

30 års erfaring med bakteriæmier

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Aalborg Universitet

1. møde i Arbejdsgruppen for bakteriæmi under DSKM

The interrelationship
between systemic
inflammation
response syndrome
(SIRS), sepsis, and
infection.

Kilde: Chest 1992;
101: 1644-55.

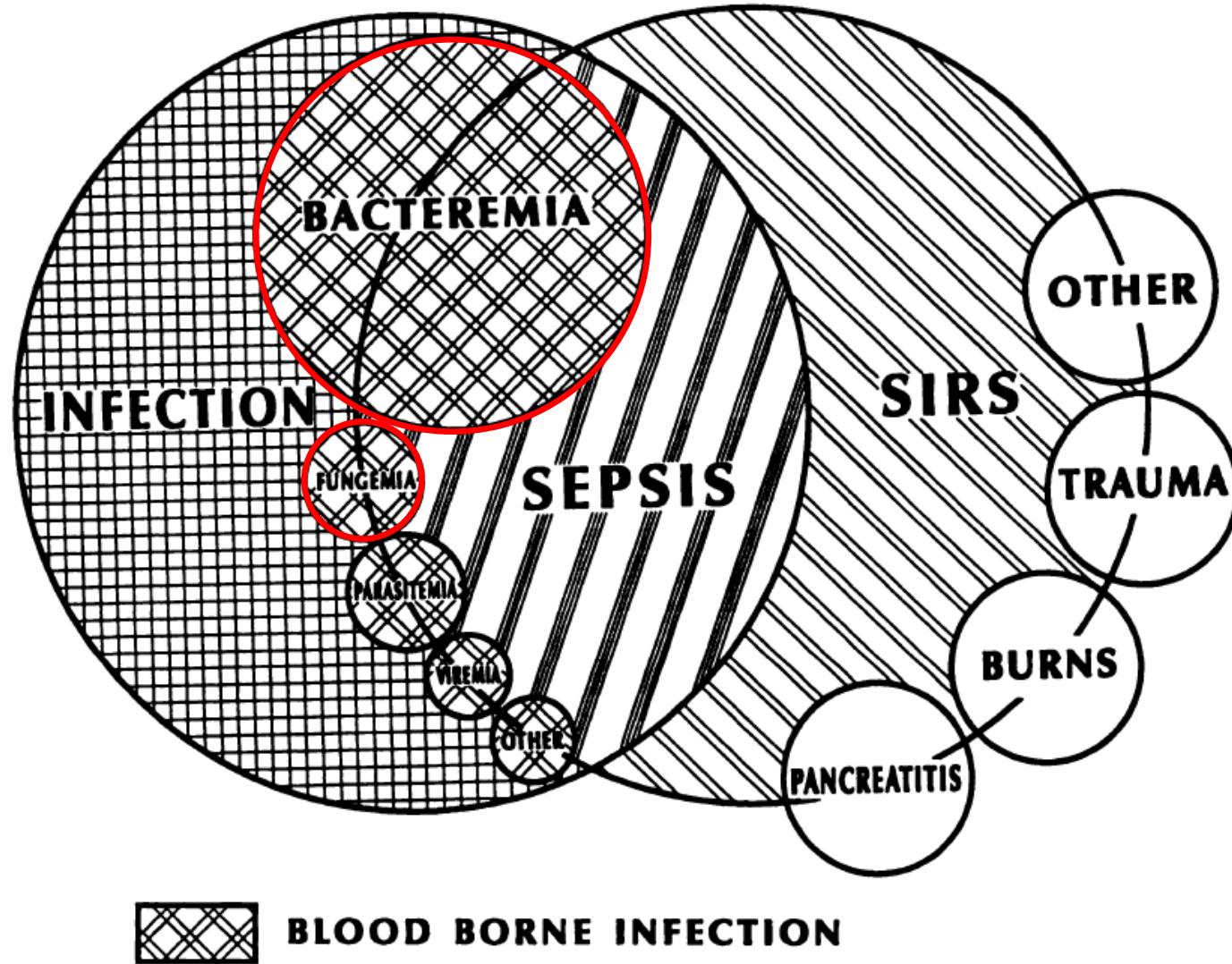
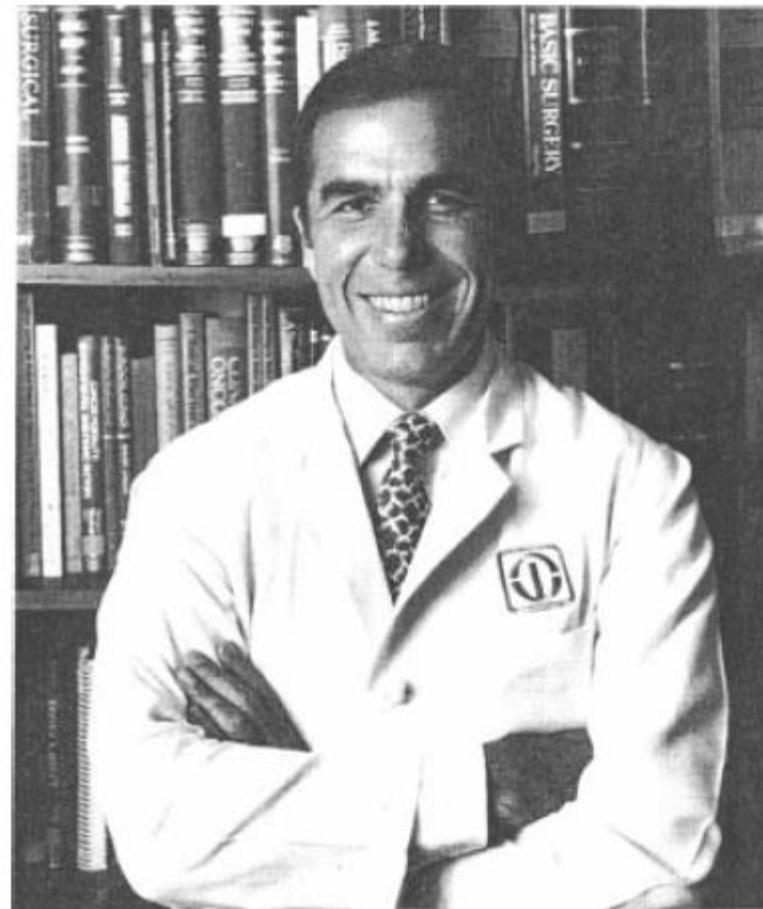


FIGURE 1. The interrelationship between systemic inflammatory response syndrome (SIRS), sepsis, and infection.

1. møde i Arbejdsgruppen for bakteriæmi under DSKM

IN MEMORIAM



Roger C. Bone MD

Død 1997

Biografi 1980'erne

Denne afhandling er i forbindelse med de nedenfor anførte, tidligere offentligjorte afhandlinger af det lægevidenskabelige fakultet ved Aarhus Universitet antaget til offentligt at forsvares for den medicinske doktorgrad.

Århus, den 10.5.1987

Palle Juul-Jensen, dekan

This thesis is based on the following papers. In the text these papers are referred to by capital letters.

- A. Schønheyder H, Andersen P. An indirect immunofluorescence study of antibodies to *Aspergillus fumigatus* in sera from children and adults without aspergillosis. *Sabouraudia* 1982; 20: 41–50.
- B. Schønheyder H, Andersen P, Stenderup A. Serum antibodies to *Aspergillus fumigatus* in patients with pulmonary aspergillosis detected by immunofluorescence. *Acta Pathol Microbiol Immunol Scand Sect B* 1982; 90: 273–279.
- C. Schønheyder H, Andersen P. Complement-binding antibodies to *Aspergillus fumigatus* in patients with pulmonary aspergillosis. *Acta Pathol Microbiol Immunol Scand Sect B* 1983; 91: 1–7.
- D. Schønheyder H, Andersen P, Pedersen AT. Antibodies against *Aspergillus fumigatus* in bronchial secretions from patients without aspergillosis. *Acta Pathol Microbiol Immunol Scand Sect B* 1983; 91: 187–191.
- E. Schønheyder H, Andersen P. Determination of antibodies to partially purified aspergillus antigens by an enzyme-linked immunosorbent assay. *Int Arch Allergy Appl Immunol* 1983; 70: 108–111.
- F. Schønheyder H, Andersen P. Fractionation of *Aspergillus fumigatus* antigens by hydrophobic interaction chromatography and gel filtration. *Int Arch Allergy Appl Immunol* 1984; 73: 231–236.
- G. Schønheyder H, Andersen P. IgG antibodies to purified *Aspergillus fumigatus* antigens determined by enzyme-linked immunosorbent assay. *Int Arch Allergy Appl Immunol* 1984; 74: 262–269.
- H. Schønheyder H, Andersen P. Serum IgA antibodies to *Aspergillus fumigatus* determined by enzyme-linked immunosorbent assay. *Acta Pathol Microbiol Immunol Scand Sect B* 1984; 92: 279–282.
- I. Schønheyder H, Jensen T, Høiby N, Andersen P, Koch C. Frequency of *Aspergillus fumigatus* isolates and antibodies to aspergillus antigens in cystic fibrosis. *Acta Pathol Microbiol Immunol Scand Sect B* 1985; 93: 105–112.
- J. Schønheyder H, Møller-Hansen C, Andersen P, Stenderup A. Serum antibodies to *Aspergillus fumigatus* in Danish farmers. *Sabouraudia J Med Vet Mycol* 1985; 23: 93–99.
- K. Schønheyder H, Andersen P, Munck Petersen JC. Rapid immuno-electrophoretic assay for detection of serum antibodies to *Aspergillus fumigatus* catalase in patients with pulmonary aspergillosis. *Eur J Clin Microbiol* 1985; 4: 299–303.
- L. Schønheyder H, Storgaard L, Andersen P. Variation of a 470 000 daltons antigen complex and catalase antigen in clinical isolates of *Aspergillus fumigatus*. *Sabouraudia J Med Vet Mycol* 1985; 23: 339–349.



