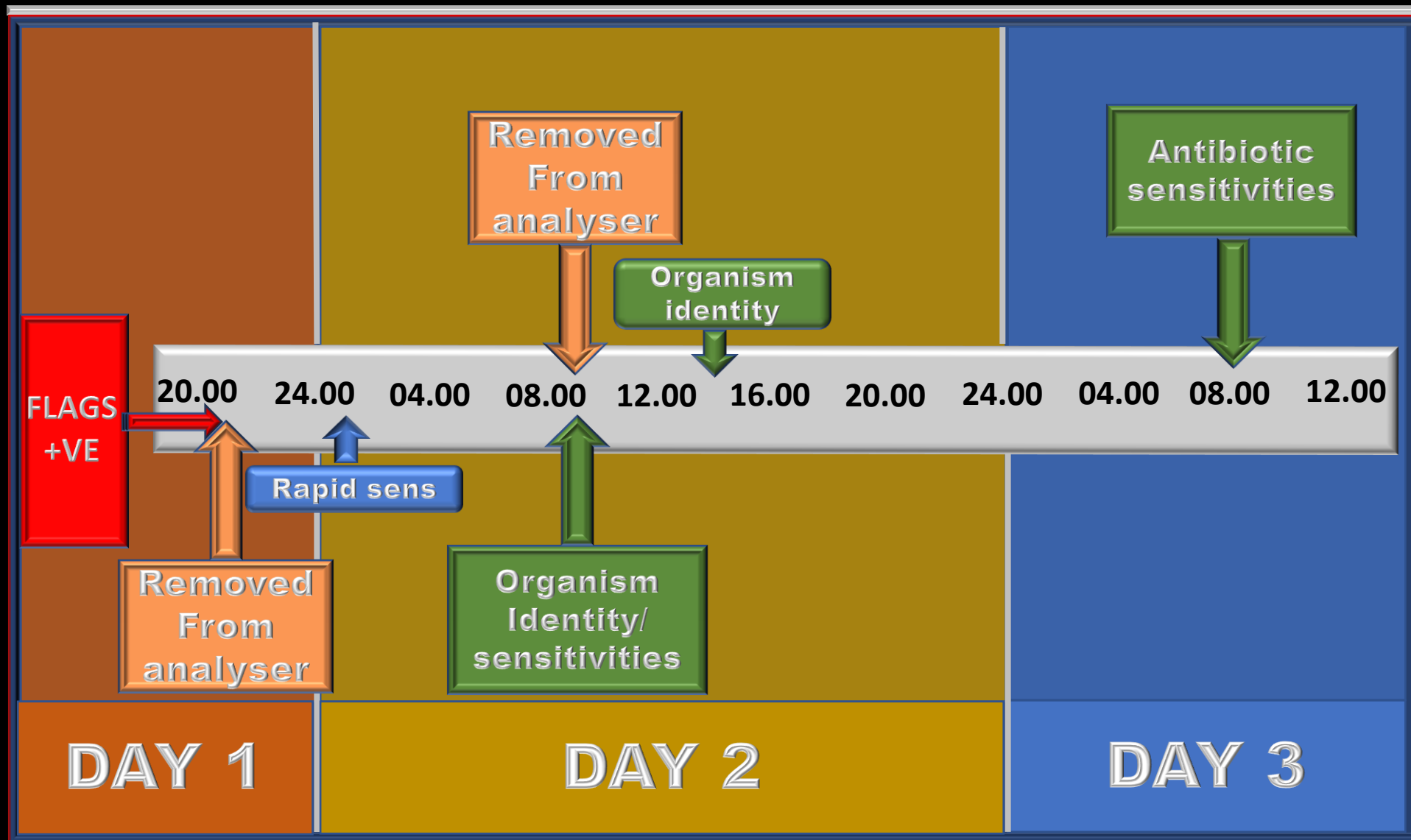
Average time in hours from collection to flagging positive and removal for *E. coli* positive blood culture

