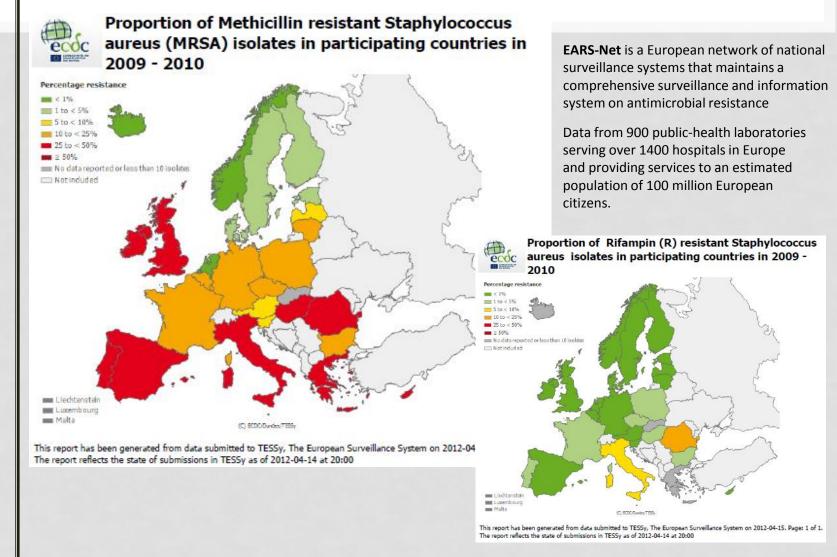


Central Hospital of the German Federal Armed Forces, Koblenz, Germany
Department Trauma surgery and Orthopedics, Reconstructive,
Hand and Plastic surgery, Burn medicine
Medical director: Colonel Privatdozent Dr. med. E. Kollig

INFECTIONS AFTER ABROAD SITUATIONS



European Antimicrobial Resistance Surveillance Network (EARS-Net)



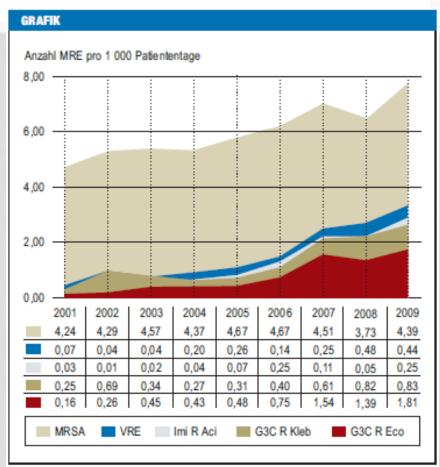


Most suspected in GE: MRSA

TABELLE 1							
Definition der Multiresistenz von gramnegativen Erregern (MR-ONE)							
Penicilline	Cephalosporine	Carbapeneme	Fluorchinolone				
Enterobacteriaceae (z. B. E. coli, K. pneumoniae, E. cloacae)							
Leitsubstanz Piperacillin- Tazobactam* ¹	Leitsubstanz Cefotaxim Cefotaxim Leitsubstanz Impenem oder Meropenem oder Ertapenem* ¹		Leitsubstanz Ciprofloxacin				
R	R	S	R				
R	R	R	S				
R	S	R	R				
R	R	R	R				
	Pseudomona	as aeruginosa					
Leitsubstanz Piperacillin oder Piperacillin- Tazobactam	Leitsubstanz Ceftazidim	Leitsubstanz Meropenem	Leitsubstanz Ciprofloxacin				
R	R	S	R				
R	R	R	S				
R	S	R	R				
R	R	R	R				
	Acinetobacte	r baumannii*2					
-	-	Leitsubstanz Imipenem	Leitsubstanz Ciprofloxacin				
-	-	S	R				
-	-	R	S				
-	-	R	R				
		cictant: S cancibal: _ kain					