1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Pathogenetic and Serological Aspects of Pulmonary Aspergillosis

By

Henrik Schønheyder

Institute of Medical Microbiology, University of Aarhus, DK-8000 Aarhus C, Denmark

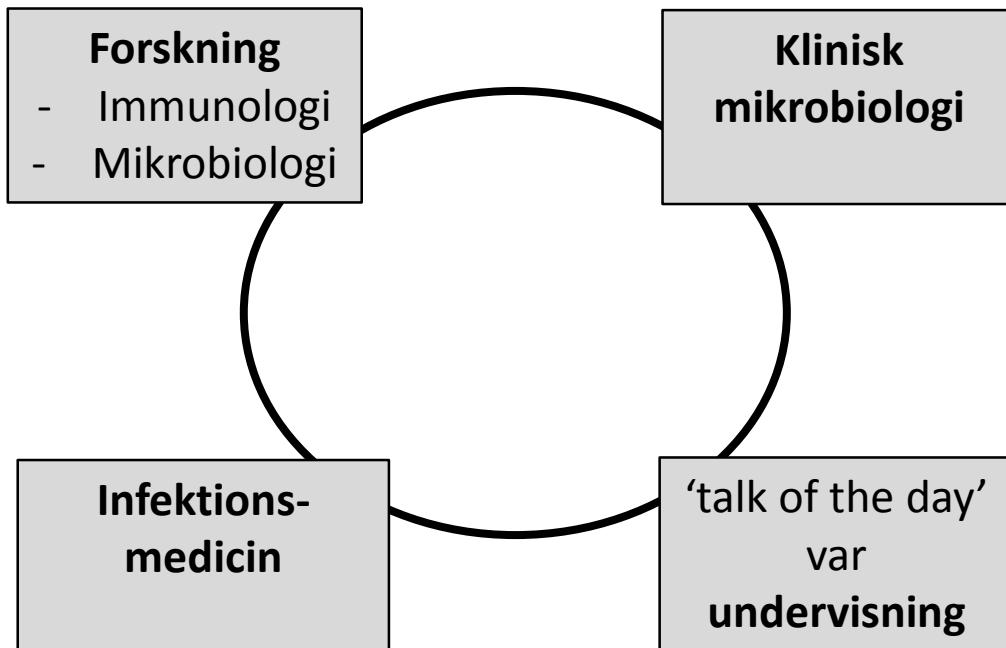
Scand J Infect Dis
Suppl. 51 1987

St. Mary's Hospital – Wright-Fleming Institute



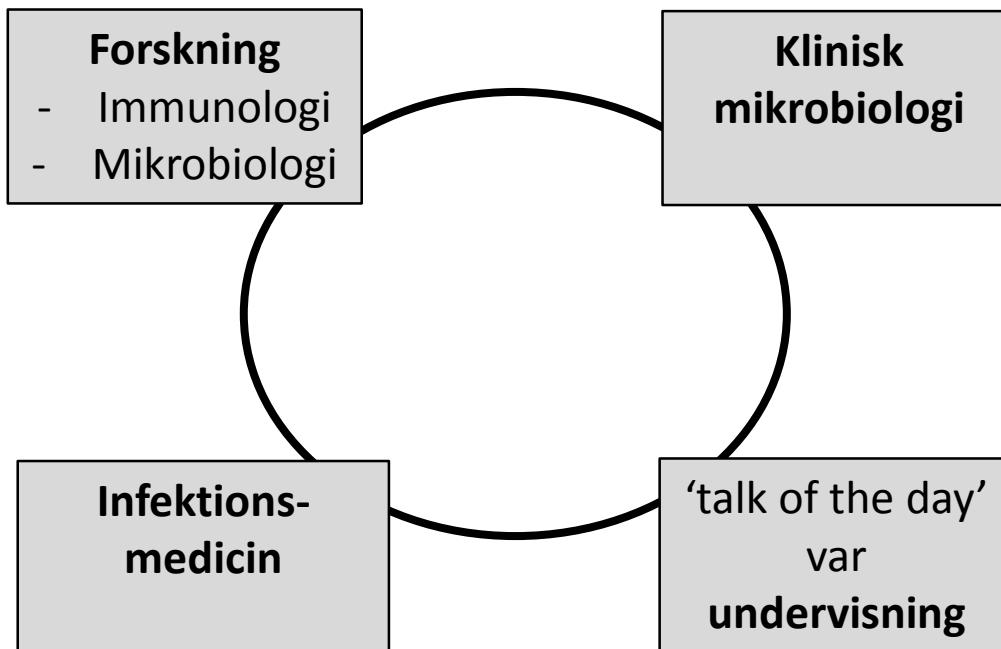
1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Karriereskifte



1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Karriereskifte



1987

Skifte af karrierespor

Uddannelse i klinisk mikrobiologi

Aalborg : Klinisk Mikrobiologisk Afdeling

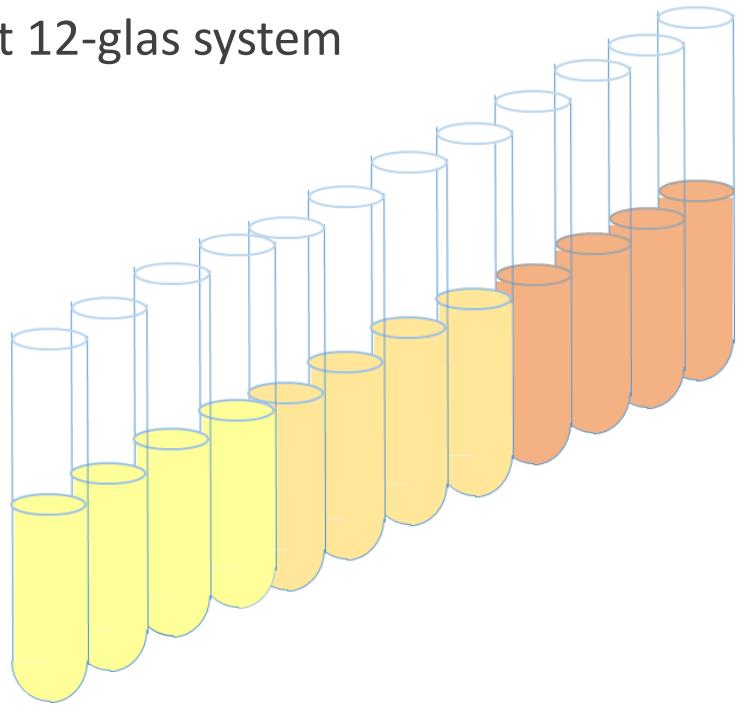
Indtil midt-80'erne regional afdeling af SSI

Klinisk og farmakologisk orienteret

"Evidence-based medicine før det blev moderne"

Hvorfra den særlige interesse for bakteriæmier?

Som yngre læge udsåede man venüler i vagten med et 12-glas system



Kjeld Truberg Jensen & Olaf Scheel

- Bacteremia in the County of North Jutland.
- Antibiotic resistance of gram-negative bacteria from blood cultures
- Resistance to antibiotics in gram-positive bacteria from blood cultures

Artiklerne er publiceret i Ugeskr. Læger I 1988

De første artikler relateret til bakteriæmi

Ebbesen F, Schønheyder HC

Tidligt indsættende neonatal sepsis med ikke-kapsulate *Haemophilus influenzae* - et nyt problem i Danmark. Ugeskrift for Læger **1991**; 153: 1271-1273

Schønheyder HC, Ebbesen F, Grunnet N, Ejlertsen T.

Non-capsulated *Haemophilus influenzae* in the genital flora of pregnant and post-puerperal women. Scandinavian Journal of Infectious Diseases **1991**; 23: 183-187

Prognose for bakteriæmi, de første trin

Schønheyder H, Gottschau A, Friland A, Rosdahl VT.

Mortality rate and magnitude of *Staphylococcus aureus* bacteraemia as assessed by a semiquantitative blood culture system. Scand. J. Infect. Dis. **1995**; 27: 19-21

Grad af bakteriæmi målt ved proportion af positive glas	Antal bakteriæmi episoder	Case fatality (%)
0,30	62	18
0,30 – 0,69	49	23
0,70 – 0,99	59	30
1,00	83	37

Chi2 for trend

DF = 1

P = 0,0095

Prognose for bakteriæmi, det første trin

Schønheyder HC, Højbjerg T

The impact of the first notification of positive blood cultures on antibiotic therapy. A one-year survey. *APMIS* **1995; 103: 37-44**

Ændring af behandling ved 1. kontakt	Total	Behandling Dækende	Behandling Ikke optimal	Ingen behandling
Antal bakteriæmier	694	418	90	186
Ingen ændring	379	299	32	48
Start eller justering	315	199	58	138
+ Aminoglykosid	109	25	21	63
+ <i>S. aureus</i> dækning	73	25	22	24

1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Søgning efter danske studier i PubMed

1. Søgeperiode fra PubMeds start i 60'erne til ultimo 2017 (>50 år)
2. MeSH terms og fritekst dækkende bakteriæmi, candidæmi, "bloodstream infection" (mulighed for flere stavemåder), og "blood culture".
3. Termer for bakterier og svampe koblet med blod eller bloddyrkning.
4. Søgning på "Denmark" i alle felter for at identificere danske studier.
5. Håndplukkede referencer fra 60'erne og enkelte senere år.

