

# Breakthrough Sepsis Pathogen Detection

Corporate Presentation  
August 2019  
(NASDAQ: TTOO)

# Forward-Looking Statements

This presentation contains forward-looking statements. Such statements reflect the current views of senior management of T2 Biosystems, Inc. ("we", "us", "our", "T2", "T2 Biosystems" or the "Company") and include those about T2's goals, strategies, plans, objectives, prospects, milestones, future operations, business and industry, anticipated product benefits, future events and conditions and potential scenarios. Such statements and those that include the words "expect," "intend," "plan," "believe," "project," "forecast," "estimate," "may," "should," "anticipate" and similar statements of a future or forward-looking nature identify forward-looking statements for purposes of the federal securities laws or otherwise. Forward-looking statements address matters that involve risks and uncertainties. Each forward-looking statement is subject to risks and uncertainties that could cause actual results to differ materially from those expressed or implied in such statement, including, for example: (i) our status as an early commercial-stage company and expectation to incur losses in the future; (ii) our ability to obtain marketing authorization from the FDA or regulatory clearance for additional product candidates in the United States or abroad; (iii) the market acceptance of our technology; (iv) our ability to timely and successfully develop and commercialize existing and future product candidates; (v) our lengthy and variable sales cycle and lack of sales history; (vi) our ability to successfully manage growth; (vii) federal, state and foreign regulatory requirements; (viii) our uncertain future capital needs and ability to raise future capital; (ix) dependence on third parties; (x) recruiting, training and retaining key personnel; (xi) competitive factors; (xii) manufacturing and other product risks; (xii) risks related to intellectual property; and (xiii) other risk factors included in our annual report on form 10-K filed with the Securities and Exchange Commission (SEC) on March 14, 2019 and other documents we file with the SEC from time to time. Accordingly, there are or will be important factors that could cause our actual results to differ materially from those indicated in these statements. The statements made herein speak only as of the date of this presentation. We do not undertake, and specifically disclaim, any obligation to update any forward-looking statements contained in this presentation.

# Why Are We Here Today?

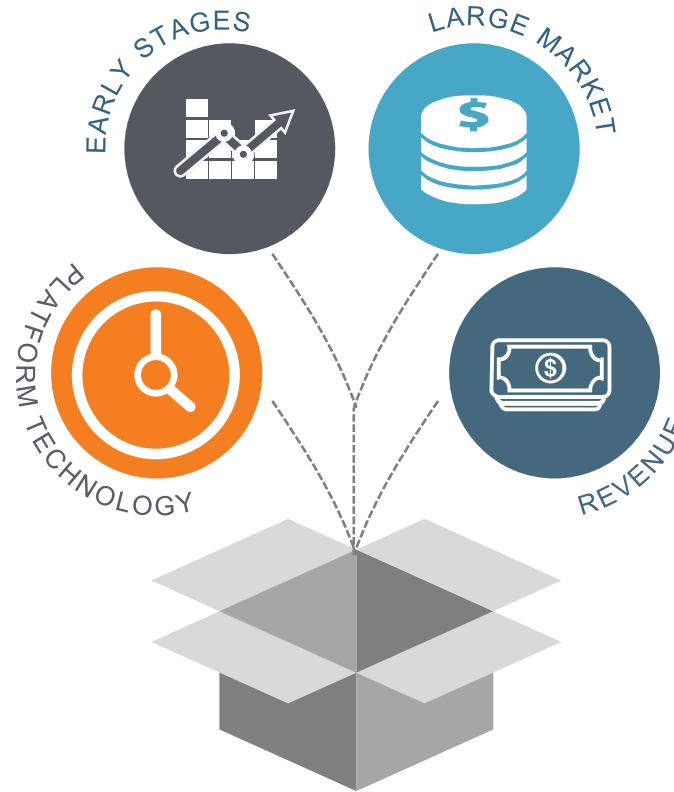
T2 has reached a tipping point for broad adoption of the T2Dx<sup>®</sup> technologies

## Early Stages

- Proven with T2Candida<sup>®</sup>
- Expanding with T2Bacteria<sup>®</sup>

## Platform Technology

- Market expansion over time



## Large Market

- Global unmet need

## Revenue

- Growth opportunity with attractive recurring model

# Sepsis is a Deadly and Frustrating Global Problem

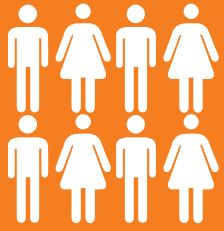
A recognized, but unsolved global crisis



1 death every  
**5 seconds**  
(more or less)

# Sepsis is a Deadly and Frustrating Global Problem

A critical part of the solution is now available



Potentially  
**>40,000**  
preventable deaths in  
the U.S. with T2



# The Facts About Sepsis

Most expensive hospital-treated condition in the U.S.



Contributes to  
1 in 2-3 hospital  
deaths<sup>1</sup>



Representing  
\$27B in U.S.  
healthcare  
costs<sup>2,3</sup>



**Claims more  
lives than breast  
cancer, prostate  
cancer and  
AIDS, combined<sup>4</sup>**



1 in 5 surviving  
sepsis patients  
die within 2 years  
due to sepsis<sup>5</sup>



Kills ~250,000  
Americans  
annually and ~6  
million people  
worldwide<sup>6,7</sup>



Most prevalent  
and costly cause  
of hospital  
readmissions<sup>8</sup>

1. Liu, V., Escobar, G. J., Greene, J. D., et al. (2014). Hospital deaths in patients with sepsis from 2 independent cohorts. *Jama*, 312(1), 90-92.

2. Torio, C. M. and Moore, B. J. (2016). Statistical Brief# 204. Healthcare Cost and Utilization Project (HCUP). May.

3. McDermott, K. W., Elixhauser, A., Sun, R. (2017). Statistical Brief# 225. Healthcare Cost and Utilization Project (HCUP). June.

4. National Institute of General Medical Sciences. National Institutes of Health. Sepsis fact sheet. 2014.

5. Prescott, H. C., Osterholzer, J. J., Langa, K.M., et al. (2016). Late mortality after sepsis: propensity matched cohort study.

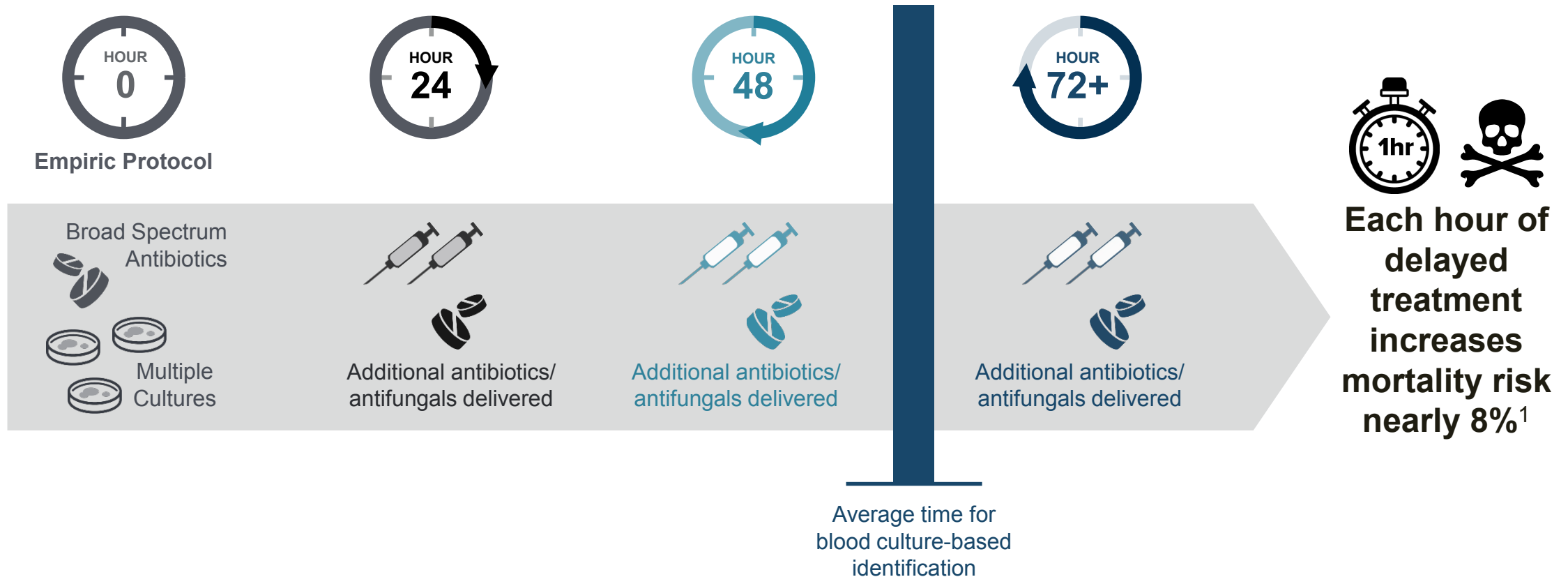
6. Centers for Disease Control and Prevention.