R, resistent; 5, sensibel; -, keine ausreichende Wirksamkeit

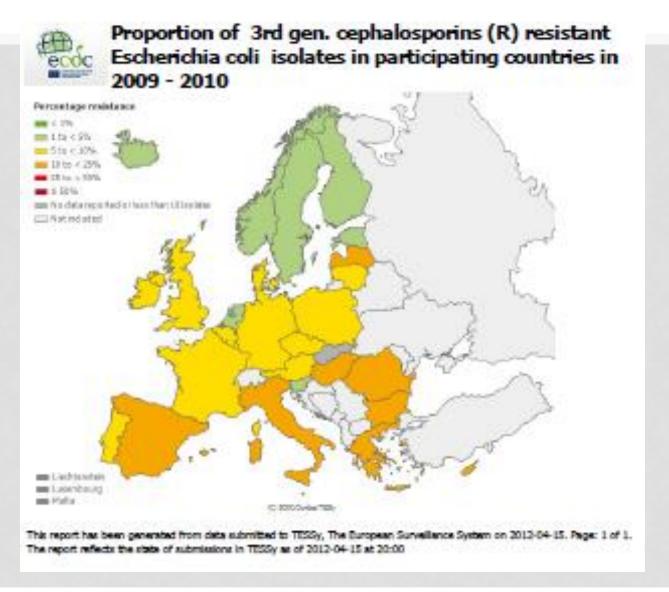
Deutsches Ärzteblatt | Jg. 109 | Heft 3 | 20. Januar 2012



Darstellung von Inzidenzdichten von 55 Intensivstationen, die am SARI-Projekt teilnehmen. Patienten mit multiresistenten Erregem (MRE) pro 1 000 Patiententage (2). SARI, Surveillance der Antibiotika-Anwendung und der bakteriellen Resistenzen auf Intensivstationen; MRSA, Methicillin-resistenter S. aureus; VRE, Vancomycin-resistenter Enterococcus faecium; Imi R Aci, Imipenem-resistenter Acinetobacter baumanii; G3C R Kleb, Gruppe-3-Cephalosporin-resistente Klebsiella pneumoniae; G3C R Eco, Gruppe-3-Cephalosporin-resistente Escherichia coli