1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Søgeresultater

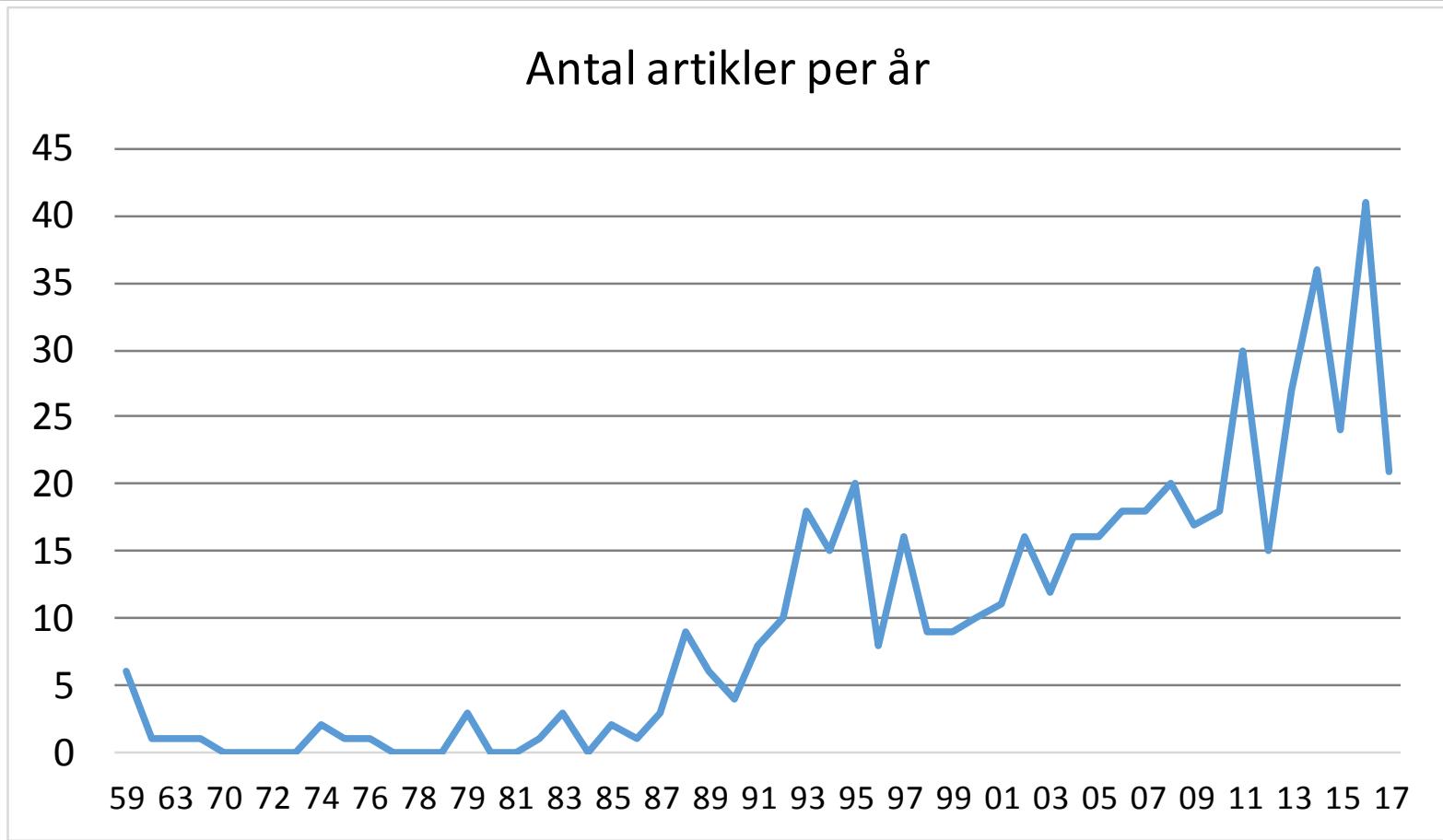
	Emne	Antal
Primært søgeresultat	Bakterier	502
- do -	Svampe	36
Supplerende udtræk	S. pneumoniae m.fl.	30
Supplerende udtræk	Tidlige S. aureus studier	11
I alt		579

Nettoresultat

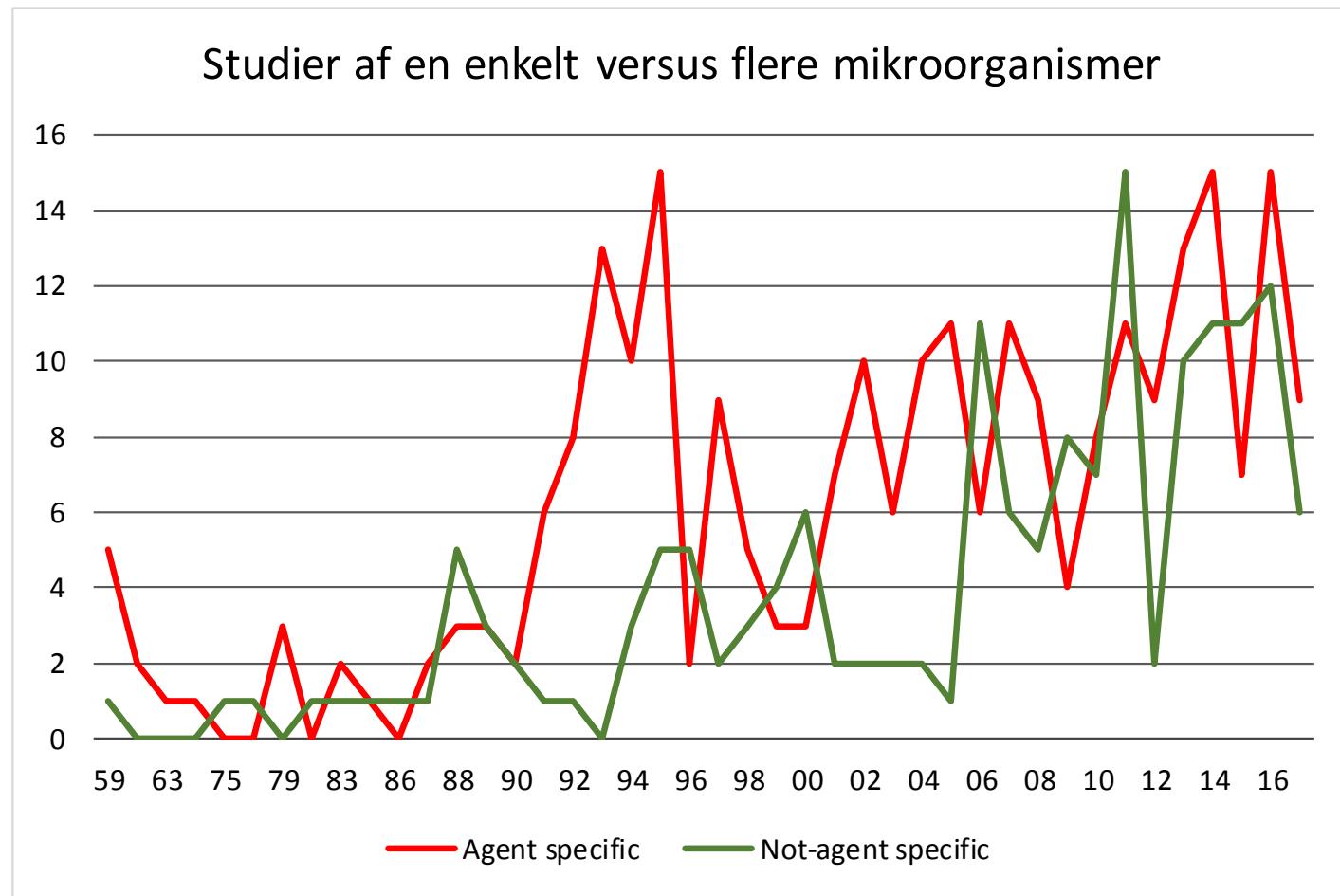
Renset for "Denmark Str.", reference stammer o. lign.	530
"Not eligible"	25
"Not known yet"	2
"Inflammation\sepsis"	46
"Animal models"	6
I alt	451

1. møde i Arbejdsgruppen for bakteriæmi under DSKM

Artikler



Emnevalg



Studie typer *

Cohorter	121
Case serier	38
Kasuistikker	32

* Baseret på titel eller/og abstract

DEBATE

Open Access



CrossMark

Clarifying the distinction between case series and cohort studies in systematic reviews of comparative studies: potential impact on body of evidence and workload

Tim Mathes^{*} and Dawid Pieper

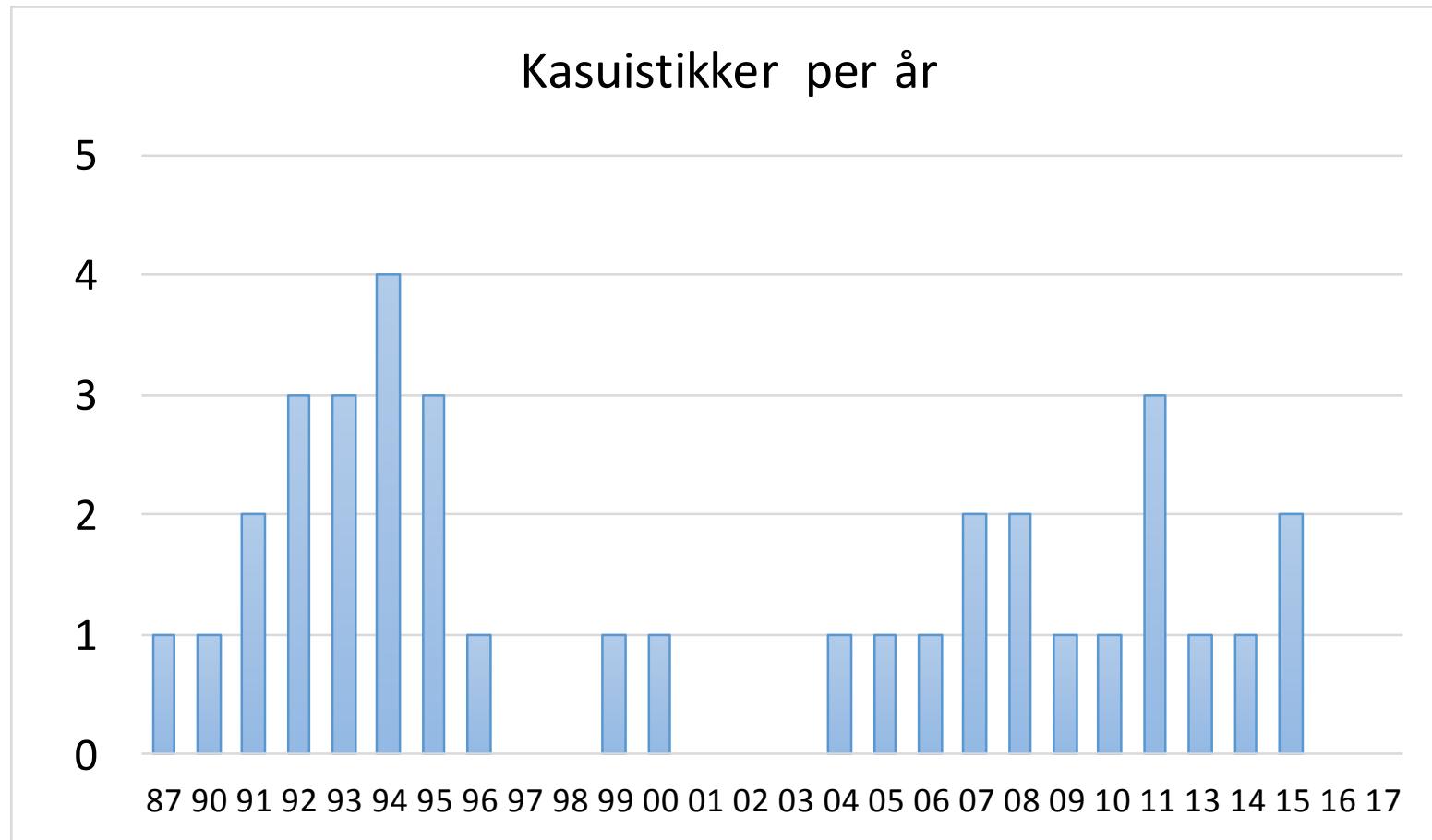
De mest studerede agens (alle typer studier)