7. Gilbert, J. A. (2018). Sepsis care bundles: a work in progress. *The Lancet Respiratory Medicine*.

8. Mayr, F. B., Talisa, V. B., Balakumar, V., et al. (2017). Proportion and cost of unplanned 30-day readmissions after sepsis compared with other medical conditions. *JAMA*, 317(5), 530-531.

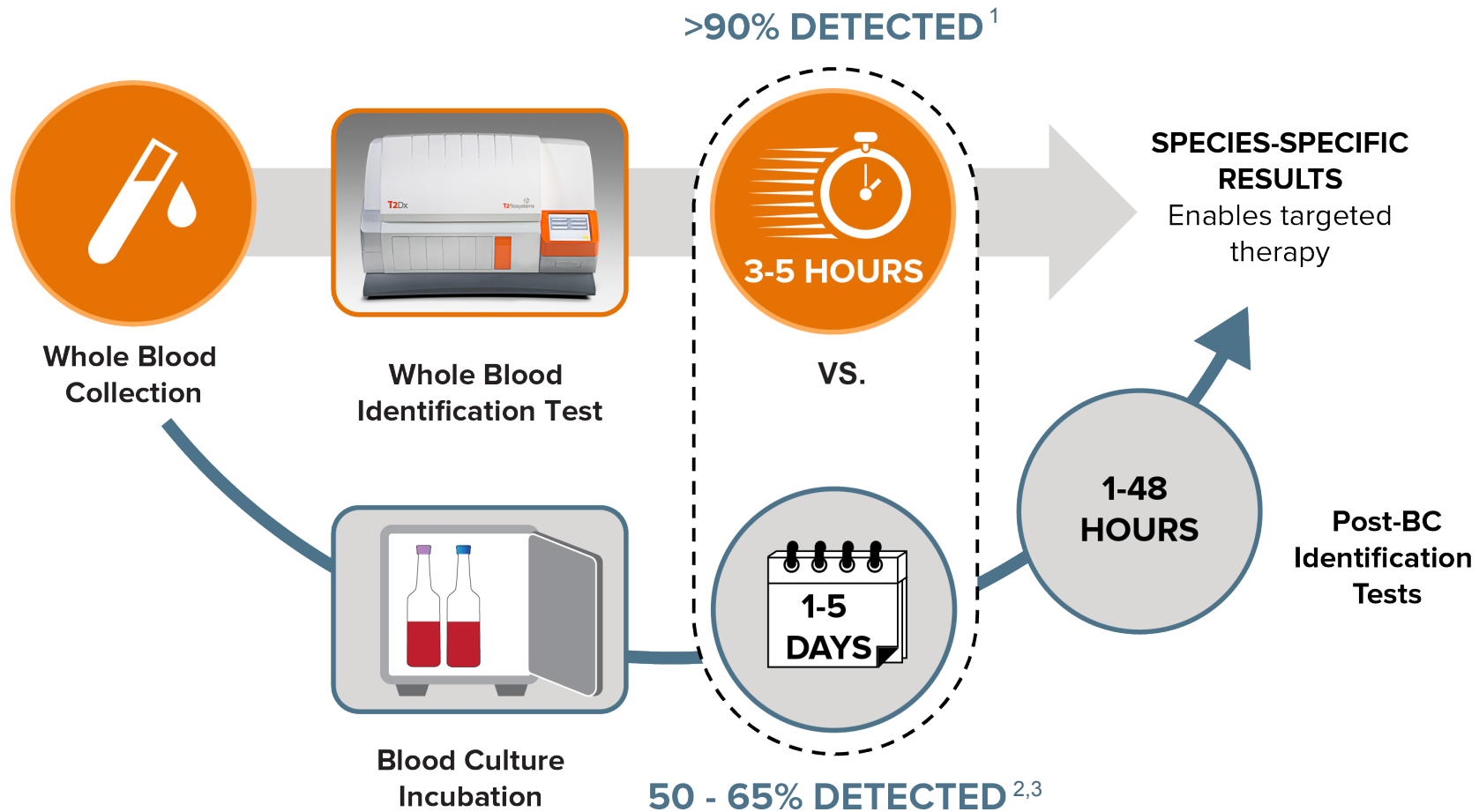
# Sepsis Poses an Hourly Challenge that Relies on Probability-Based Protocols

Patient journey: Current pathway and empiric “process”



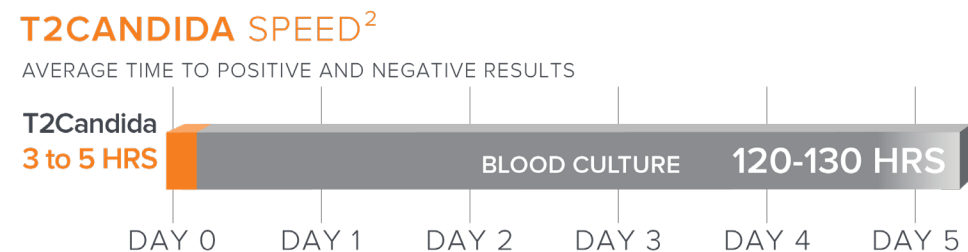
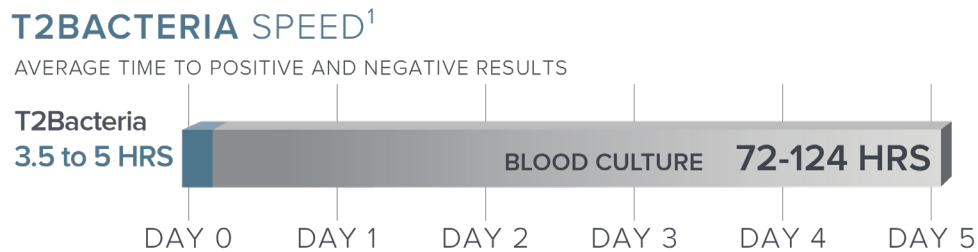
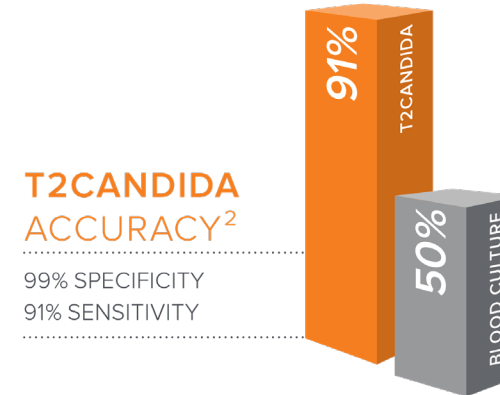
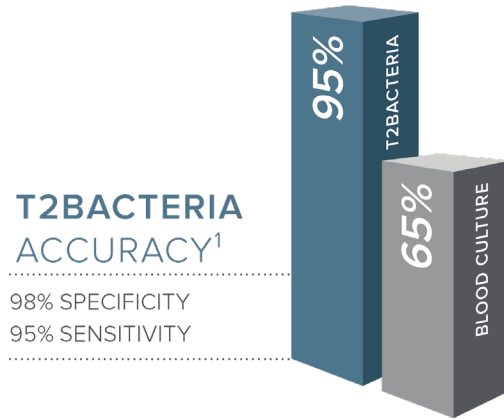
# T2MR: New Standard in Detecting Sepsis Causing Pathogens