12.79

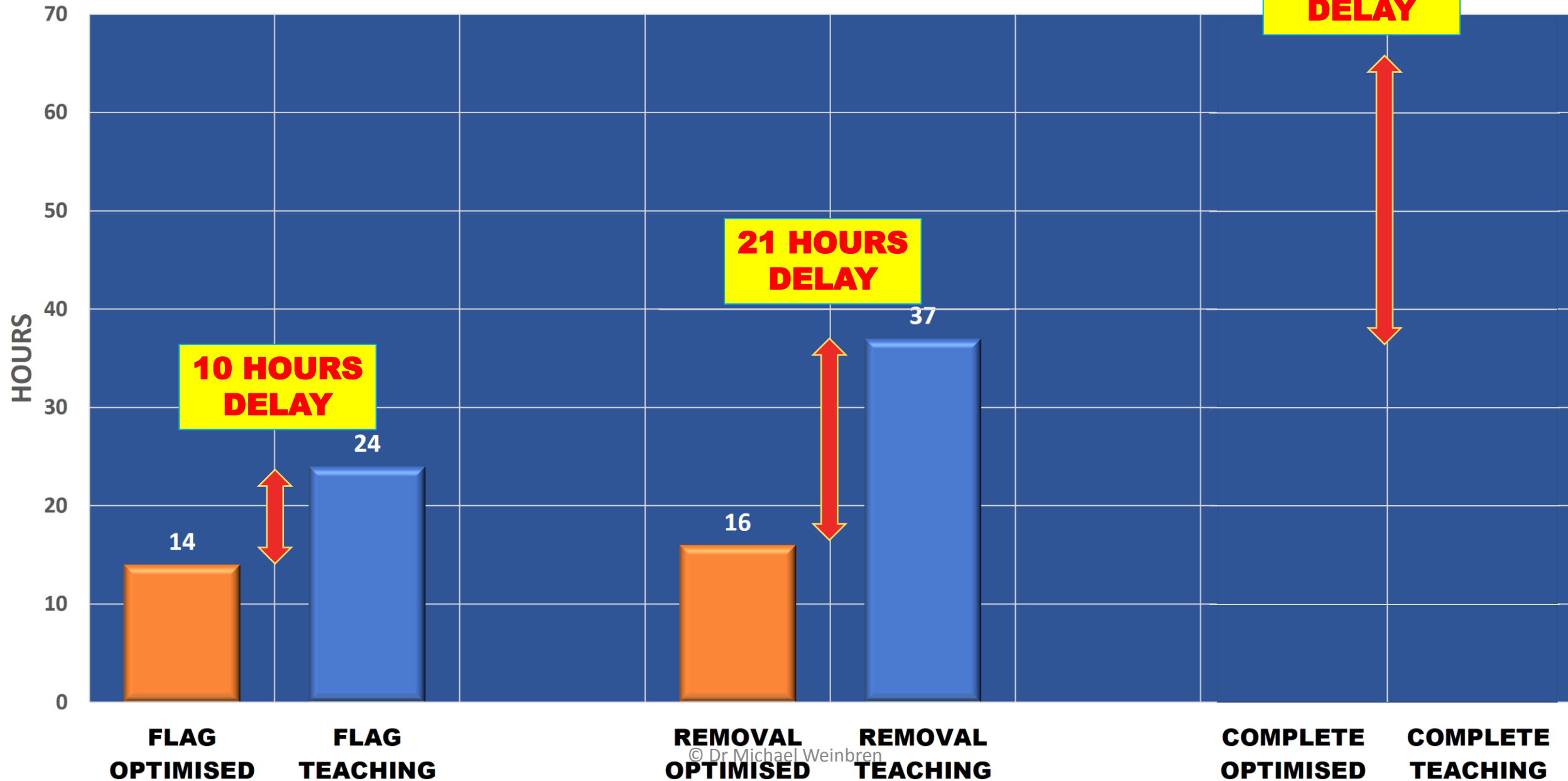
OPTIMISED

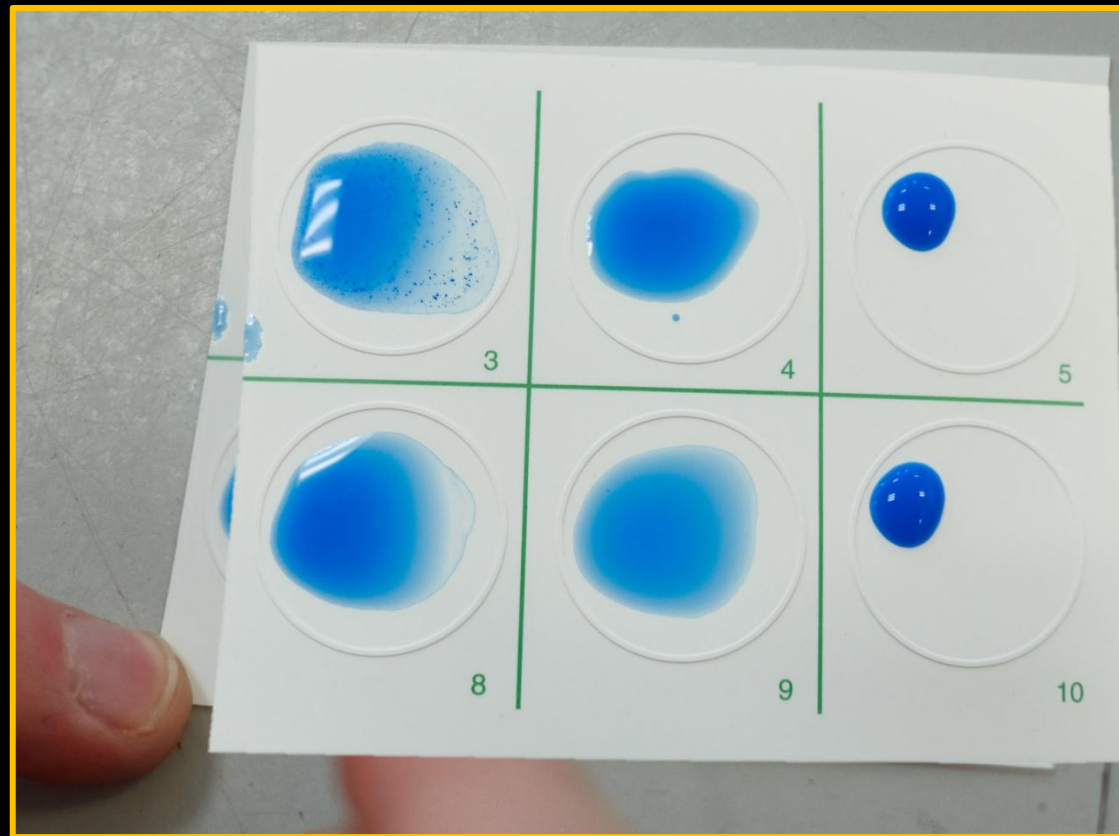
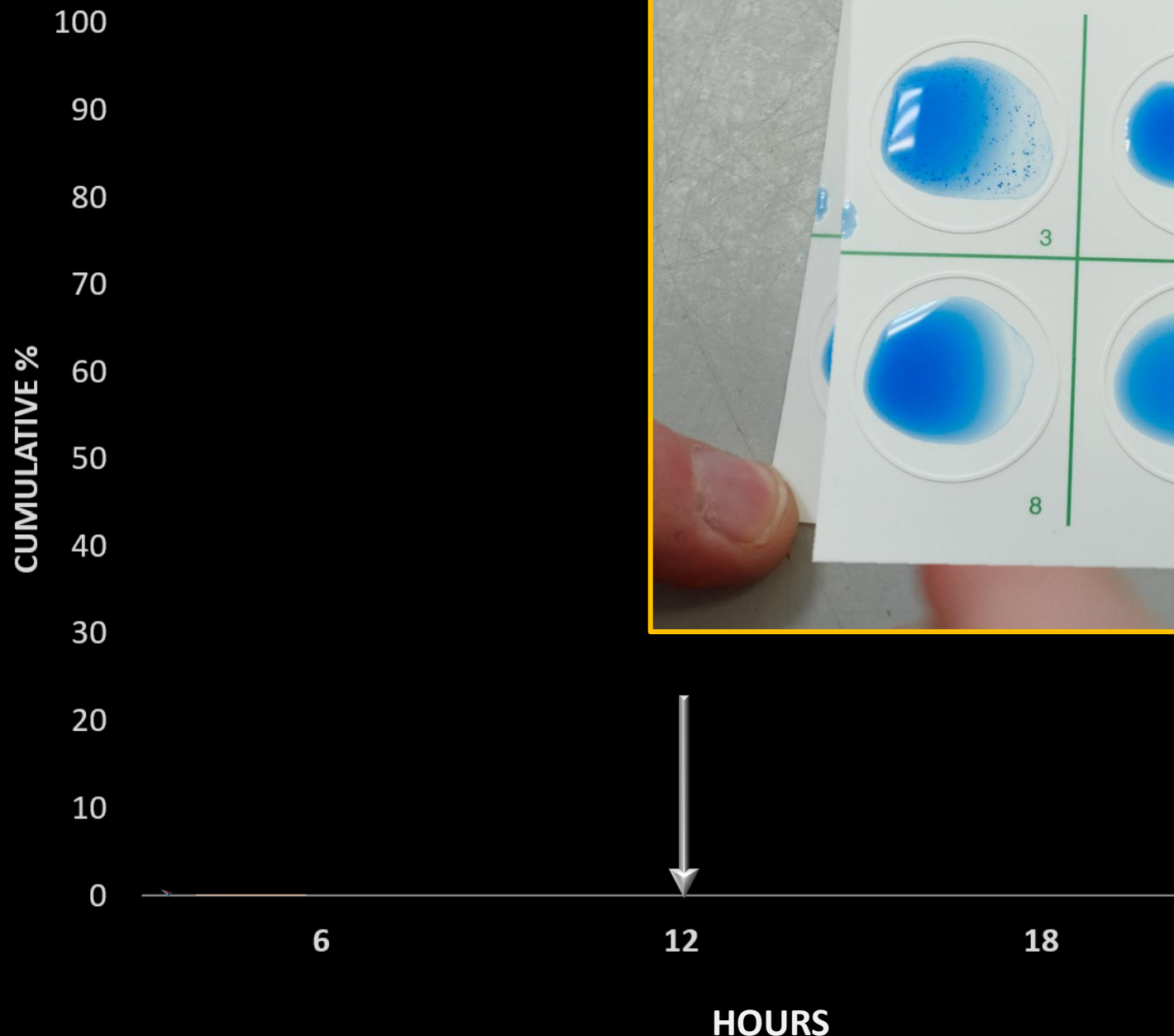
Time in hours to complete >85% of E. coli positive blood cultures at various stages in processing