Agens	
Staphylococcus aureus	75
Streptococcus pneumoniae	25
Anaerobe bakterier	18
Escherichia coli	17
Hæmolytiske streptokokker	16
Zoonotiske Salmonella	13
Listeria monocytogenes	8
Aerococcus	5
Capnocytophaga/Helicobacter	4

NB Forbehold for
mindre fejl i
optællingen.

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Kasuistikker

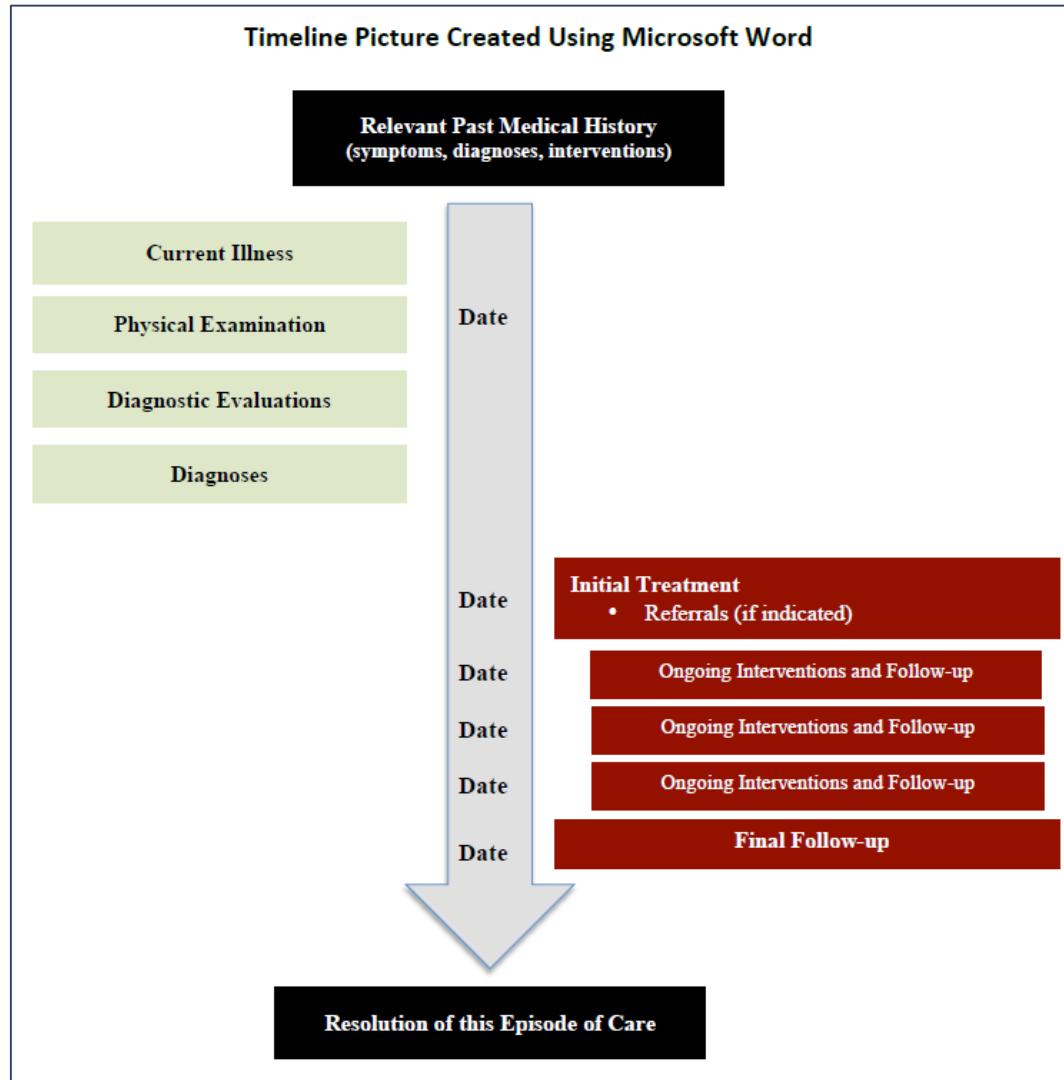




BMJ CASE REPORTS

The CARE guidelines: consensus-based clinical case reporting guideline development

Joel J Gagnier,^{1,2} Gunver Kienle,³ Douglas G Altman,⁴ David Moher,⁵ Harold Sox,⁶ David Riley,⁷ the CARE Group



1. møde i Arbejdsgruppen for bakteriæmi under DSKM

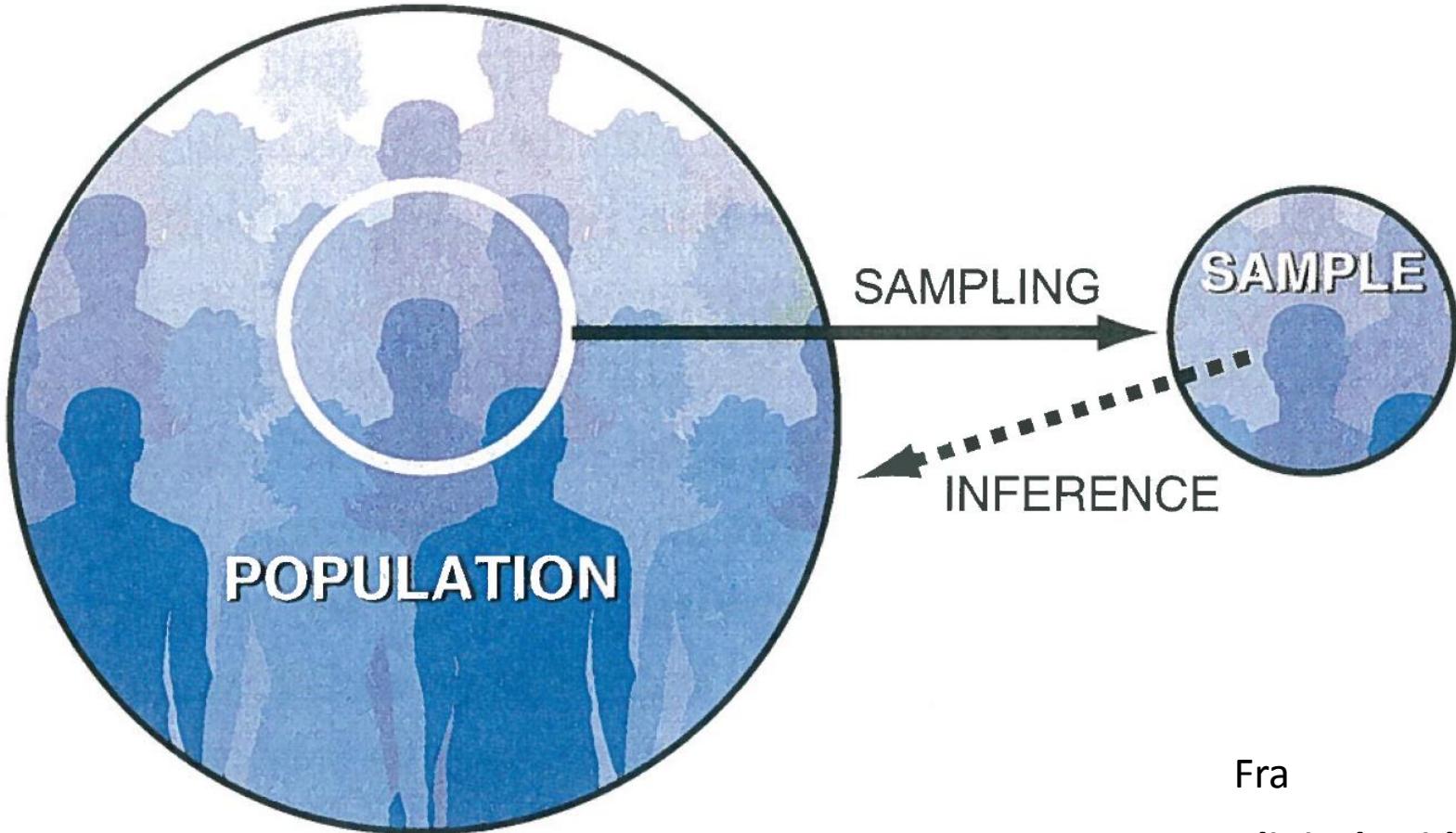


Figure 1.2 ■ Population and sample.