T2Direct Diagnostics provide faster and more accurate detection



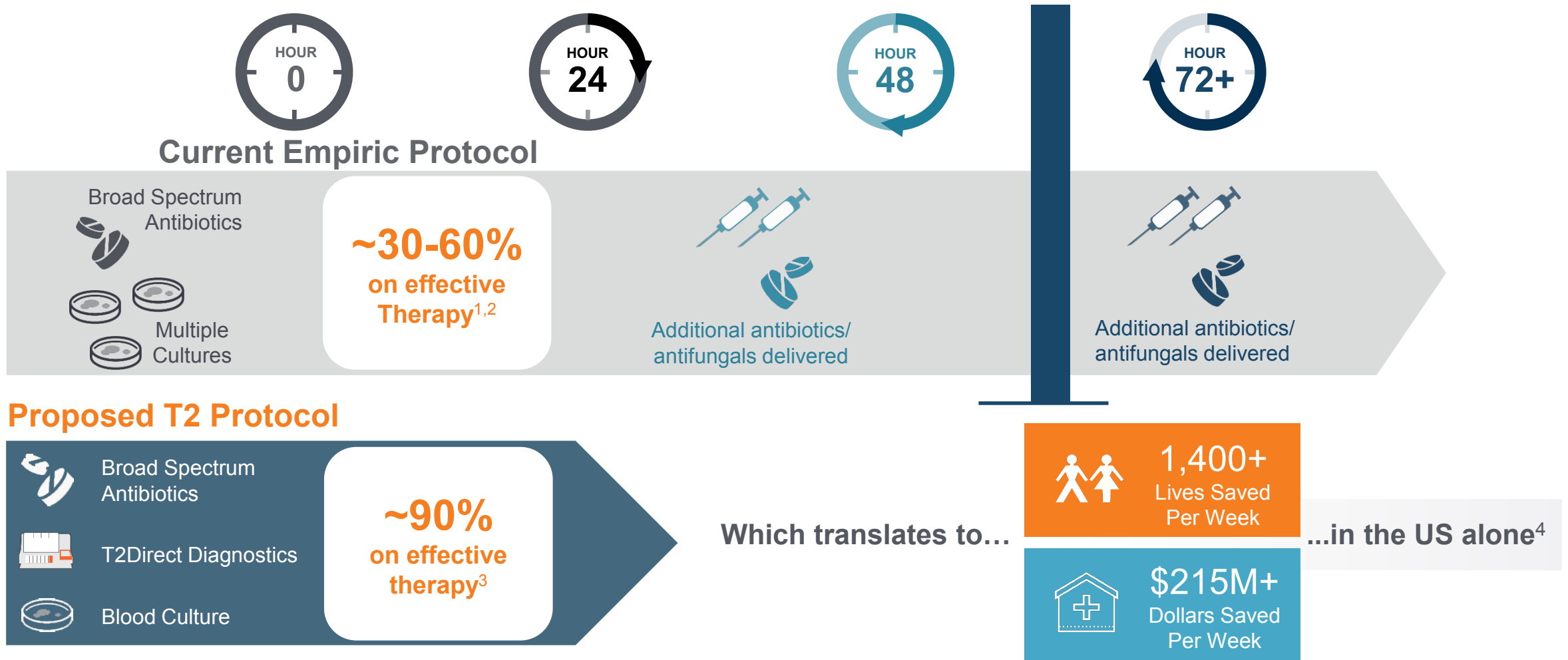
1. Mylonakis, E., Clancy, C. J., Ostrosky-Zeichner, L., et al. (2015). T2 magnetic resonance assay for the rapid diagnosis of candidemia in whole blood: a clinical trial. *Clinical Infectious Diseases*, ciu959.
2. Clancy, C. J., & Nguyen, M. H. (2013). Finding the "missing 50%" of invasive candidiasis: how nonculture diagnostics will improve understanding of disease spectrum and transform patient care. *Clinical infectious diseases*, 56(9), 1284-1292.
3. Cockerill III, F. R., Wilson, J. W., Vetter, E.A., et al. (2004). Optimal testing parameters for blood cultures. *Clinical Infectious Diseases*, 38(12), 1724-1730.

# The Blood Culture Divide



1. T2Bacteria Pivotal Clinical Study. Overall average sensitivity of 90% in prospective arm and 97% PPA in contrived arm.  
2. Mylonakis, E., Clancy, C. J., Ostrosky-Zeichner, L., et. al. (2015). T2 magnetic resonance assay for the rapid diagnosis of candidemia in whole blood: a clinical trial. Clinical infectious diseases, ciu959.

# A Simple Change, an Immense Impact



1. T2Bacteria Clinical Pivotal Trial Data.

2. Buehler, S. S., Madison, B., Snyder, S. R., et al. (2016). Effectiveness of practices to increase timeliness of providing targeted therapy for inpatients with bloodstream infections: a laboratory medicine best practices systematic review and meta-analysis. Clinical microbiology reviews, 29(1), 59-103.

3. Kumar, A., Ellis, P., Arabi, Y., et al. (2009). Initiation of inappropriate antimicrobial therapy results in a fivefold reduction of survival in human septic shock. CHEST Journal, 136(5), 1237-1248.

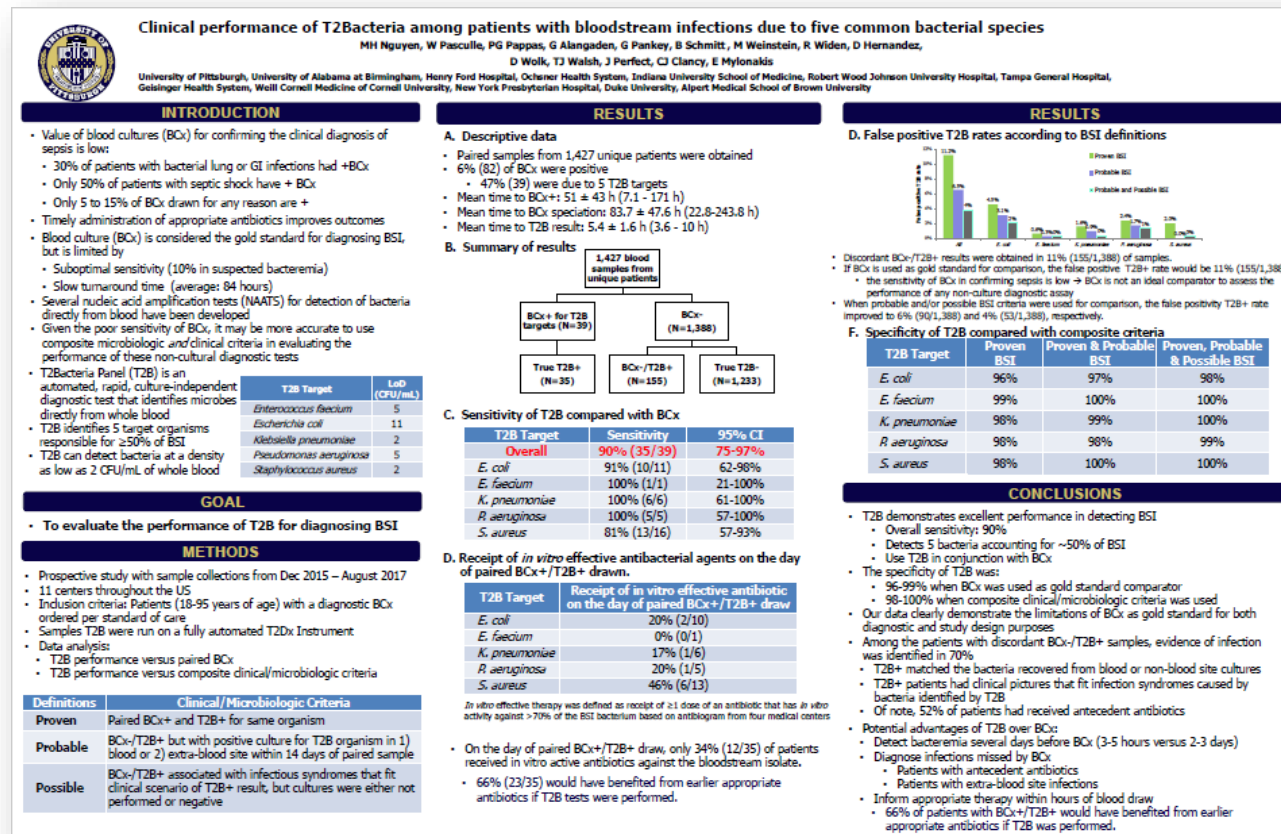
4. Represents the potential healthcare savings and lives saved using the T2Direct Diagnostic to test high risk patients based on assumed levels of total annual patients assuming all high-risk sepsis patients are tested with T2Direct Diagnostics and assuming (i) 90% of high risk patients receive appropriate therapy within hours of the presentation of symptoms, (ii) a 50% mortality rate reduction for patients who receive rapid appropriate therapy, and (iii) that each new detected patient saves \$22,800. This slide contains T2's estimates, which are not based on historical results and constitute forward-looking statements that are subject to risks and uncertainties that could cause actual results to differ materially from those expressed or implied by such statement.