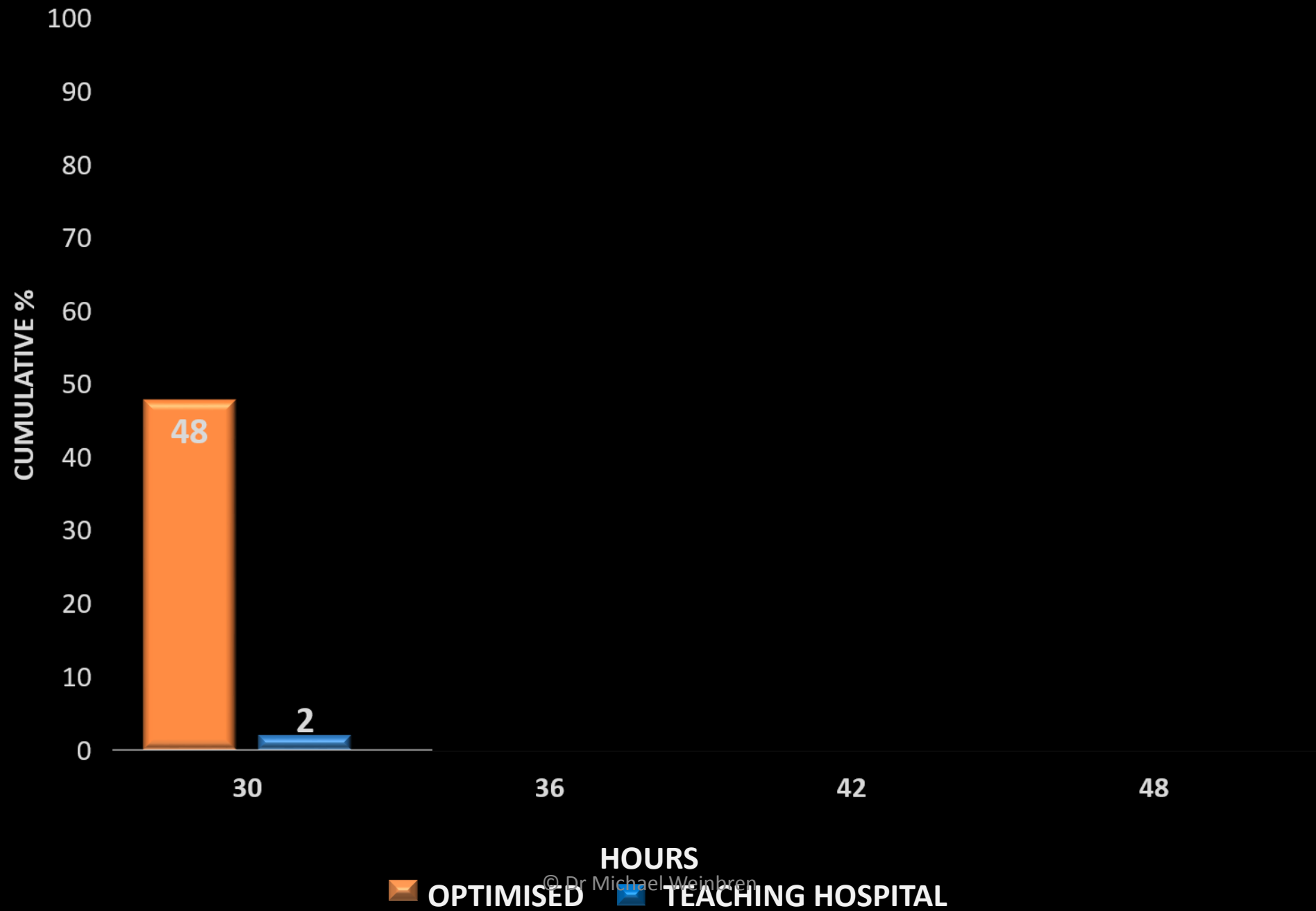
Time in hours to complete >85% of E. coli positive blood cultures at various stages in processing