Fra

Clinical Epidemiology, the essentials

Fletcher, Fletcher & Fletcher

Seneste udgave 2014

Clin Infect Dis 2011

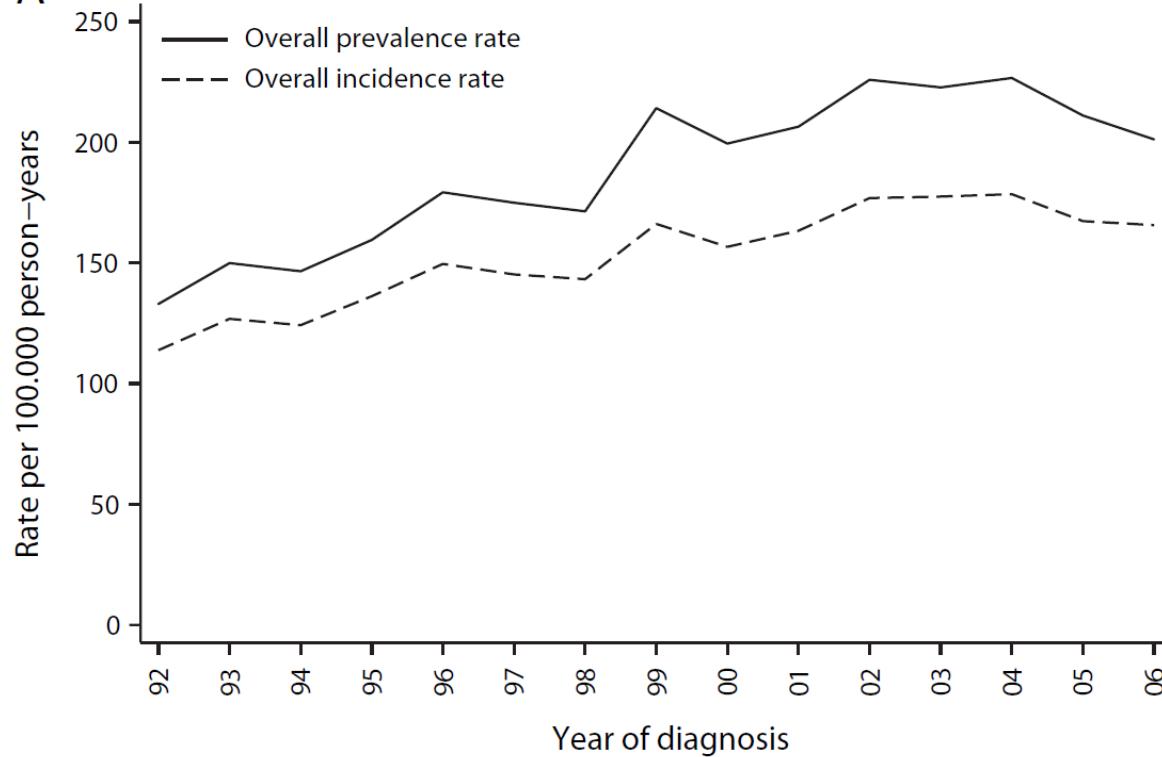
Temporal Changes in the Incidence and 30-Day Mortality associated with Bacteremia in Hospitalized Patients from 1992 through 2006: A Population-based Cohort Study

Mette Søgaard,^{1,2} Mette Nørgaard,² Claus Dethlefsen,³ and Henrik Carl Schønheyder¹

¹Department of Clinical Microbiology, Aarhus University Hospital, Aalborg, ²Department of Clinical Epidemiology, Clinical Institute, Aarhus University Hospital, Aarhus, Denmark, and ³Department of Cardiology, Center for Cardiovascular Research, Aarhus University Hospital, Aalborg

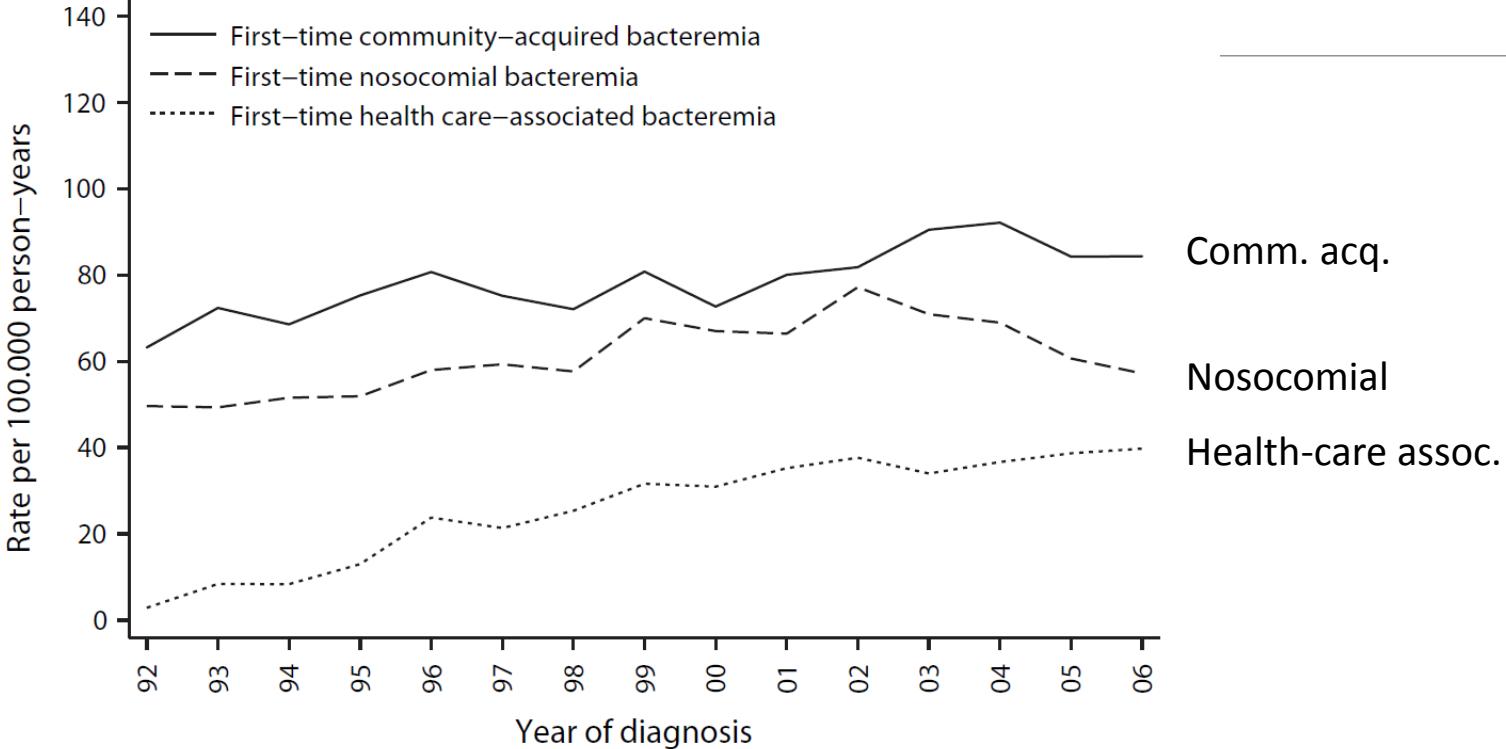
Incidence and prevalence rates – North Jutland 1992-2006

A

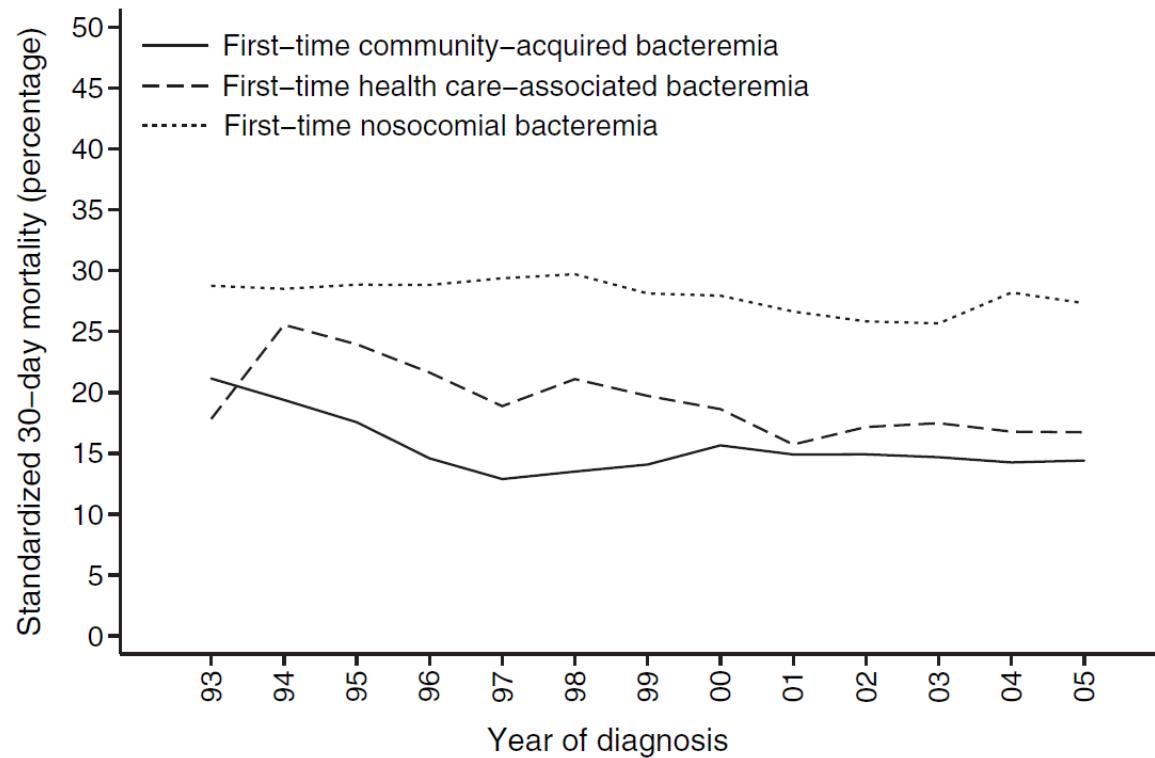


Incidence rates comm.-acq., nosocom. & health-case assoc.