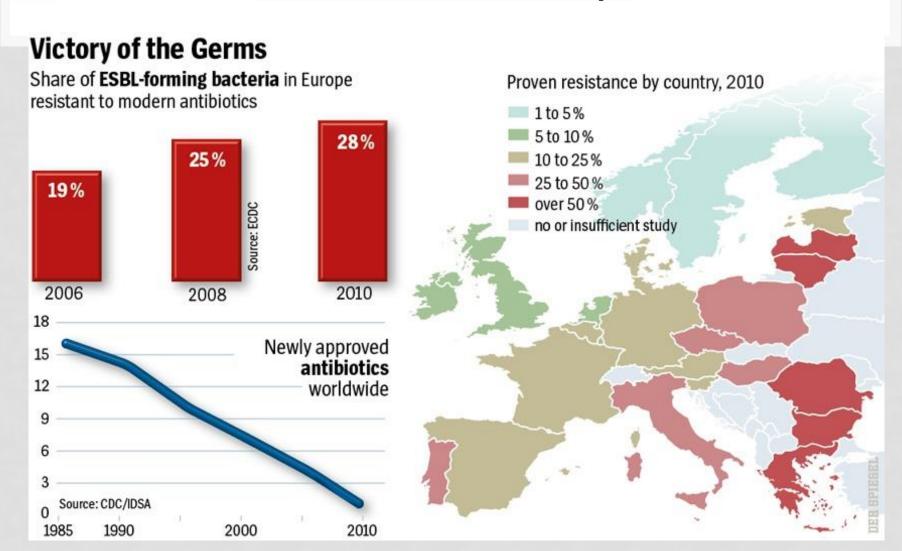


Gram-negative bacteria: situation in Europe





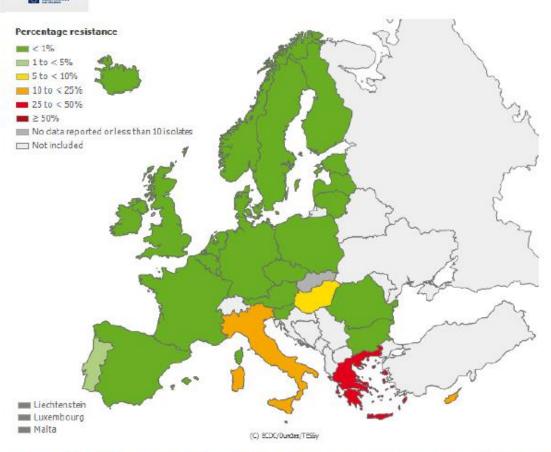
ESBL: situation in Europe





Carbapenem-resistance: the European issue



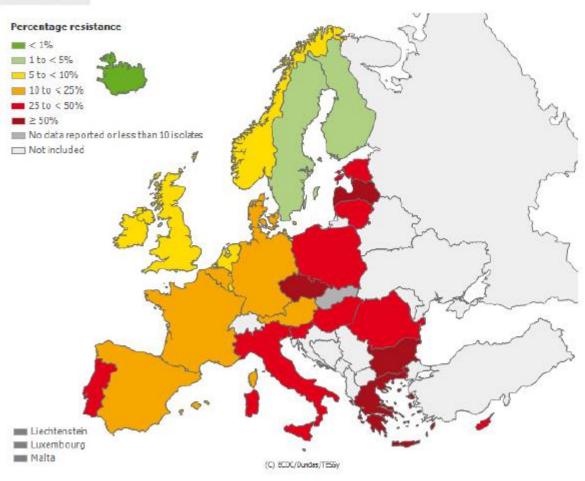


This report has been generated from data submitted to TESSy, The European Surveillance System on 2012-04-24. Page: 1 of 1. The report reflects the state of submissions in TESSy as of 2012-04-24 at 20:00

the metamorphosis of Klebsiella P.



Proportion of Fluoroquinolones (R) resistant Klebsiella pneumoniae isolates in participating countries in 2010



This report has been generated from data submitted to TESSy, The European Surveillance System on 2012-04-24. Page: 1 of 1. The report reflects the state of submissions in TESSy as of 2012-04-24 at 20:00



<u>Different scenarios – different germs</u>

One year ago not business as usual: Wound management, infection and psychoemotional control during tertiary medical care following the 2004 Tsunami disaster in southeast Asia

Marc Maegele^{1,2}, Sven Gregor³, Nedim Yuecel¹, Christian Simanski¹, Thomas Pattrath¹,
Dieter Rixen¹, Markus M Heiss³, Claudia Rudrott³, Stetan Saad³, Walter Perbix⁴, Frank Wappler⁵,
Andreas Harzheim⁶, Rosemarie Schwarz⁷ and Bertil Bouillon¹

Critical Care 2006, 10:R50 (doi:10.1186/cc4868)



Trauma-related Infections in Battlefield Casualties From Iraq

Annals of Surgery * Volume 245, Number 5, May 2007

Kyle Petersen, DO, * Mark S. Riddle, MD, MPH, TM, † Janine R. Danko, MD, MPH, *

David L. Blazes, MD, MPH, ‡ Richard Hayden, MS, Mt(ASCP)SBB,§ Sybil A. Tasker, MD, *

and James R. Dunne, MD|

Bacterial wound cultures:

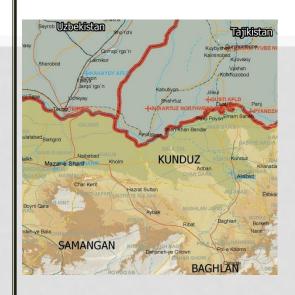
- Acinetobacter baumanii
- Enterococcus faecium
- Escherischia coli
- Proteus vulgaris
- Pseudomonas aeruginosa
- MRSA
- Stenotrophomonas maltophila
- Candida, Aspergillus, Fusaria

TABLE 3.	Distribution	of the 7	Most	Common	Organisms	bν	Culture S	ite
I ADEL J.	DISCIDLATION	Of the /	IVIOSE	CONTINUOU	Organisms	L y	Culture 3	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,

	Bacteria Isolates (N = 120)							
Specimen Site [no. (%)]	Acinetobacter spp.	E. coli	Pseudomonas spp.	Coagulase-Negative Staphylococcus	Enterobacter spp.	Klebsiella spp.	Proteus spp.	Total
Wound	33 (75)	18 (90)	17 (85)	3 (25)	9 (100)	4 (50)	7 (100)	91 (76)
Blood	11 (25)	1(5)	3 (15)	9 (75)	0	3 (37.5)	0	27 (23)
CSF	0	1 (5)	0	0	0	1 (12.5)	0	2(2)
Total	44	20	20	12	9	8	7	120

spp. indicates species; CSF, cerebrospinal fluid.