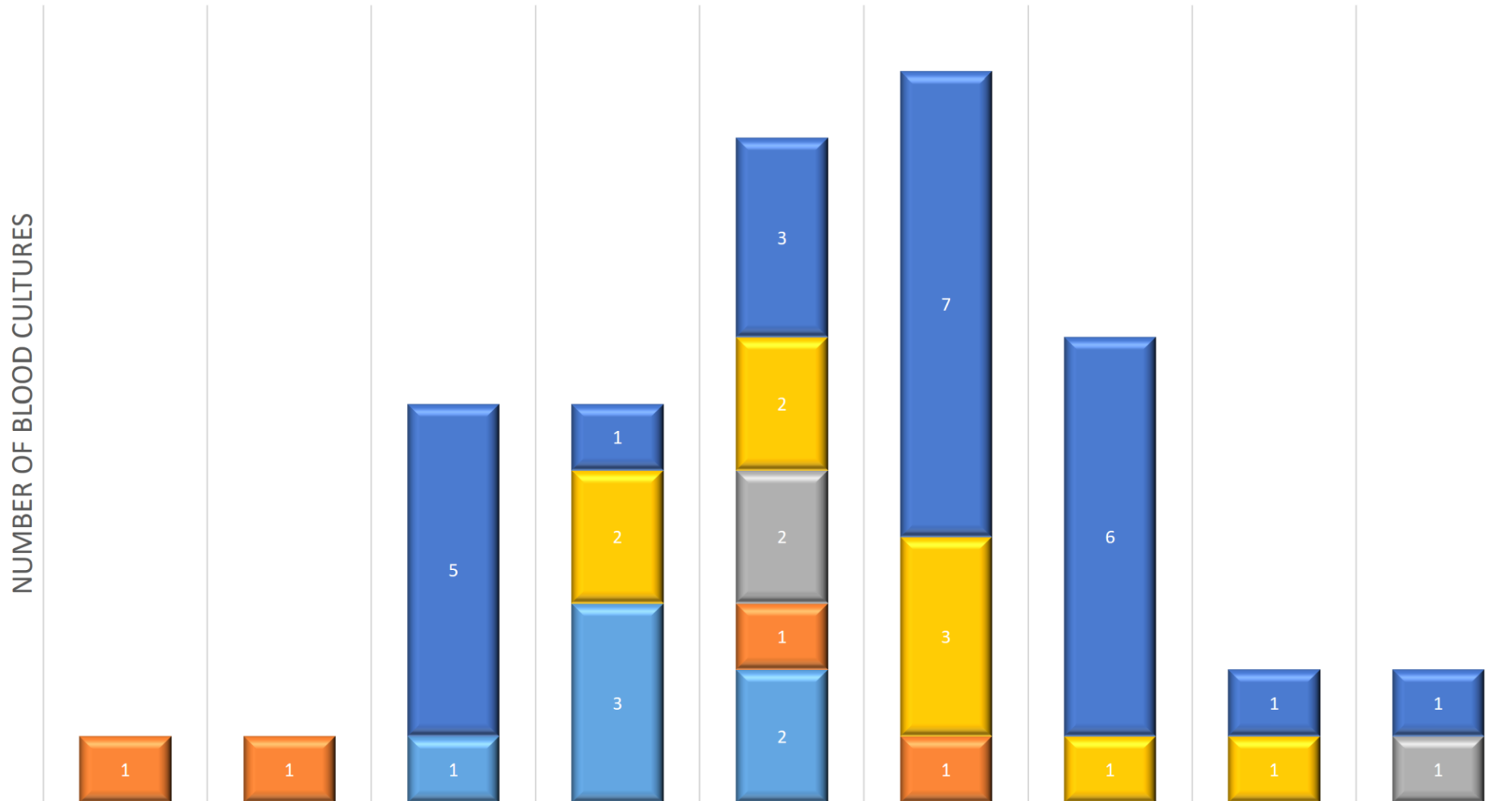
NHS Number
Hospital Number
Surname
Forename
Date of Birth
Address
Clinical Details
sepsis ? source, che
Signature
Tests Requested
Adult Blood cultures
For this request you v
1 x Adult Blood culture bottles