# T2Bacteria Pivotal Data Published in Annals of Internal Medicine

1,427 patient samples collected across 11 hospitals

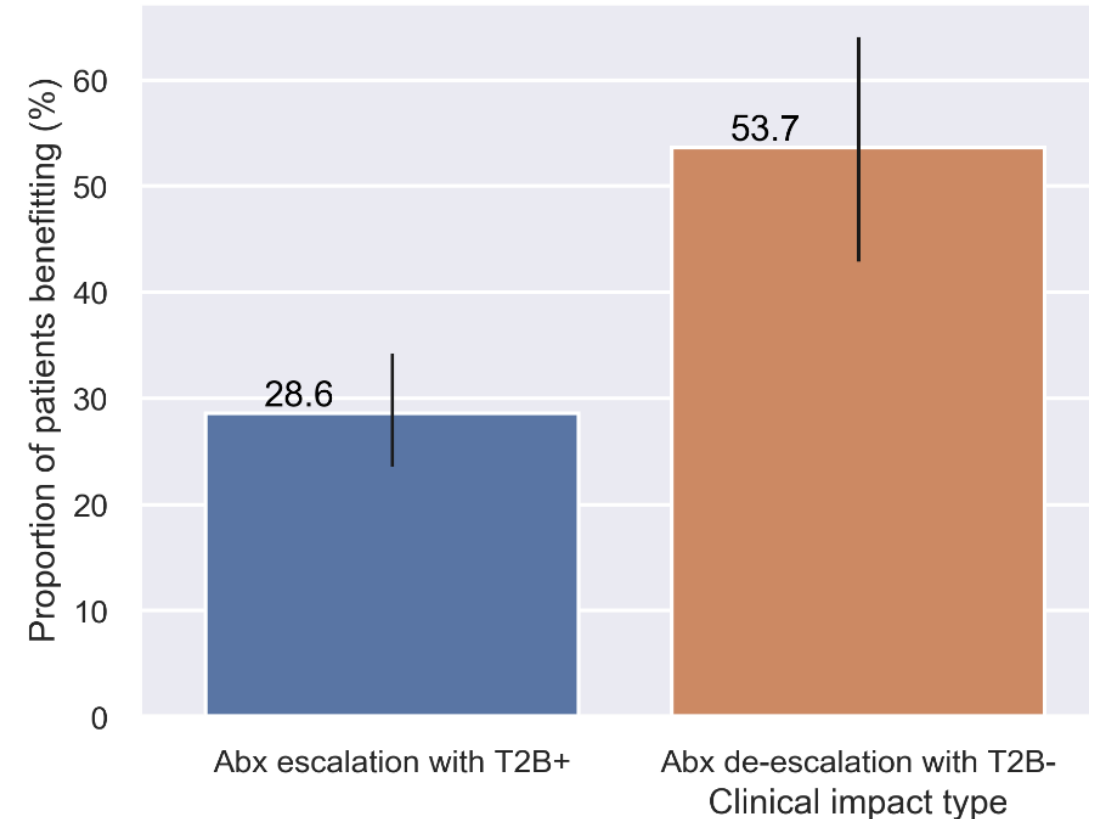
## Comparisons to Blood Culture:

- Detected **69 patient** infections not detected by culture
- Provided results more than **2.5 days faster than culture** (5.4 hours)
- 68%** of patients with a BSI confirmed by T2 and blood culture could have benefitted from earlier rapid diagnostic result
- Noted advantage in detecting infected patients on antibiotics who were missed by blood culture



# Emerging T2Bacteria Clinical Utility Data

- Cases from 7 ED & ICU studies aggregated here.
- 125/299 (42%) cases showed substantial clinical benefit of those evaluated under chart review
  - T2+: faster time to effective therapy
  - T2-: earlier antibiotic de-escalation
  - T2+: avoiding premature ED discharge/readmission
- 16% T2Bacteria positivity rate, range 13%-35% across all patients tested
- 2 cases of T2Bacteria helping avoid premature discharge and readmission in the ED
- A significant percentage of patients benefit from T2Bacteria results.



# New Technology Add-on Payment by CMS



- The T2Bacteria Panel is the first, and only, in-vitro diagnostic test to ever receive approval from the US Centers for Medicare & Medicaid Services (CMS) for New Technology Add-on Payment (NTAP)
- CMS is the single largest payer for health care in the US
- The payment is in addition to the current diagnosis-related group (MS-DRG) reimbursement

## CMS Found that T2Bacteria Met its Criteria For Additional Payment For Medical Advances

1

New

2

Cases are Inadequately  
Paid Under the Existing  
MS-DRG System

3

Significant  
Clinical Improvement  
Over Existing  
Technologies

*“The T2Bacteria Test Panel represents a substantial clinical improvement over existing technologies because it reduces the proportion of patients on inappropriate therapy, thus reducing the rate of subsequent diagnostic or therapeutic intervention as well as length of stay and mortality rates caused by sepsis causing bacterial infections.”*

-United States CMS FY 2020 inpatient prospective payments system final rule

# T2Candida Panel is Changing Treatment Protocols

Growing number of real-world T2Candida success stories



- Study demonstrated \$2.3M in annual hospital savings
- Reduced median ICU length of stay by 7 days; overall stay by 4 days
- Most negative patients had antifungals discontinued or de-escalated saving \$\$.<sup>1</sup>



- Median length of stay reduced by 7 days
- Unnecessary antifungal therapy was avoided in >50% of patients
- Average net antifungal savings of ~\$195 for every patient tested<sup>2</sup>



- Pharmacy savings of ~\$280 per patient
- T2Candida detected 56% more positive patients than blood culture<sup>3</sup>



- 100% of patients who tested positive received appropriate therapy in <9 hours
- Therapy was discontinued for all patients who tested negative<sup>4</sup>

1. Wilson, N.M., Kenney, R.M., Tibbetts, R.J., et. al. T2 Magnetic Resonance Improves the Timely Management of Candidemia. Poster Presentation IDWeek 2016.

2. Estrada, S. J. Real World Value of T2Candida Lee Memorial Hospital. Slide Presentation ASM 2016.

3. Kateon, H., Edwards, J., Sawyer, A., et al. Utilization of T2Candida Panel for the rapid detection of *Candida* species in a large community hospital. Poster Presentation IDWeek 2016.

4. Patel, F. and Young, E. Antifungal Prescribing During Initial Implementation of Candidemia Early Detection and Species Identification Testing with T2Candida Panel. Poster Presentation IDWeek 2016.

# Significant Burden of Bacterial Infection and Sepsis

Payors should support and incentivize revised protocols

**Add  
T2Bacteria  
&  
T2Candida**

**>90% of  
patients on the  
right targeted  
therapy within  
6 to 8 hours**



Representing \$27B in U.S. healthcare costs<sup>1,2</sup>

**~\$25,000 Cost Savings**

Per patient if on right therapy  
within 24 hours<sup>3</sup>

**Billions of Dollars**

In savings for hospitals, including  
decreased readmissions<sup>4</sup>

**50% Reduction**

In mortality for patients with rapid  
effective treatment<sup>5</sup>

**Patients Benefit**

From reduction in long-term  
side-effects

1. Torio, C. M. and Moore, B. J. (2016). Statistical Brief# 204. Healthcare Cost and Utilization Project (HCUP). May.

2. McDermott, K. W., Elixhauser, and A., and Sun, R. (2017). Statistical Brief# 225. Healthcare Cost and Utilization Project (HCUP). June.