B



Standardized 30-day mortality



Nosocomial

Health-care assoc.
Comm. acq.

Table 3. Crude and Adjusted 30-Day Mortality in Patients with a First-Time Diagnosis of Bacteremia from 1992 through 2006 according to Study Period and Place of Acquisition

Variable	Study period	No. of patients	30-Day mortality		
			No. (%) of patients who died	Crude OR (95% CI)	Adjusted ^a OR (95% CI)
Overall incident episodes of bacteremia	1992–1996	3267	742 (22.7)	1.0 (Reference)	1.0 (Reference)
	1997–2001	3934	835 (21.2)	0.92 (0.82–1.03)	0.77 (0.68–0.88)
	2002–2006	4502	926 (20.6)	0.88 (0.79–0.98)	0.73 (0.64–0.82) 
First-time community-acquired bacteremia	1992–1996	1772	336 (19.0)	1.0 (Reference)	1.0 (Reference)
	1997–2001	1923	291 (15.1)	0.76 (0.64–0.91)	0.66 (0.54–0.80)
	2002–2006	2246	345 (15.4)	0.78 (0.66–0.91)	0.63 (0.52–0.76) 
First-time nosocomial bacteremia	1992–1996	1277	356 (27.9)	1.0 (Reference)	1.0 (Reference)
	1997–2001	1609	468 (29.1)	1.06 (0.90–1.25)	0.97 (0.81–1.16)
	2002–2006	1733	479 (27.6)	0.99 (0.84–1.16)	0.84 (0.70–1.00) 
First-time health care-associated bacteremia	1992–1996	278	65 (23.4)	1.0 (Reference)	1.0 (Reference)
	1997–2001	730	161 (22.1)	0.93 (0.67–1.29)	0.72 (0.50–1.03)
	2002–2006	967	212 (21.9)	0.92 (0.67–1.26)	0.72 (0.51–1.03) 

NOTE. CI, confidence interval; OR, odds ratio.

^a Adjusted for place of acquisition, age, sex, level of comorbidity, marital status, and appropriateness of empirical treatment.

Overall reduction in 30-day mortality: 27%

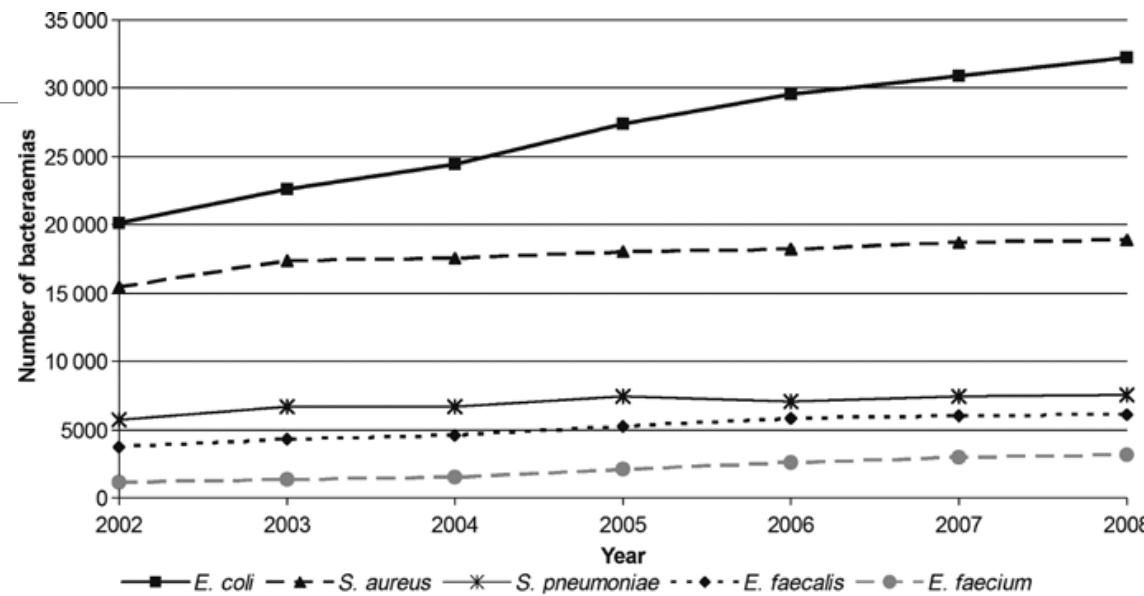
Adj. for age, sex, comorbidity, marital status and empirical treatment

The changing epidemiology of bacteraemias in Europe: trends from the European Antimicrobial Resistance Surveillance System

M. E. A. de Kraker^{1,2}, V. Jarlier³, J. C. M. Monen¹, O. E. Heuer⁴, N. van de Sande⁵ and H. Grundmann^{1,2}

Clin Microbiol Infect 2012

The changing epidemiology of bacteraemias in Europe: trends from EARSS



Pathogen	2002	2003	2004	2005	2006	2007	2008
<i>E. coli</i>	20 151	22 620	24 371	27 328	29 543	30 931	32 194
<i>S. aureus</i>	15 422	17 325	17 550	18 003	18 210	18 665	18 894
<i>S. pneumoniae</i>	5725	6674	6721	7458	7051	7421	7555
<i>E. faecalis</i>	3679	4309	4625	5201	5837	6012	6105
<i>E. faecium</i>	1118	1374	1563	2091	2541	2938	3128

Species	Change per year in % (95% CI)	p-value	Relative change per year in % (95% CI)	p-value
<i>E. coli</i>	8.1 (7.8–8.3)	<0.01	1.7 (1.4–1.9)	<0.01
<i>S. aureus</i>	2.8 (2.5–3.1)	<0.01	-3.4 (-3.7 to -.1)	<0.01
<i>Streptococcus pneumoniae</i>	3.8 (3.4–4.3)	<0.01	-2.4 (-2.9 to -.0)	<0.01
<i>Enterococcus faecalis</i>	8.7 (8.2–9.3)	<0.01	2.3 (1.7–2.8)	<0.01
<i>Enterococcus faecium</i>	19.3 (18.3–20.3)	<0.01	12.4 (11.5–13.4)	<0.01

Only laboratories that consistently reported for all years were included. Results are based on generalized linear models with Poisson distribution.

Defining the epidemiology of bloodstream infections: the ‘gold standard’ of population-based assessment

K. B. LAUPLAND*

Kevin Laupland highlights 5 themes

- Value of a population denominator
- Importance of age and gender standardization for calculation and comparison of incidence rates
- Implications of selected hospital admission
- Implications of referral bias



Project Note

Open Access

Rationale for and protocol of a multi-national population-based bacteremia surveillance collaborative

Kevin B Laupland^{*1}, Henrik C Schønheyder², Karina J Kennedy³,
Outi Lytykäinen⁴, Louis Valiquette⁵, John Galbraith⁶, Peter Collignon³,
Deirdre L Church¹, Daniel B Gregson¹ and Pamela Kibsey⁶

BMC Research Notes 2009, **2**:146 doi:10.1186/1756-0500-2-146

This collaboration now has participation from regions in Canada, England, Denmark, Sweden, Finland, Germany and Australia.

The group plans to meet in Copenhagen in April 2015 in connection with the 25th ECCMID.

The group is open for new participants!

Utilization of blood cultures in Danish hospitals: a population-based descriptive analysis

S. Gubbels¹, J. Nielsen¹, M. Voldstedlund¹, B. Kristensen², H. C. Schønheyder^{4,5}, C. M. J. E. Vandebroucke-Grauls⁶, M. Arpi⁷, M. K. Björnsdóttir³, J. Dahl Knudsen⁸, R. B. Dessau⁹, T. Gorm Jensen¹⁰, P. Kjældgaard¹¹, L. Lemming¹², J. K. Møller¹³, D. Schrøder Hansen¹⁴ and K. Mølbak¹

1) Department of Infectious Disease Epidemiology, 2) Department of Microbiology and Infection Control, Statens Serum Institut, 3) Department of Clinical Microbiology, Rigshospitalet, Copenhagen, 4) Department of Clinical Microbiology Aalborg University Hospital, Aalborg, 5) Department of Clinical Medicine, Aalborg University, Aalborg, Denmark, 6) Department of Medical Microbiology and Infection Control, VU University Medical Center, Amsterdam, The Netherlands, 7) Department of Clinical Microbiology, Herlev Hospital, Herlev, 8) Department of Clinical Microbiology Hvidovre Hospital, Hvidovre, 9) Department of Clinical Microbiology, Zealand Region, Slagelse, 10) Department of Clinical Microbiology, Odense University Hospital, Odense, 11) Department of Clinical Microbiology, Sygehus Sønderjylland, Sønderborg, 12) Department of Clinical Microbiology, Aarhus University Hospital, Aarhus, 13) Department of Clinical Microbiology, Vejle Hospital, Vejle and 14) Department of Clinical Microbiology, Hillerød Hospital, Hillerød, Denmark