10/8/20
2/4
2/4



TIME FROM COLLECTION TO FLAGGING POSITIVE STREPTOCOCCI

GROUP A GROUP B GROUP C GROUP G S. PNEUMO



6-7
6

7-8
8

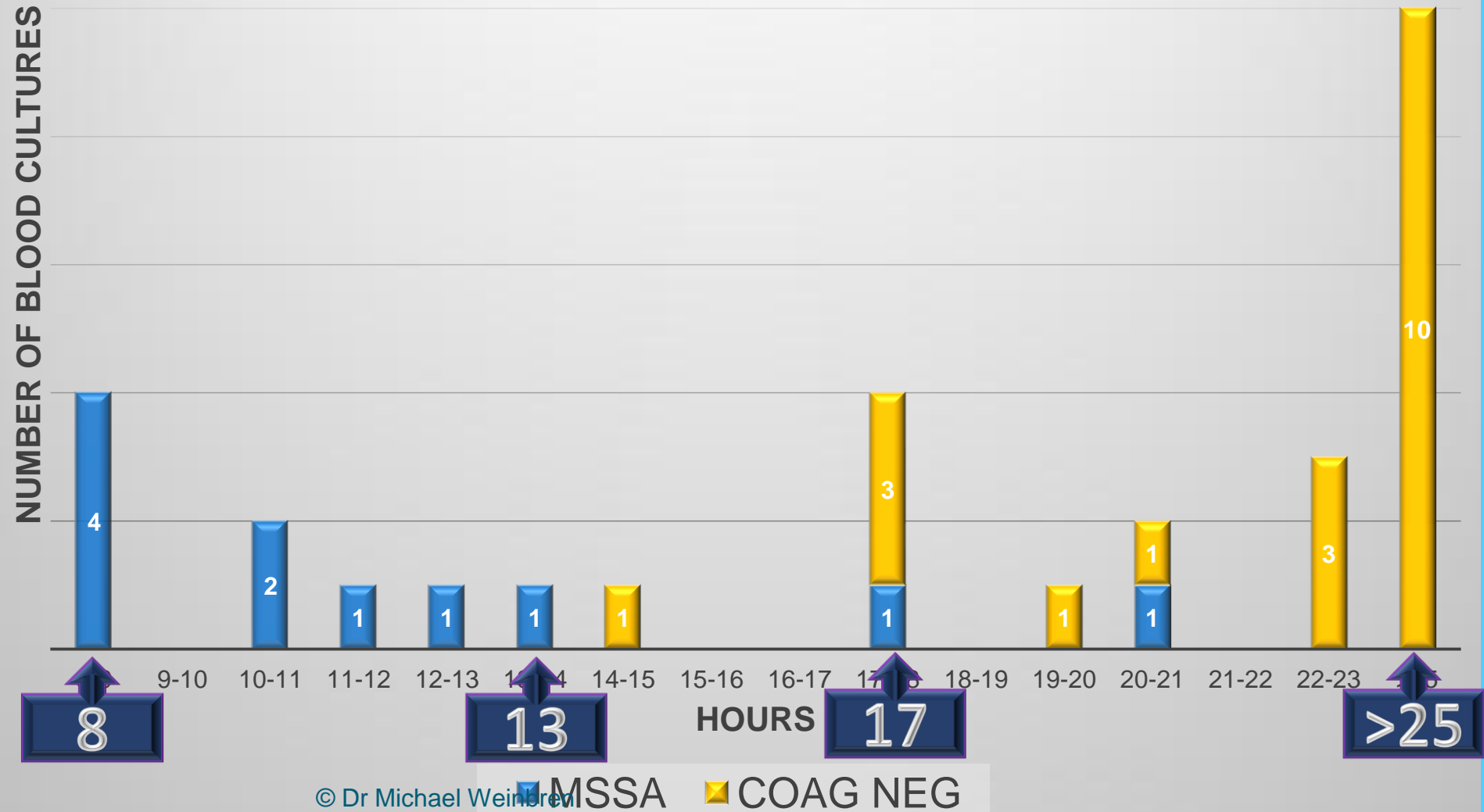
9-10
10

11-12
12

© Michael Weinbren

HOURS

TIME FROM COLLECTION TO FLAGGING POSITIVE STAPHYLOCOCCI



Clinical cases

- Mr RB
- ? Chest infection
- Oral clarithromycin
- 8hours 36 minutes- group G strep
- Teicoplanin and clindamycin (Clari res)
- Stormy next 48 hours
- Home day 6 -cellulitis

Benefits

- Real time 36 hour negative neonatal blood cultures
- Improved management of patients- earlier correction of deficiencies in empirical antibiotic therapy
- Earlier de-escalation of antibiotics
- Real time results gives clue to relevance of organism
- Minimises risks of organisms auto-lysing
- Improvements mostly achieved through better use of existing resources
- Essential pathway is optimised to maximise benefits of new rapid diagnostic methodology

EMPIRICAL TREATMENT OF GRAM NEGATIVE SEPSIS

ANTIBIOTIC	AVG % SENS	MIN % SENS	MAX %SENS
CO-AMOX	57.0	49.8	65.2
GENT	92.0	85.0	94.5
PIP-TAZO	87.2	81.7	94.2
CIP	86.4	81.4	93.0
MERO	99.4	98.8	100.0

Data courtesy of Naomi Thompson

Analysis (106 positive blood cultures)

- 33% patients (36) not on an effective antibiotic
- 6 patients not on antibiotics

Gram-negative bacteraemia; a multi-centre prospective evaluation of empiric antibiotic therapy and outcome in English acute hospitals

ARTICLE *in* CLINICAL MICROBIOLOGY AND INFECTION · NOVEMBER 2015

Impact Factor: 5.77 · DOI: 10.1016/j.cmi.2015.10.034

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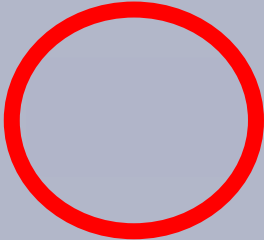
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34% not on an Effective agent

CORRECTION OF INITIAL ANTIBIOTIC THERAPY

	GRAM STAIN	ORGANISM IDENTITY	ANTIBIOTIC SENSITIVITIES (RESISTANCE)
NOT ON ONE EFFECTIVE ANTIBIOTIC			

Blood culture collection

Load times and critical processing times for >85% of E. coli Positive blood cultures for teaching hospital vs hospital with Optimised pathway