3. Estimated economic impact based on customer experience with T2Candida Panel; Bilir, S. P., Ferrufino, C. P., Pfaller, M. A., and Munakata, J. (2015); and studies for target bacterial species.

4. See slide 11.

5. Leibovici, L., Shraga, I., Drucker, M., et al. (1998). The benefit of appropriate empirical antibiotic treatment in patients with bloodstream infection. Journal of internal medicine, 244(5), 379-386.

# Established Reimbursement Across Multiple Care Environments

Financially attractive in all settings

## Point-of-Care Testing

Emergency Room  
Outpatient Settings

- CPT 87640, 87798
- Coverage if not admitted; other outpatient settings
- ER is most common setting

	T2Bacteria
Reimbursement	\$220
Cost of Test	\$150

## In-Patient Hospital

Admitted from ER  
Admitted for Unrelated Procedure

- DRG 870, 871, 872; example DRG reimbursement: \$35,000<sup>1</sup>
- NTAP max reimbursement of \$97.50 (65% of list price of T2Bacteria), starting 10/1/19

	T2Bacteria	T2Candida
Cost of Test	\$150	\$200
Cost After NTAP	\$52.50	
Percent of DRG	0.15%	0.6%

0.75% of DRG

# The T2Dx Impact

Improve the quality of patient care while reducing healthcare costs

## Targeted Rx

- Reduced resistance
- Reduced length of stay
- Potential reduction in morbidity and mortality



## Efficient use of limited resources

- Reduced repeat testing
- Reduced unnecessary Rx
- Reduced time waiting for diagnostic test results

# Adoption Drives Revenue and Rapid Pay Back

Doing well by doing good

## Typical High Risk Patients In Target Market

Patients Suspected of Sepsis	3,000
Patients Suspected of Fungal Infections	375

## Potential Hospital Utilization Scenario

	Patients Tested	Price per Test	Total Revenue
T2Bacteria	1,500	\$150	\$225,000
T2Candida	375	\$200	\$75,000
<b>Annual Recurring</b>	<b>1,875</b>		<b>\$300,000</b>
<b>T2Dx Instrument</b>		\$100,000 unit price	

In this example, patients suspected of sepsis are screened with the T2Bacteria Panel in the ER and throughout portions of the hospital as part of a sepsis protocol.

# Commercial Strategy

Global expansion of T2Direct Diagnostics driven by T2Bacteria Panel launch

## Expanding on the existing T2Dx installed base



### United States

#### Direct Sales

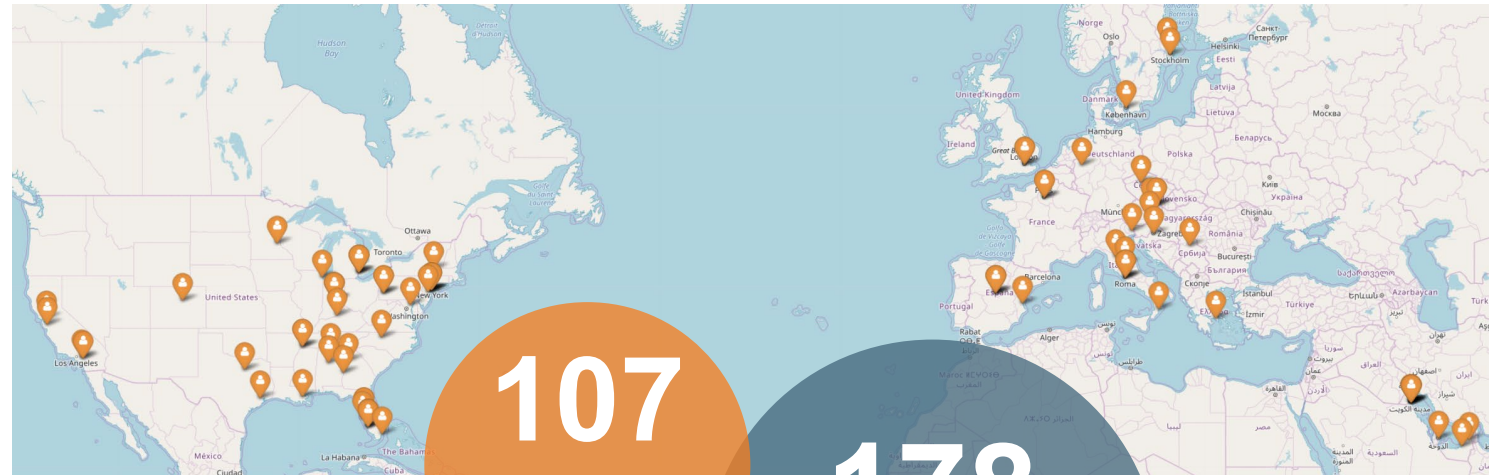
- **Organization:** 15 sales reps and 6 medical affairs liaisons
- **Target:** 1,200 hospitals with the highest concentration of patients at risk for sepsis-related infections



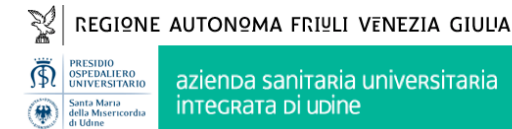
### International

#### Distributor Sales in 35 Countries

- 14 distribution partners supported by small team of direct sales/marketing and field service personnel



# Selected T2 Biosystems Customers



## Global expansion of T2Direct Diagnostics driven by T2Bacteria Panel launch

## Medical Meetings & Conferences



**29th ECCMID** Amsterdam, Netherlands  
13 – 16 April 2019

# Digital Marketing



**T2 Biosystems**  
4,779 followers  
6d

WEBINAR: Sepsis Management: The Emergence of Rapid Diagnostics in the ED.\* Sepsis, the #1 cause of death in US hospitals, has a mortality that exceeds prostate cancer, breast cancer and AIDS combined. As many as 92% of s ...see more



LIVE LECTURE

Sepsis Management:  
The Emergence of Rapid Diagnostic  
in the Emergency Department

SPEAKER: W. Frank Peacock IV, MD, FACP, FCCP  
Professor of Emergency Medicine, Associate Chief, and  
Research Director for the Department of Emergency Medicine  
at the Baylor College of Medicine

WEBINAR

SEPT. 18, 2018 11:00 AM ET


**T2Biosystems**

37 Likes · 3 Comments

### Targeted Social Media and Email Campaigns



invited guest speaker Prof. Mustafa Bangash, MD of Denver Hospital, [year presented at the CMAA conference 2013](#) has T2Bacteremia can help critical care and emergency medicine clinicians to:

- Rapidly diagnose and treat some of the most common and deadly pathogenic organisms.
- Demonstrate how rapid diagnostics can be a "game changer" in the microbiology lab.

The delay of microbial culture results often leave them unhelpful for the clinician, but T2Bacteremia achieves the "need for speed" in the clinical setting. Prof. Bangash presented CMAA Researcher T2Bacteremia data recently published in the [Journal of Antimicrobial Pharmacology](#), demonstrating:



Please contact us to request a meeting on T28est@esig.

Best Regards,  
Karl M. Gonyea  
Senior Director Marketing, Repco  
72 Bixbylane, Inc.

[Download Prof. Sanguinetti's Presentation](#)

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[Schedule a Meeting on T2Bacteria](#)

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www.elsevier.com/locate/jbiotec

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## Publications / Economic Models

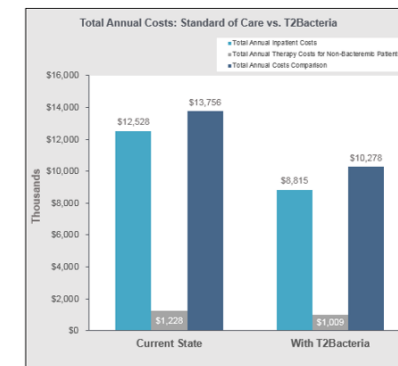
## Utilizing Same-Day Sepsis Pathogen ID for Faster Therapy, Enhanced Stewardship, Improved Outcomes

Cornelius J. (Neil) Clancy, M.D.