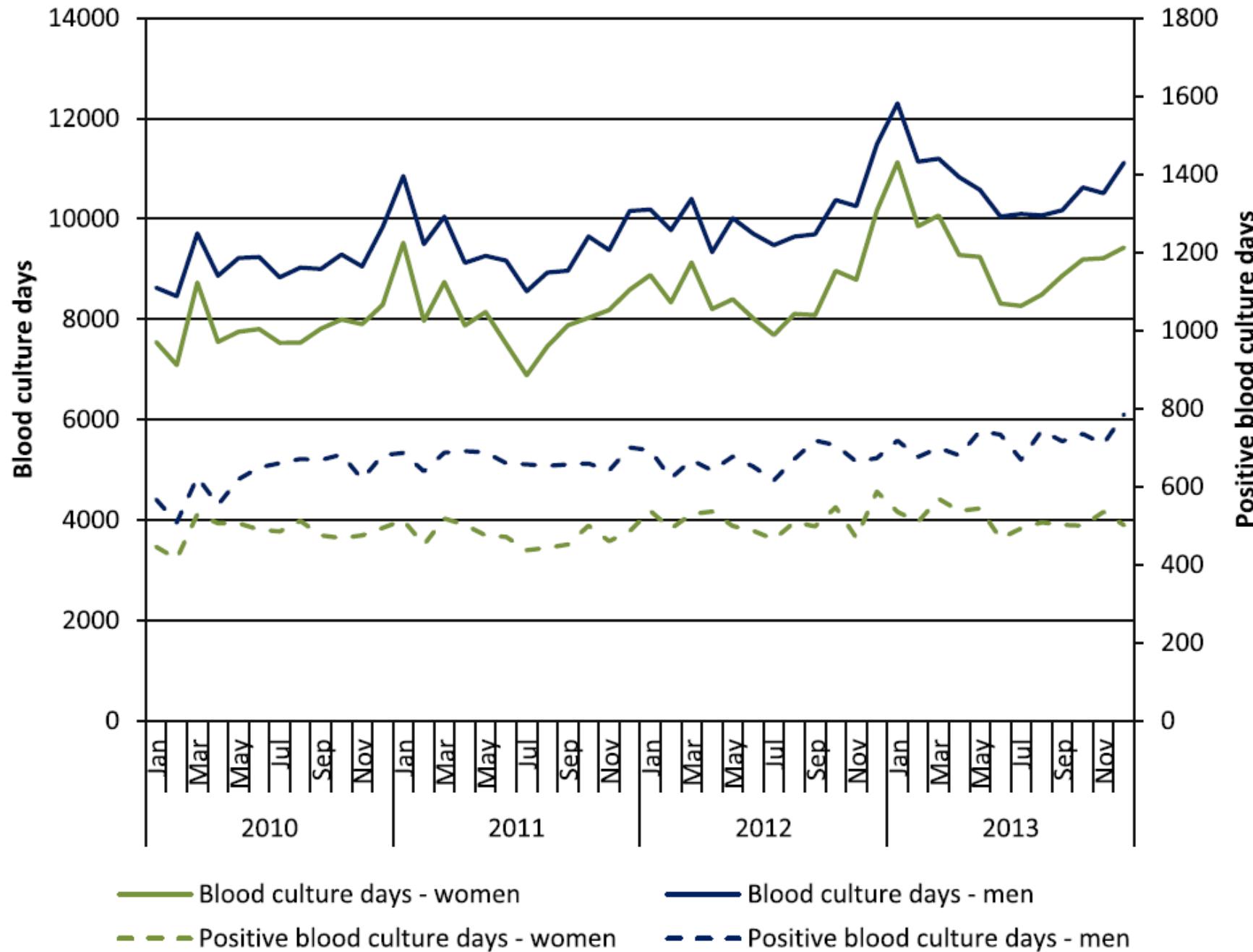
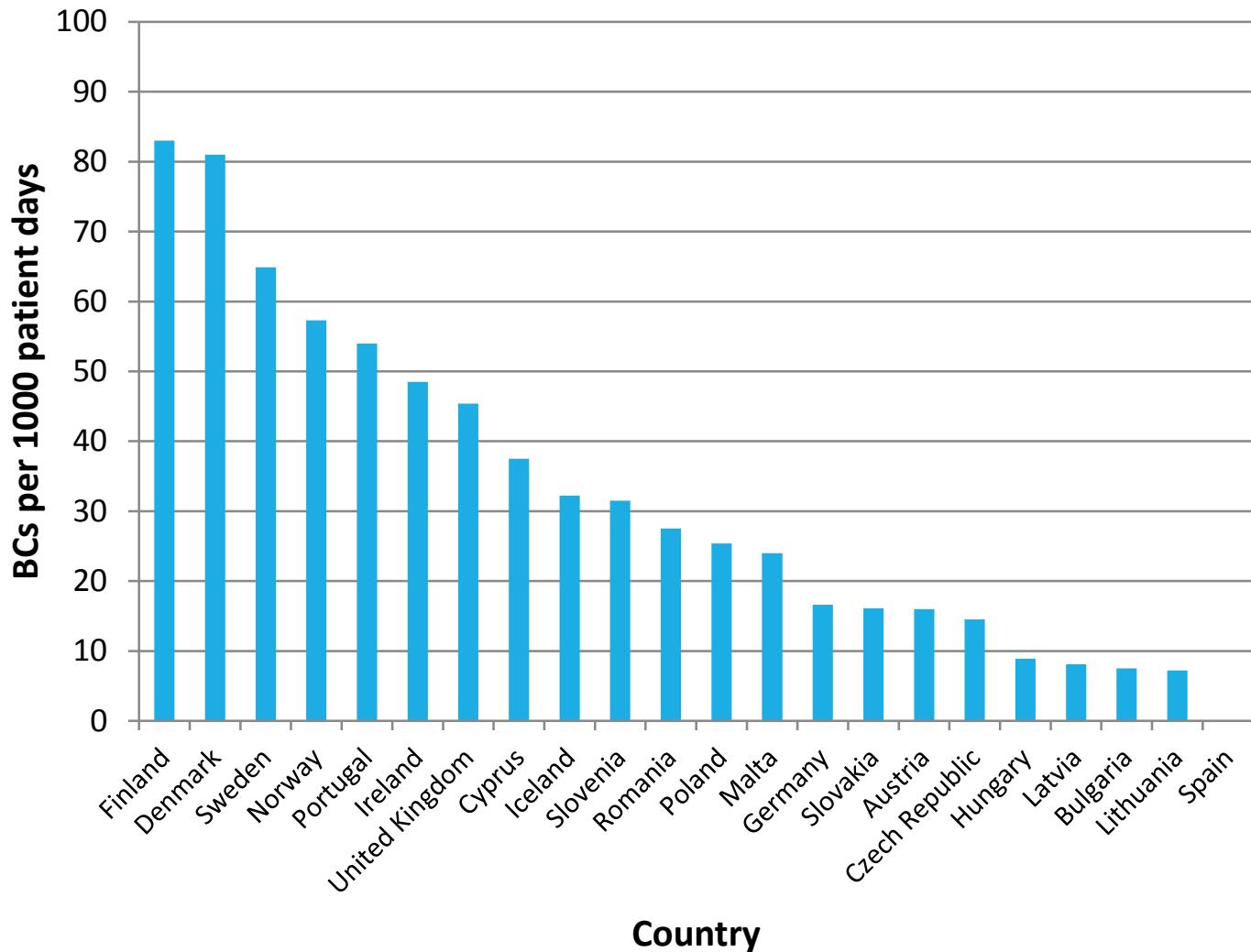


FIG. I. Number of blood culture (BC) days and positive BC days stratified by sex between 2010 and 2013.

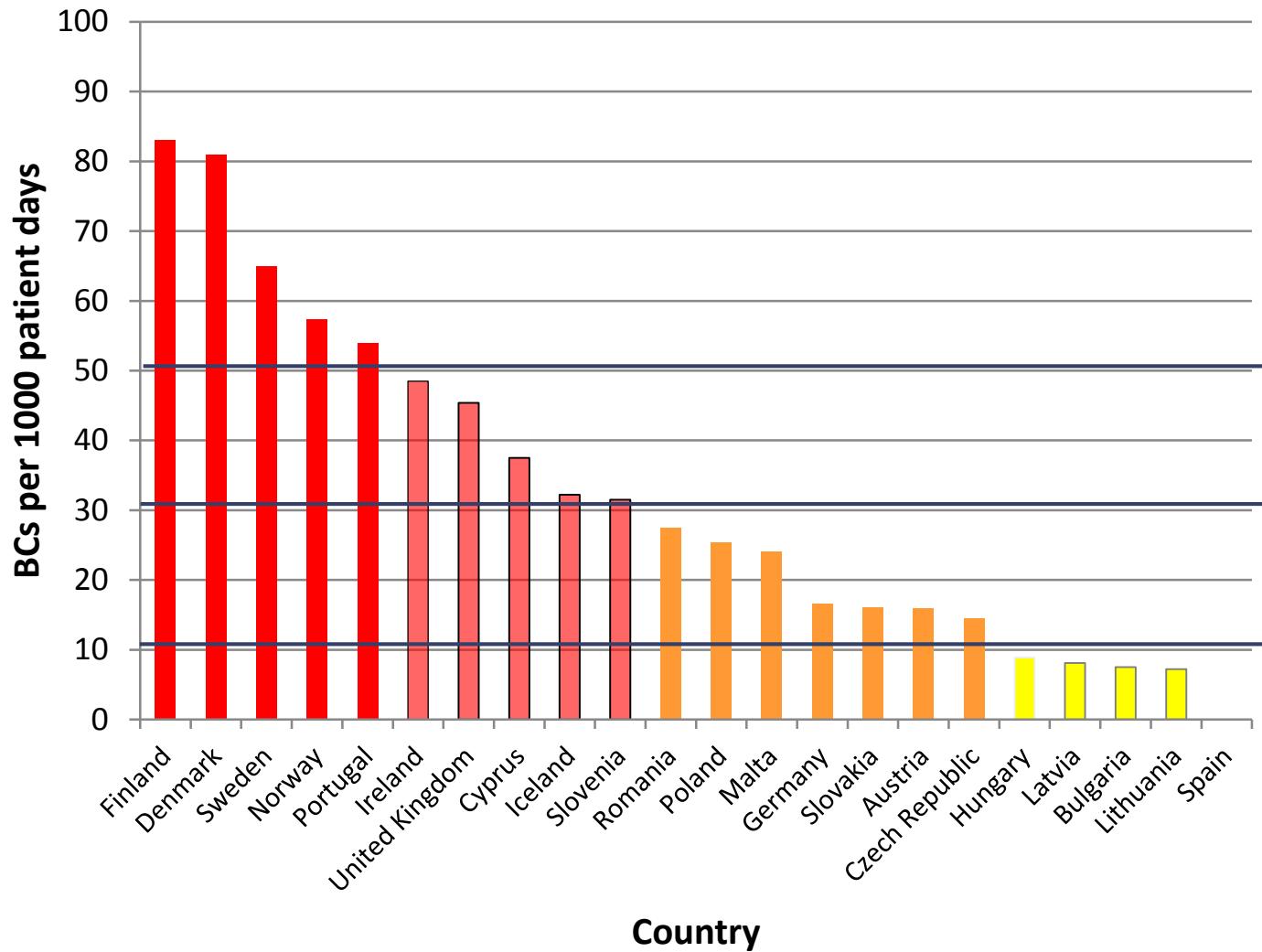
TABLE I. Ten most common pathogenic microorganisms per blood culture day according to department of clinical microbiology between 2010 and 2013