KEY



95% of bottles loaded



100% of bottles loaded



85% E. coli bottles Removed from analyser



85% E. coli positive Blood cultures completed processing

Optimised pathway

Teaching hospital

0

5

10

20

30

40

50

60

70

Time in hours

© Dr Michael Weinbren

Blood culture collection

Effect new technology could have on turnaround times of
1 Gram negative organisms (providing rapid sensitivities),
2 Gram positive organisms (providing organism ID)
Compared to an already optimised pathway

Optimised pathway plus new technology Gram negatives

1

Optimised pathway

0

5

10

20

30

40

50

60

70

Time in hours

KEY



95% of bottles loaded



100% of bottles loaded



85% E. coli bottles
Removed from analyser



85% E. coli positive
Blood cultures completed
processing



>50% of significant Gram
positive blood cultures



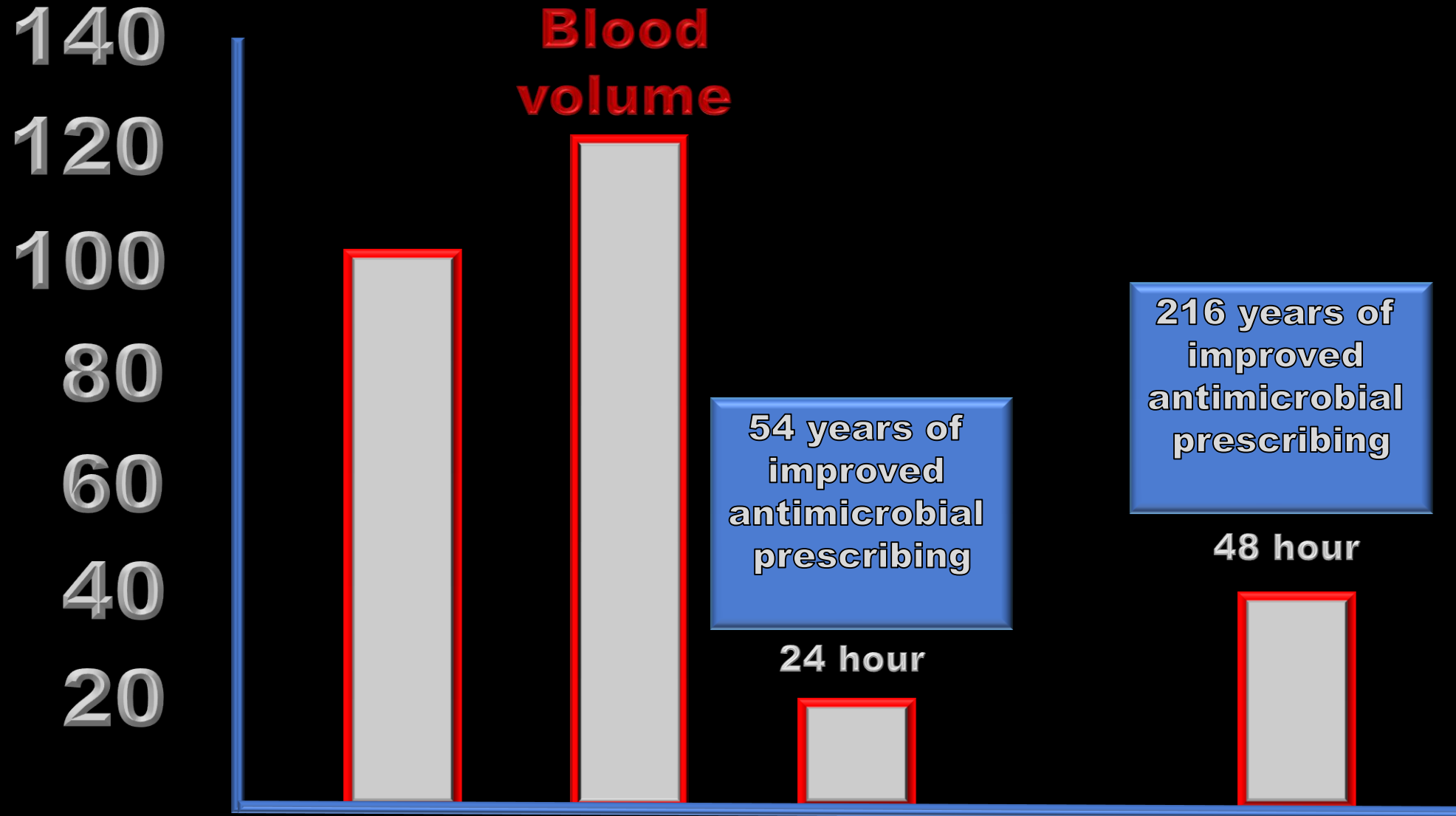
New tech rapid ID in
1-2 hours



Time of conventional
ID

Thousands of
bacteraemias / year

Diagnostic stewardship





0.00

4.05



Case 2-

MRS DM DOB 28/12/1930

Admitted with sepsis ? Biliary/ urinary source

Started on co-amox but changed to tazocin as systemically unwell

Blood culture collected 8/11/2015 at 13.32

On blood culture machine at 14.49

Flagged positive 09/11/15 03.04

Subcultured 03.20

09.00

09.45 changed to meropenem

15.00 E. coli gent and tazocin resistant



What do these have in common?