20 June 2018



## T2Direct Diagnostics™ Economic Model

[illegible]

# T2Resistance Panel

The first direct-from-blood detection of resistance markers

- Detection of **13 resistance genes** from both Gram-positive and Gram-negative pathogens from a single patient blood sample, without the wait for blood culture, in 3-5 hours
- Covers the most clinically important genes, including several listed on the **CDC's Urgent Threat list** for antibiotic resistance
- Utilizes **same T2Dx Instrument** as the T2Bacteria and T2Candida Panels
- Developed with the help of an award from CARB-X (funded by BARDA), the Wellcome Trust, and the National Institute of Allergy and Infectious Diseases (NIAID)
- Expected to be available for research use only in Q3 2019 and to receive CE Mark for commercial availability outside U.S. by the end of 2019

## FDA Breakthrough Designation

- Granted “Breakthrough Device” designation by the FDA
- Allows T2 Biosystems to work closely with the FDA during the premarket review phase to ensure patients can have access to the benefits of this innovation as soon as possible

Powered by **CARB-X**



# T2 Product & Pipeline Highlights

Enabled by Highly Sensitive Detection

Powered by **CARB-X**

## T2Candida®

Sensitivity: 91.1%<sup>2</sup>  
Specificity: 99.4%<sup>2</sup>

*C. albicans*  
*C. tropicalis*  
*C. parapsilosis*  
*C. krusei*  
*C. glabrata*

New FDA Product Code  
1-3 CFU/mL LoD

## T2Bacteria®

Sensitivity: 95.4%<sup>1</sup>  
Specificity: 98.0%<sup>1</sup>

*E. faecium*  
*S. aureus*  
*K. pneumoniae*  
*P. aeruginosa*  
*E. coli*

New FDA Product Code  
2-11 CFU/mL LoD

## T2Resistance

FDA Breakthrough Device  
CE Mark/RUO 2019

*mecA/C*  
*vanA/B*  
CTXM-14/15  
KPC  
OXA-48 Group  
NDM, VIM, IMP  
AmpC (CMY/DHA)

In Development  
2-5 CFU/mL LoD

## T2Candida auris

Sensitivity: ≥89%  
Specificity: 98%

*C. auris*  
*C. duobushaemulonii*  
*C. haemulonii*

RUO; Validated by CDC for  
patient swabs, demonstrated  
performance in blood;  
≤5 CFU/mL LoD

## T2Lyme

PPA: 78%  
NPA: 100%

*B. burgdorferi*  
*B. afzelii*  
*B. garinii*  
*Borellia spp.*

In pivotal study;  
3-10 cells/mL LoD

1. T2Bacteria Pivotal Clinical Study. This is a combination of samples run in both prospective and contrived arms of study. T2Bacteria showed an overall average sensitivity of 90% in the prospective arm of the study and the contrived arm an overall average PPA of 97%.

2. Mylonakis, E., Clancy, C.J., Ostrosky-Zeichner, L., et al. (2015). Clinical Infectious Diseases

3. Research reported in this presentation is supported by the Cooperative Agreement Number IDSEP160030 from ASPR/BARDA and by an award from Wellcome Trust, as administrated by CARB-X. The content is solely the responsibility of T2 Biosystems and does not necessarily represent the official views of the Department of Health and Human Services Office of the Assistant Secretary for Preparedness and Response, other funders, or CARB-X.

# Financial Summary<sup>1</sup>

June 30, 2019		
Revenue	2Q19	\$1.8M
	2Q18	\$3.9M
	FY18	\$10.5M
Product Revenue	2Q19	\$1.3M
	2Q18	\$1.2M
	FY18	\$4.8M
Product Growth	YoY	8%
Cash Burn	2Q19	\$8.9M
Cash <sup>4</sup>		\$28.6M
Common Shares Outstanding	2Q19	44.4M

>5% Investors – As of June 30, 2018 <sup>2,3</sup>	
Canon Life Sciences	13.6%
Goldman Sachs	9.6%
Senvest Management	6.4%

1. All amounts are rounded to the nearest hundred thousand.
2. Based on 44,535,572 shares outstanding as of June 30, 2019.
3. Source SEC filings as of July 25, 2019.
4. Includes \$180k restricted cash.

# Guidance

2019 Guidance	
<b>2019 total revenue</b>	<b>\$8.7 - \$9.6 million</b>
Product revenue	\$5.7 - \$6.1 million
<b>3Q 2019:</b>	
Product revenue	\$1.4 - \$1.5 million
<b>2019 T2Dx new contracts:</b>	<b>43 – 53</b>
<b>3Q/4Q 2019 T2Dx new contracts:</b>	<b>10 – 15</b>
<b>Quarterly operating expense:<sup>1</sup></b>	<b>\$10.5 - \$11.5 million<sup>2</sup></b>

Breakeven Model	
Total revenue	\$65 - \$75 million
Gross margin	~45 - 50%

1. Excluding cost of product revenue.

2. Including non-cash depreciation and stock based compensation of approximately \$3.0 million; contingent on closing a research collaboration

3. \* This slide contains T2's future goals and aspirations, which constitute forward-looking statements that are subject to risks and uncertainties that could cause actual results to differ materially from those expressed or implied by such statements. See "Forward-Looking Statements" on slide 2.