Departments of clinical microbiology	<i>Escherichia coli</i>		<i>Staphylococcus aureus</i>		<i>Klebsiella pneumoniae</i>		<i>Streptococcus pneumoniae</i>		<i>Enterococcus faecium</i>		<i>Enterococcus faecalis</i>		<i>Pseudomonas aeruginosa</i>		<i>Candida albicans</i>		<i>Enterobacter cloacae</i>		<i>Klebsiella oxytoca</i>		Total
	n	%	n	%	n	%	n	%	n	%	n	%	n	%	n	%	n	%	n	%	n
Aalborg	1814	41.2	738	16.7	395	9.0	391	8.9	198	4.5	240	5.4	202	4.6	154	3.5	139	3.2	137	3.1	4408
Aarhus	2040	37.0	931	16.9	508	9.2	476	8.6	437	7.9	330	6.0	229	4.1	217	3.9	179	3.2	173	3.1	5520
Esbjerg	628	39.8	279	17.7	139	8.8	122	7.7	67	4.2	102	6.5	69	4.4	85	5.4	42	2.7	44	2.8	1577
Herlev/Hvidovre/ Hillerød	5339	42.1	2201	17.3	1110	8.7	1145	9.0	702	5.5	812	6.4	439	3.5	298	2.3	317	2.5	326	2.6	12 689
Herning/Viborg	1332	38.0	634	18.1	345	9.8	345	9.8	160	4.6	285	8.1	121	3.4	70	2.0	97	2.8	120	3.4	3509
Odense	1616	32.8	787	16.0	415	8.4	324	6.6	568	11.5	437	8.9	211	4.3	233	4.7	182	3.7	147	3.0	4920
Rigshospitalet	567	20.0	610	21.5	327	11.5	78	2.7	523	18.4	245	8.6	114	4.0	168	5.9	132	4.6	78	2.7	2842
Slagelse	2676	42.1	1077	16.9	630	9.9	586	9.2	272	4.3	363	5.7	230	3.6	152	2.4	178	2.8	198	3.1	6362
Sønderborg	577	45.2	224	17.5	116	9.1	108	8.5	48	3.8	77	6.0	37	2.9	22	1.7	36	2.8	32	2.5	1277
Vejle	923	44.2	324	15.5	179	8.6	187	9.0	127	6.1	118	5.6	90	4.3	39	1.9	47	2.2	55	2.6	2089
Total	17 512	38.7	7805	17.3	4164	9.2	3762	8.3	3102	6.9	3009	6.7	1742	3.9	1438	3.2	1349	3.0	1310	2.9	45 193

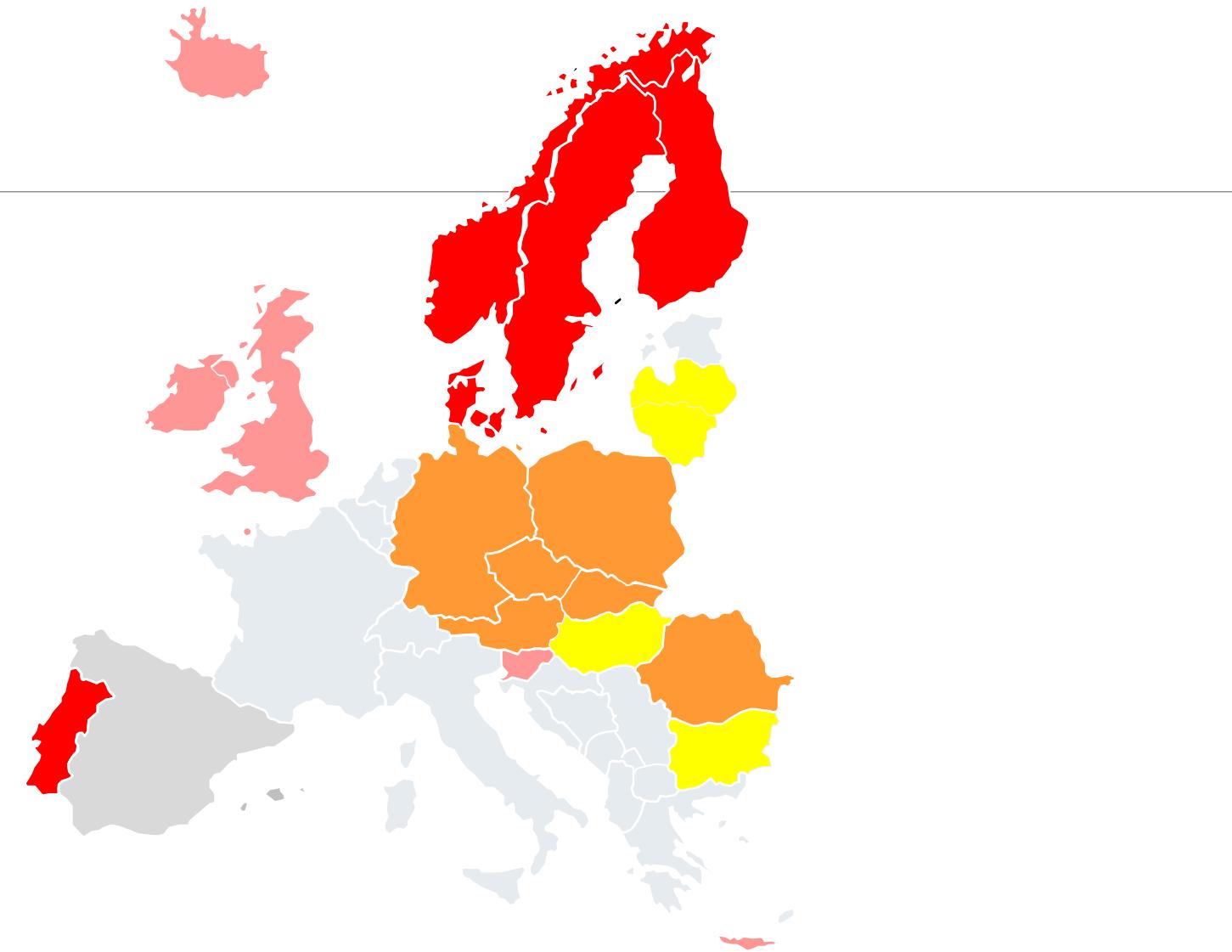
Data from the EARSS 2012 Surveillance Report



Data from the EARSS 2012 Surveillance Report



Per 1000 patient days	
Red	>50 BCs
Pink	31-50 BCs
Yellow	10-30 BCs
Light Yellow	>10 BCs
Grey	No statistics
Light Grey	No data



Monitoring the epidemiology of bloodstream infections: aims, methods and importance

Expert Rev. Anti Infect. Ther. 11(12), 1281–1290 (2013)

Mette Søgaard^{*1},
Outi Lyytikäinen²,
Kevin B Laupland³ and
Henrik Carl
Schönheyder⁴

Bloodstream infections (BSI) are a major cause of mortality, morbidity and medical cost. Even though monitoring activities have been on-going for decades, it is difficult to depict a full picture of the burden of BSI. The main reasons for shortcomings include varying study aims, definitions and inclusion criteria for both microorganisms and patients. Incidence studies are commonly hampered by difficulties in delineating the population at risk. The objective of this review was to provide a framework for comprehensive BSI monitoring systems in the future. We highlight the importance of standardized definitions and acquisition of data combined with cautious statistical analyses. Hereby, valid data on BSI can be provided for clinicians and decision makers and ultimately contribute to improvement of the quality of care for BSI patients.

KEYWORDS: bacteremia • epidemiology • monitoring • outcome • surveillance

Box 1. Desirable features of bloodstream infections monitoring programs.

- Clearly defined objectives
- Unambiguous case definition
- High degree of data completeness and accuracy
- Well-defined population denominator data
- Reproducible data
- Data on vital status
- Standardized coding schemes and terminology
- Minimal extra resources
- Integration with other health care databases
- Rapid feedback of information to providers
- Data informing local services and commissioners

Table 3. Bloodstream infections that are currently targeted for monitoring in selected European countries. Supply of data is either mandatory (M) or voluntary (V).

Agents	DK [104]	FI [102]	NO [105]	SE [106]	UK [103]
All BSIs		M		V	
Polymicrobial BSI				V	
Individual pathogens					
<i>Staphylococcus aureus</i>	V	M	M	M	M
<i>Streptococcus pneumoniae</i>	M	M	M	M	V
HS	V			V	
HS group A		M	M	M	
HS group B		M	M		
<i>Listeria monocytogenes</i>	M	M	M	M	V
<i>Neisseria meningitidis</i>	M	M	M	M	M
<i>Haemophilus influenzae</i>	M	M	M	M	M
<i>Escherichia coli</i>		M			M
<i>Salmonella</i> serovar Typhi	M	M	M	M	M
<i>Salmonella</i> serovar Paratyphi	M	M	M	M	M
Zoonotic <i>Salmonella</i>			M		M
Fungemias		M		V	
Resistance phenotype					
MRSA	M	M [†]	M	M	M
GRE		M		M	M
Enterobacteriaceae producing ESBL or ESBL-CARBA		M [‡]		M	
Place of acquisition					
Nosocomial BSIs					