The Sepsis Six



1. Give high-flow oxygen
2. Take blood cultures
3. Give IV antibiotics
4. Start IV fluid resuscitation
5. Check lactate
6. Monitor hourly urine output

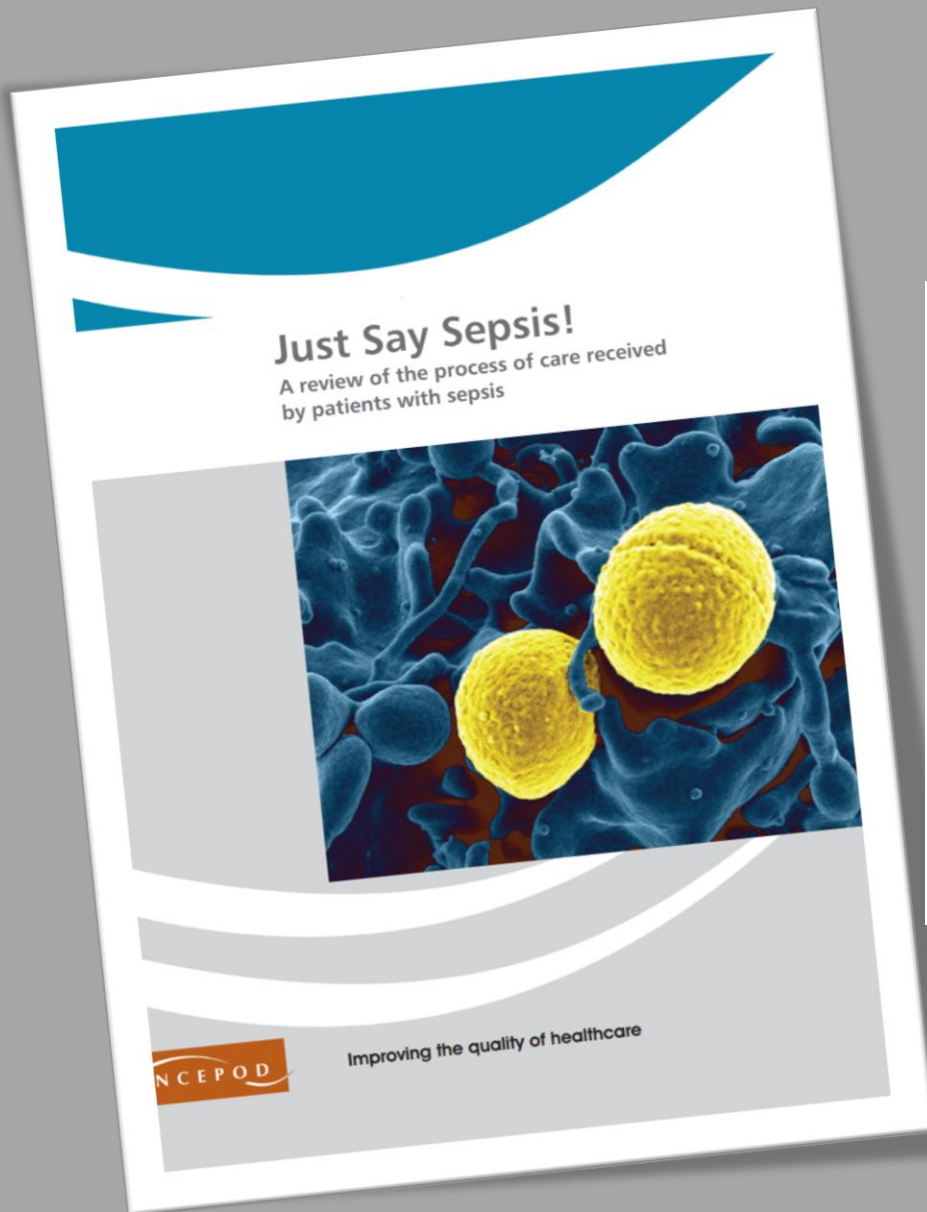
within one hour

..plus Critical Care support to complete EGDT

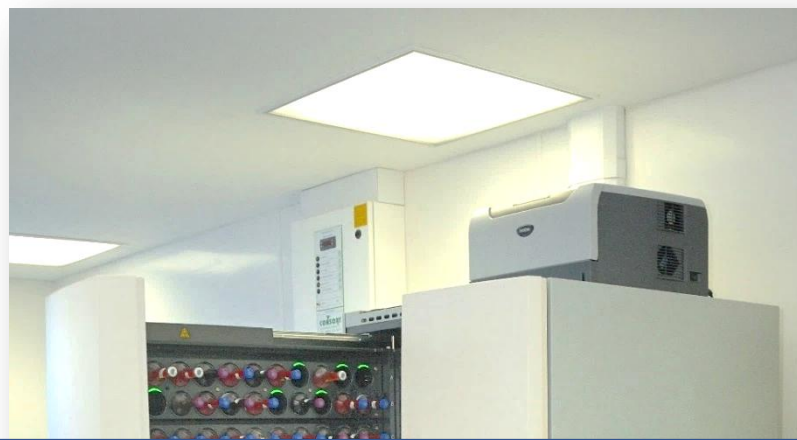
via non-rebreathe bag
and **consider source control**
according to local protocol
Hartmann's or equivalent