# Breakthroughs in Medical Diagnostics

First and only FDA-cleared diagnostic to detect pathogens directly from blood

FAST COMPANY



Early 20<sup>th</sup>  
century



Blood  
Culture

1947



Cell  
Counting

1959



Immunoassay

1968



Automated  
Chemistry  
Analyzers

1985



PCR  
Technique

## Direct Sample Analysis Identify Pathogens Using Magnetic Resonance



# Investment Highlights

A platform technology with multiple, billion-dollar franchise opportunities



## T2MR

Innovative  
technology - broad  
applications



## Market

\$2B+ Initial market  
potential



## Sepsis Causing Pathogen ID

Provide species-specific  
results, direct from whole  
blood, in 3 to 5 hours



## Reimbursement

Covered by existing  
reimbursement codes



## Robust Pipeline

A new generation of  
diagnostics



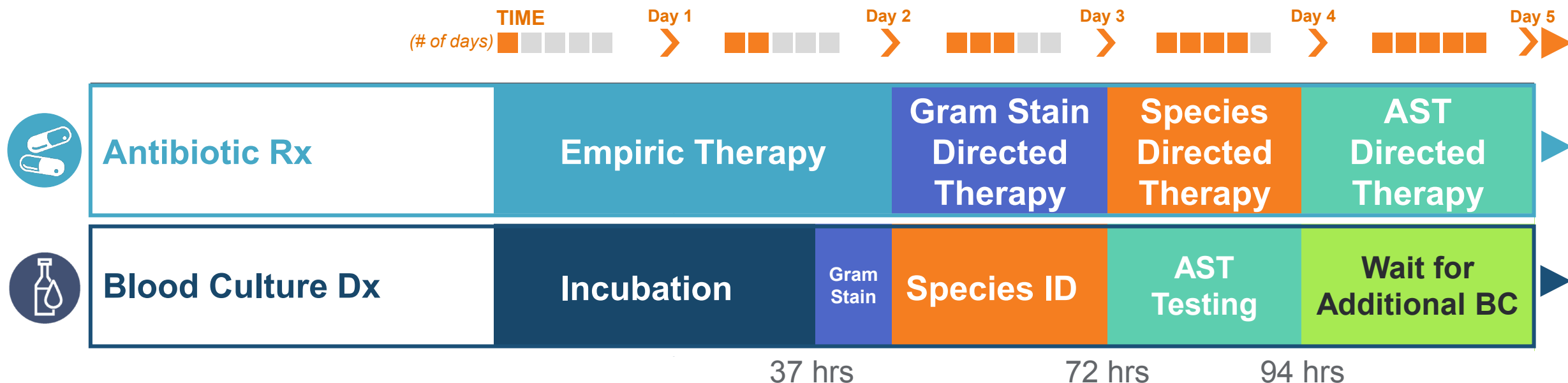
## Execution

Patient access growing,  
key collaborations  
established

# Breakthrough Sepsis Pathogen Detection

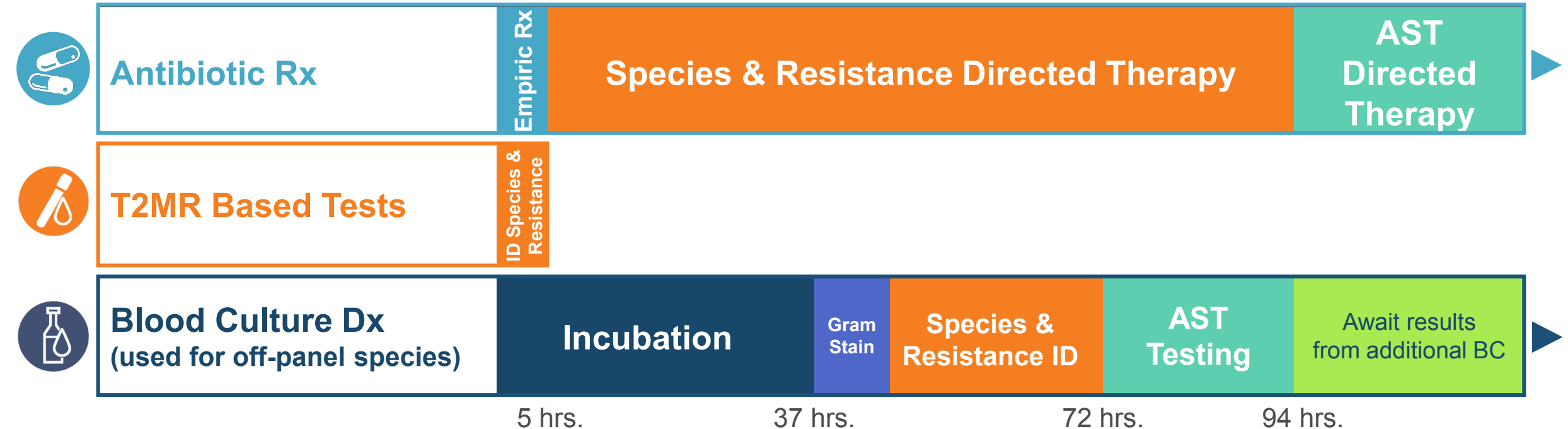
## Appendix

# Diagnostics Time to Result Influences Therapy



- Antibiotic administration rates range from 50% to 70% for patients with a blood culture draw <sup>(1-3)</sup>
- Only ~10% of patients suspected of sepsis yield a positive blood culture.
- Meta-analysis of 70 studies found empiric antibiotic therapy was inappropriate in 46.5% of patients<sup>4</sup>
- The proportion of patients on effective therapy after organism species ID has been shown to be >90%, demonstrating effectiveness of antibiogram-directed therapy based on species ID<sup>5</sup>
- Significant percentage of patients infected with resistant organisms die before AST results are available<sup>6</sup>

# T2MR & Blood Culture Results Influence Therapy



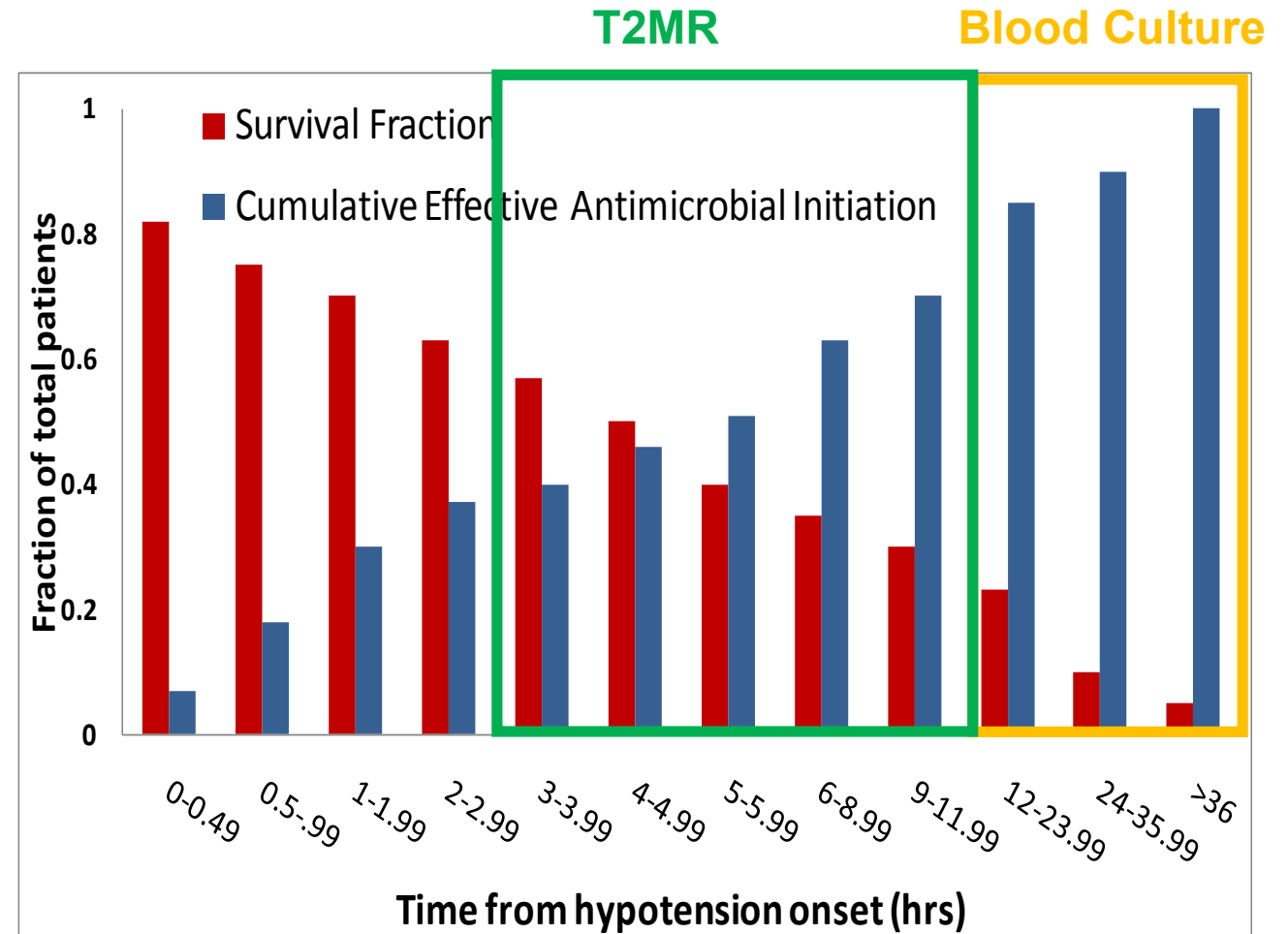
- T2MR Based tests enable more rapid targeted therapy based on species ID & resistance
- Across 3 studies, 94%-100% of patients are correctly treated after species ID, not after assessment of susceptibility testing, demonstrating effectiveness of antibiogram-directed therapy<sup>1-3</sup>
- Numerous studies indicate that this will reduce both LoS and mortality for infected patients

(1) Doern et al. *J Clin Micro* 1994; (2) Byl et al. *Clin Infect Dis* 1999; (3) Kerremans et al. *J Clin Microbiol* 2012  
CORPORATE PRESENTATION

Average Turn around times from Nguyen et al. *Annals Int Med* 2019

# Time to Appropriate Therapy Impacts Survival










- Key predictor of survival and length of stay (LoS) for patients with bacteremia is time to effective therapy
- As many as 80% of sepsis deaths could be prevented with rapid diagnosis and treatment
- For every hour delay in time to appropriate therapy:
  - Survival decreases by 7.6% during septic shock<sup>1</sup>
  - Relative odds of death increase by 4.0% during bacteremia<sup>2</sup>
- Reducing time to effective therapy has resulted in significant reductions in LoS, up to 8 days<sup>3-5</sup>
- Appropriate and rapid delivery of targeted antibiotics is critical for surviving sepsis<sup>6</sup>



Kumar A. et al., Crit Care Med 2006, 34:1286, N=2731










# “Pursuit of the Holy Grail”\*

## Rapid, Direct-from-Blood Pathogen & Resistance Results

	Development	CE-Marked	FDA Cleared	US Clinical Use
				
				
				
				
				
				
				
				
				

1. For decades, numerous companies & technologies have pursued the development of a platform for direct-from-blood pathogen & resistance results.
2. Several have stopped or shifted their development programs to post positive blood culture as a “direct” sample input.