GE - Investigations from Maza e Sharif:

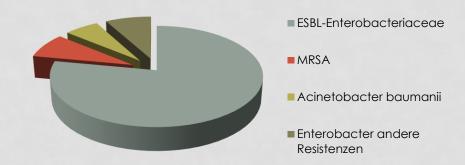


Vandersee, S. et al., HygMed 2011; 36-10, Sei. 384-392

Kolonisationsprävalenz der afg. Patienten:

29 % MRE+ (auf ICU 46%)

Anteil MREs am Gesamtprobenaufkommen 05 / 2010 - 09 / 2010



Sutter, DE. Et al., InfectControlHosp Epidemiolog 2011; 32(9): 854-60

Kolonisationsprävalenz bei 411 afg. Patienten 09/2007 – 08/2008:

- 51 % MRE+
- 76 % gram negative Keime (70% MRE's)
- Gram MRE's:
 - 53% der E. coli, 90% der Acinetobacter, 60% der Klebsiellen

→ gram-negative MRE: dominating bacteria in crisis areas?



Gram-negative bacteria in combat related wounds: - US experience -

"After leaving the combat zone, patients are presenting to US military hospitals with a much higher rate of MDR gram-negative bacteria colonizing and infecting wounds".

Murray C et al; JOT 2011; 71: 235-257

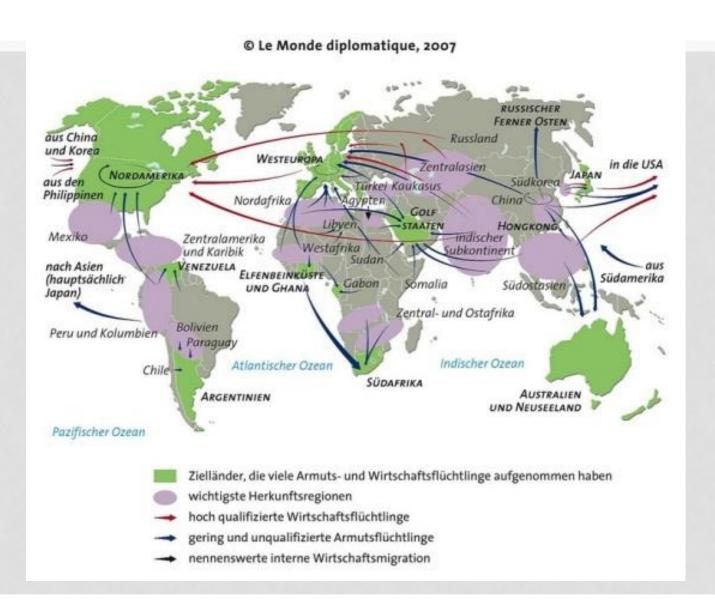


Causes of resistance?

- low standards of development and hygienics
- free available, low-cost standard antibiotics
- overuse / misuse of antibiotics
- permanent reservoir in GIT and UGT
- travelling of germ vectors
- plasmid-hospitalism
- mutations and horizontal gen-transfer



Bacterial traffic worlwide: partially migration related?





MDR identified as / to be differentiated

- colonization (skin, GIT, respir. system)
- contamination (bacterial wound cultures pre-/post debridement)
- infection (proof of bacteria after onset)





"after leaving the combat zone, patients are presenting to US military hospitals with a much higher rate of MDR gram-negative bacteria colonizing and infecting wounds."

Murray C et al; JOT 2011; 71: 235-257



Established infection after ORIF





the sword of Damocles





Civilian benchmark: Bone infection rates in Germany

- heterogenous data available
- no current data since last 5 years
- after closed fx: 0,5% 3,0%
- after open fx: 2,6% 10%
- after osteosynthesis: 0,6% 3,4%

AH Tiemann, R Braunschweig, GO Hoffmann: Knocheninfektionen, Unfallchirurg (2012), S. 480 – 488]



MDR and combat related wounds: US experience

"Increasing colonization with MDR bacteria throughout the evacuation chain from the combat zone, through Germany, to the US supports the concept, that most MDR bacteria colonization and infection is health care associated."

Hospenthal DR et al; JOT 2011; 71: 210-234



Complications in combat related injuries: the large scale (USA)

- 7,4% primary amputation
- 15% osteomyelitis
- 17% relapse of infection
- primordial no typical risk factors acc. to young, healthy milit.
 personnel