consider catheterisation



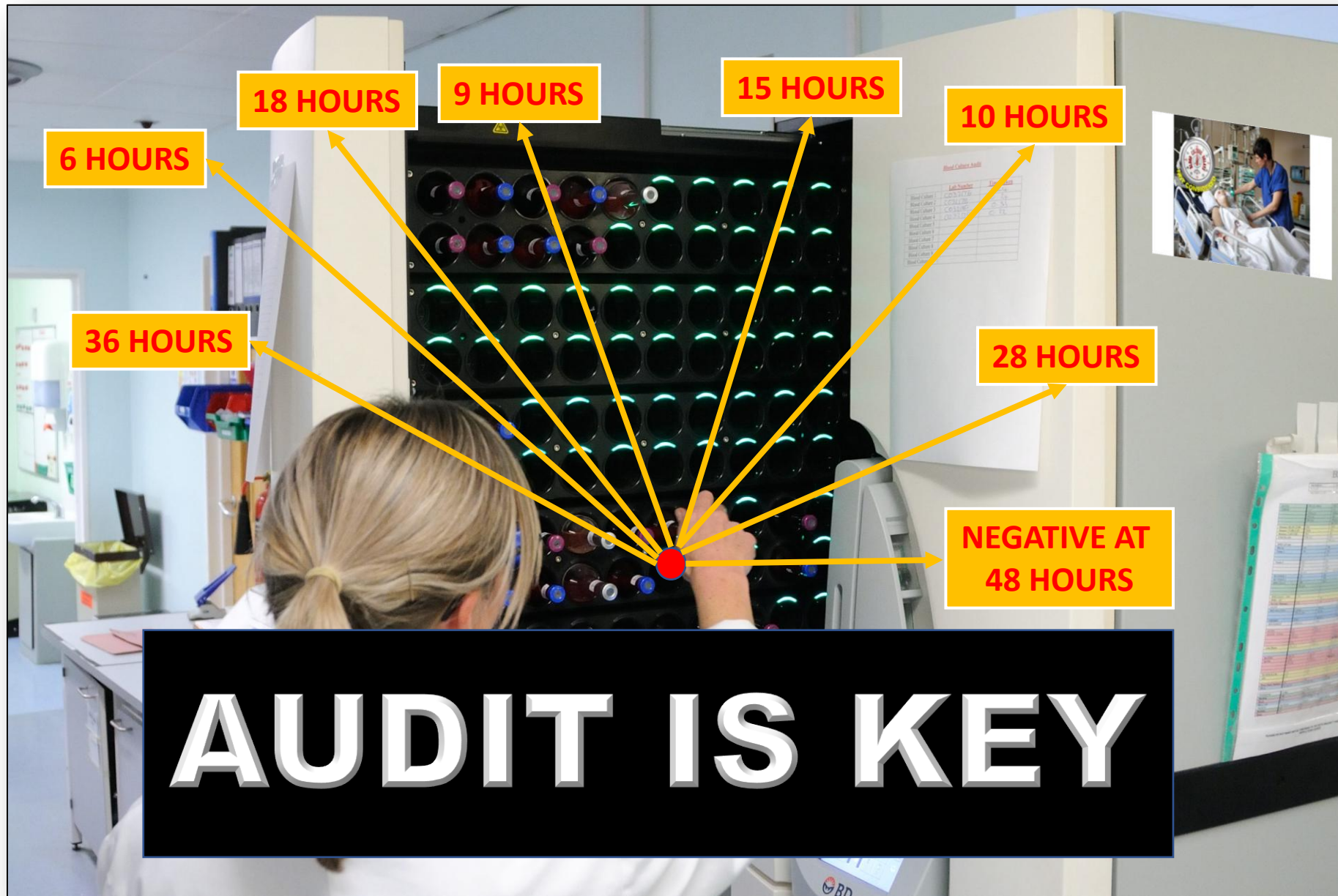


**WAS A
BLOOD
CULTURE
COLLECTED?**





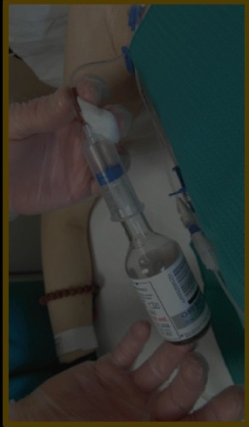






COLLECTION

1



Right patient / volume
of blood to analyser
with minimum delay

PROCESSING

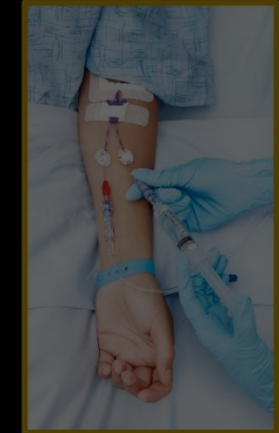
2



SHORTEST TIME
TO KEY RESULT

MANAGEMENT

3



USE KEY RESULT
TO MANAGE
PATIENT WITH
MIN DELAY

ISO STANDARDS

One frequent pitfall is that accreditation can focus excessively on technical details rather than on patient value, which results in an inappropriate clinical service, i.e. quality being disconnected from the end-point target of improved patient care.

B. Lamy CMI 2018

FINISH

A Healthcare Improvement Intervention Combining Nucleic Acid Microarray Testing with Direct Physician Response for Management of *Staphylococcus aureus* Bacteremia

Joshua C. Eby^{1#}, Morgan M. Richey², James A. Platts-Mills¹, Amy J. Mathers^{1,3}, Wendy M. Novicoff², Heather L. Cox^{1,4}

Results: 106 pre-intervention and 120 post-intervention subjects were assessed. Time to ID consultation after notification of a positive blood culture decreased 26.0 hours (95% CI 45.1 to 7.1 hours, $p<0.001$) post-intervention compared with pre-intervention. Time to initiation of targeted antibiotic decreased by a mean 21.2 hours (95% CI 31.4 to 11.0 hours, $p<0.001$) and time to targeted antibiotics for methicillin-sensitive *S. aureus* by a mean 40.7 hours (95% CI 58.0 to 23.5 hours, $p<0.001$). The intervention was associated with lower in-hospital (13.2% to 5.8%, $p=0.047$) and 30-day mortality (17.9% to 8.3%, $p=0.025$).

Conclusions: Compared to an ASP-directed response to traditionally detected SAB, an efficient physician response to NAM was associated with improved care and outcomes for SAB.

Individualized Approaches Are Needed for Optimized Blood Cultures

Ritu Banerjee,¹ Volkan Özenci,² and Robin Patel^{3,4}

¹Department of Pediatric Infections Diseases, Vanderbilt University, Nashville, Tennessee; ²Division of Clinical Microbiology, Karolinska Institutet, Karolinska University Hospital, Huddinge, Stockholm, Sweden; ³Division of Clinical Microbiology, Department of Laboratory Medicine and Pathology, and ⁴Division of Infectious Diseases, Department of Medicine, Mayo Clinic, Rochester, Minnesota

Before laboratories consider offering rapid matrix-assisted laser desorption/ionization time-of-flight mass spectrometry (MALDI-ToF MS) or expensive rapid panel-based molecular BC diagnostics, they should optimize preanalytical, analytical, and postanalytical processes and procedures surrounding BC systems.

The technology exists.

**It is only our imaginations,
beliefs which limit us**

Summary

Blood sciences made this all possible