# Progress towards Culture Independent Tests

	LoD (CFU/mL)*	Clinical Sensitivity	Clinical Specificity	Comments
	1-11	90% - per sample 90% - per result	90% - per sample 98% - per result	CE mark & FDA cleared <sup>1</sup>
	3-100	65%	86%	Manual test, CE mark only <sup>2</sup>
	1-100	44%	87%	Automated test, CE-mark only <sup>3</sup>
	10-50	85%	48%	Manual test, CE mark only <sup>4</sup>
	8-32	86% - per sample	86% - per sample	Withdrew FDA application, not on market <sup>5</sup>
	1,000,000	N/A	N/A	Focused on post-culture <sup>6</sup>
	1,000	N/A	N/A	Maintained focus on post-culture <sup>7</sup>
	N/A	N/A	N/A	Formal LoD data not disclosed <sup>8</sup>
	N/A – cfDNA	94% - per sample	40% - per sample	Send-out reference lab test <sup>9</sup>

1. Nguyen et al. Annals Int Med 2019; 2. Roche Septifast labeling; Stevenson et al Health Technol Assess 2016; 3. McKeating JCP 2018; 4. Molzyme Sepsitest product labeling; 5. Metzgar et al PLOS ONE 2016; 6. PhenoTest 510(k) summary; 7. BCID product labelling; 8. Qvella AMP 2017 presentation; 9. Blauwkamp et al. Nature Micro 2019.  
\*LoD based on ≥95% positivity rate for N≥20 replicates at a given titer level using CFU/mL assignment of cell solution spiked into blood.

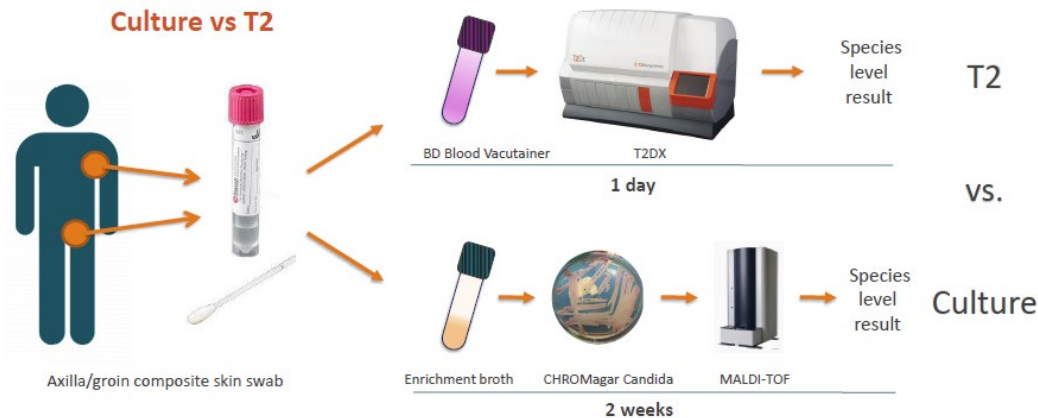
# Rapid Development of *C.auris* T2MR Panel

- ID of *Candida auris* has been hampered by the poor specificity, poor sensitivity and slow growth of organism
- Resulting in a delays in therapy for infected patients and delays in infection controls for colonized patients accelerating the spread of *Candida auris*



## *C. auris* diagnostic comparison:

### Culture vs T2



- CDC approached T2 Biosystems with problem of prolonged culture for *C. auris*
- T2Candida auris Panel RUO (5 hr TAT) vs. Patient swab cultures (14 days TAT)
- T2Dx's installed at CDC Mycology laboratory for validation of use on patient skin swab samples<sup>1-3</sup>
- 100% inclusivity of Clades I-IV,  $\leq 5$  CFU/mL LoD;  $\geq 89\%$  Sensitivity & 98% Specificity; 98% PPV, 89% NPV

1. Manning et al. "Automated Detection of Candida auris Direct from Whole Blood and Swab Specimens by T2MR" ID Week 2017  
 2. Sexton et al "Evaluation of a new T2 Magnetic Resonance assay for rapid detection of emergent fungal pathogen Candida auris on clinical skin swab samples." Mycoses 2018  
 3. Sexton et al. "Evaluation of T2 Magnetic Resonance Candida auris Panel as a Rapid Diagnostic for this Emerging Multidrug Resistant Yeast in Clinical Skin-Swab Samples." ASM 2018.

# T2Lyme Clinical Performance

N= 21 Subjects	Tissue Culture		
Diagnostic Test	PPA	NPA	OPA
T2Lyme (blood)	78%	100%	90%
Tissue PCR	67%	100%	86%
Seroconversion	67%	67%	67%
2-Tier (Serology/WB)	56%	92%	76%
Blood PCR	0%	100%	57%
T2Lyme and 2-tier IgM	100%	92%	95%

- True positive subjects were established by culturing *Borrelia* from an EM tissue sample.
- T2Lyme PPA was 78%, while CDC recommended 2-tier diagnosis PPA was 56%
- T2Lyme had higher accuracy (OPA) than all other methods; combination of T2Lyme & 2-tier IgM provided best overall clinical accuracy
- Blood PCR (PPA, 0%) failed to return any positive results
- NPA of T2Lyme was 100%, indicating greater specificity over serology results
  - We've tested >550 negative samples with no false-positive results

# Comprehensive T2MR Panel for Bloodstream Infections

Result type	N	Results
Pan-level	3	Gram positive bacteria, Gram negative bacteria, Fungi
Genus level	11	<i>Acinetobacter spp.</i> , <i>Candida spp.</i> , <i>Citrobacter spp.</i> , <i>Enterobacter spp.</i> , <i>Enterobacteraceae</i> , <i>Enterococcus spp.</i> , <i>Listeria spp.</i> , <i>Mycobacterium spp.</i> , <i>Staphylococcus spp.</i> , <i>Coag negative Staphylococcus spp.</i> , <i>Streptococcus spp.</i>
Gram pos. species	6	<i>E. faecium</i> , <i>E. faecalis</i> , <i>S. aureus</i> , <i>S. pneumoniae</i> , <i>S. pyogenes</i> , <i>S. viridans</i>
Gram neg. species	6	<i>A. baumannii</i> , <i>E. coli</i> , <i>Enterobacter cloacae</i> complex, <i>H. influenzae</i> , <i>K. pneumoniae</i> , <i>P. aeruginosa</i>
Fungal Species	7	<i>C. albicans</i> , <i>C. tropicalis</i> , <i>C. dublinensis</i> , <i>C. parapsilosis/C. metapsilosis/C. orthopsilosis</i> , <i>C. krusei</i> , <i>C. glabrata</i> , <i>C. auris</i>
Resistance genes	13	<i>mecA/C</i> , <i>mefA/E</i> , <i>vanA/B</i> , <i>ermA/B</i> , KPC, NDM, VIM, IMP, OXA, SHV, AmpC (CMY, DHA), CTX-M 14/15, TEM

- T2MR Feasibility data in hand to support T2MR panel of ≥40 reported results
- Achieves inclusivity of ≥ 99% of infections caused by blood-borne bacterial and fungal pathogens
- “Pan-level” channels detect >250 pathogen species; ≤10 CFU/mL detection demonstrated for pan gram+/-
- Resistance markers provide coverage for all blood-borne CDC antibiotic resistance threats
- New T2MR detection technology developed to shorten turn-around time and reduce cost
- T2MR is proven for direct-from-whole-blood; new data shows feasibility for positive culture detection