Murray C et al; JOT 2011; 71: 235-257

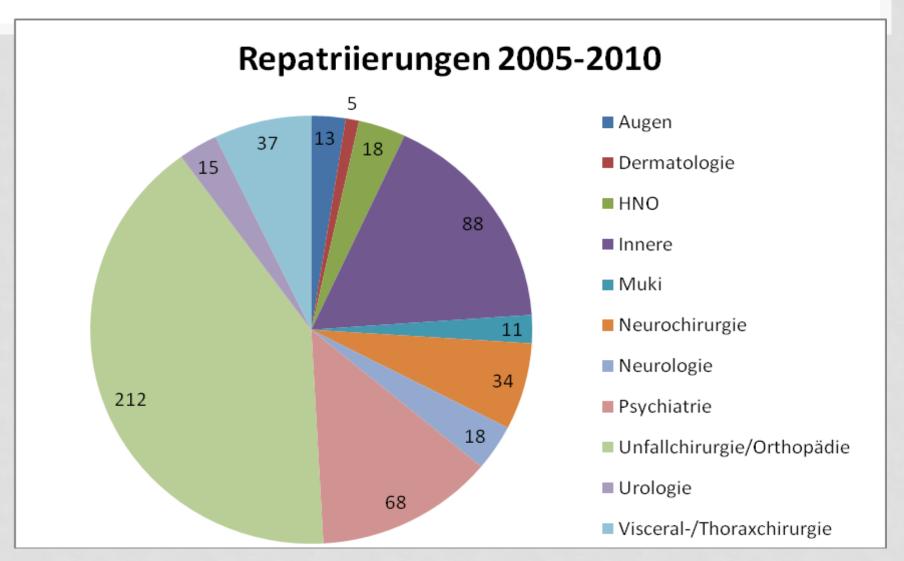


PRINCIPLES OF COMBAT RELATED INJURIES:

- in gunshot and blast-injuries gross contamination is obligatory
- open wounds, damage in the deep easily underestimated
- So called "developing wounds"
- Soft tissue determines preservation of bone and so called guiding structures, thereby the fate of a limb









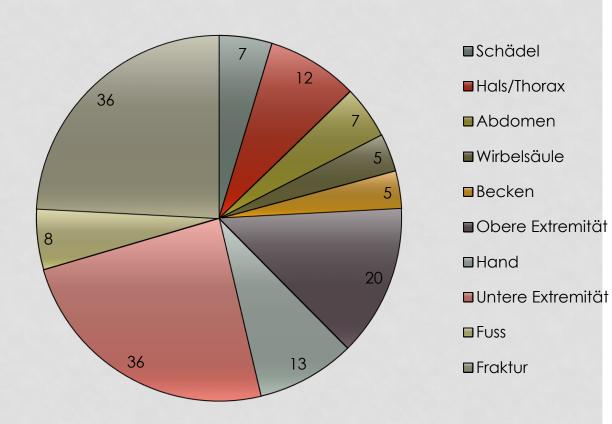


Lokalisation Schussverletzungen (113 Verletzungen bei 79 Patienten)

2004 – 2015

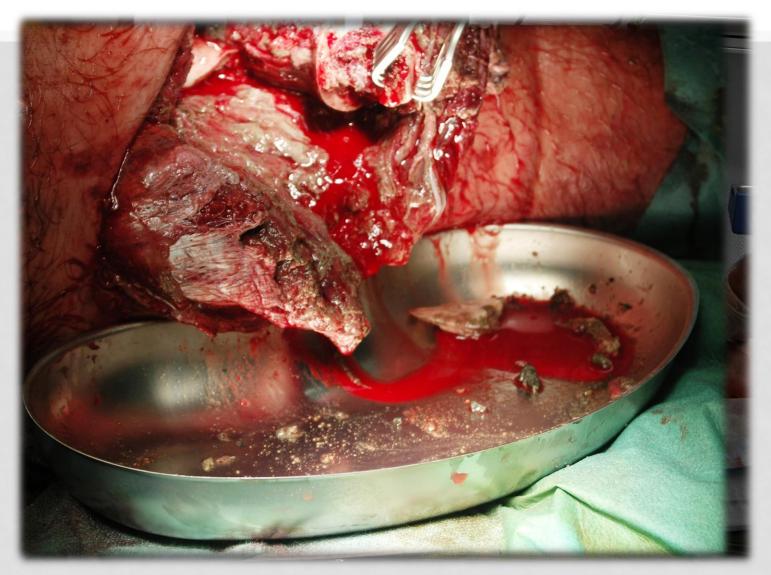
combat: 40

accident: 37



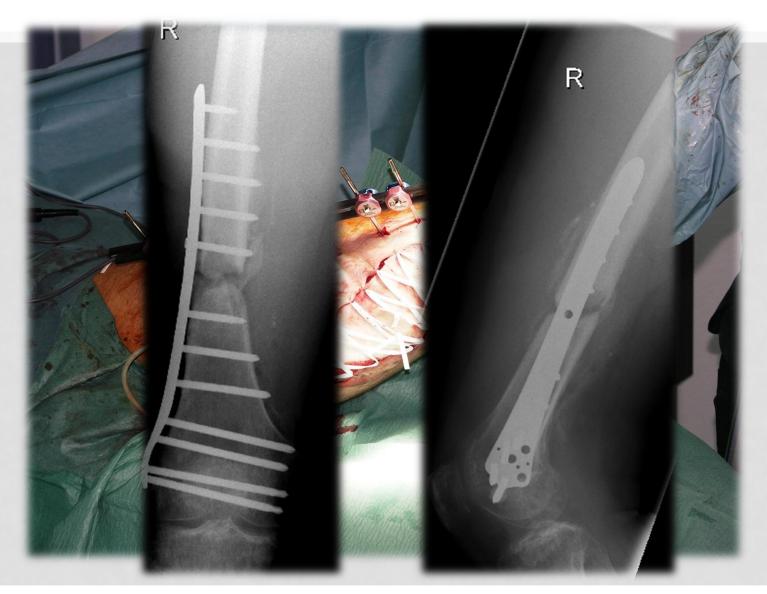


Blast injury caused by IED





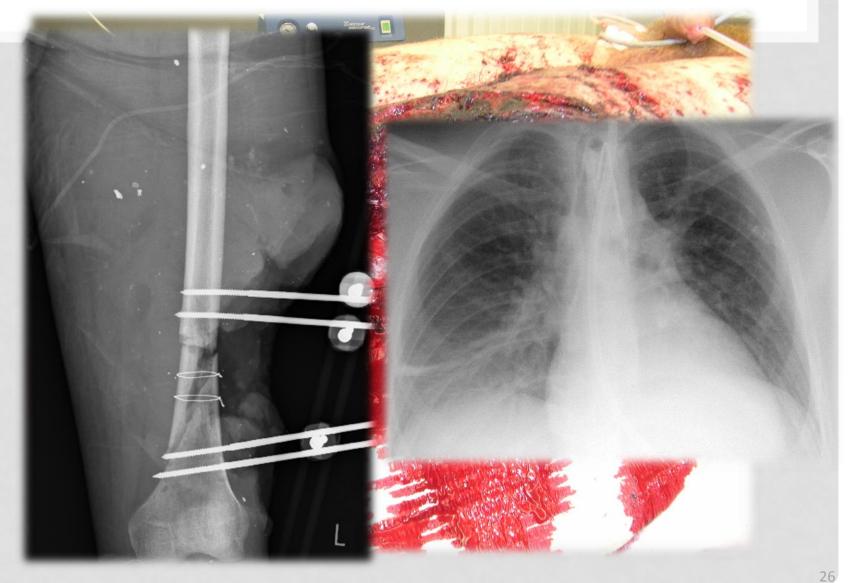
Blast injury caused by IED





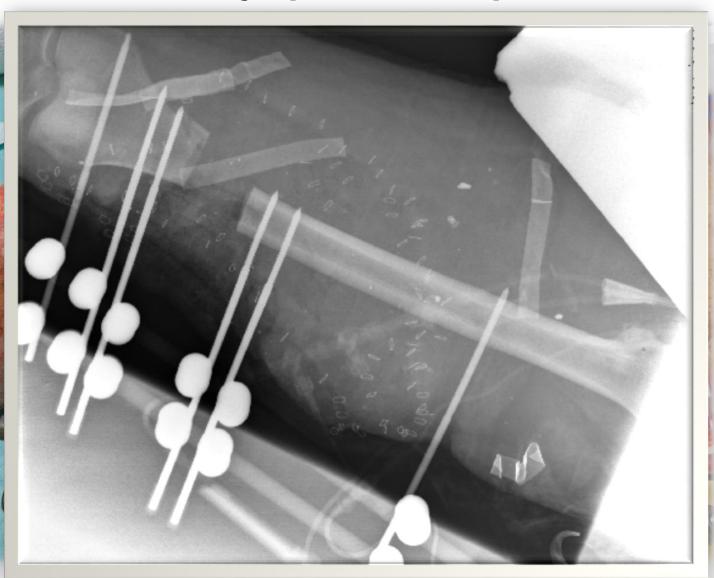






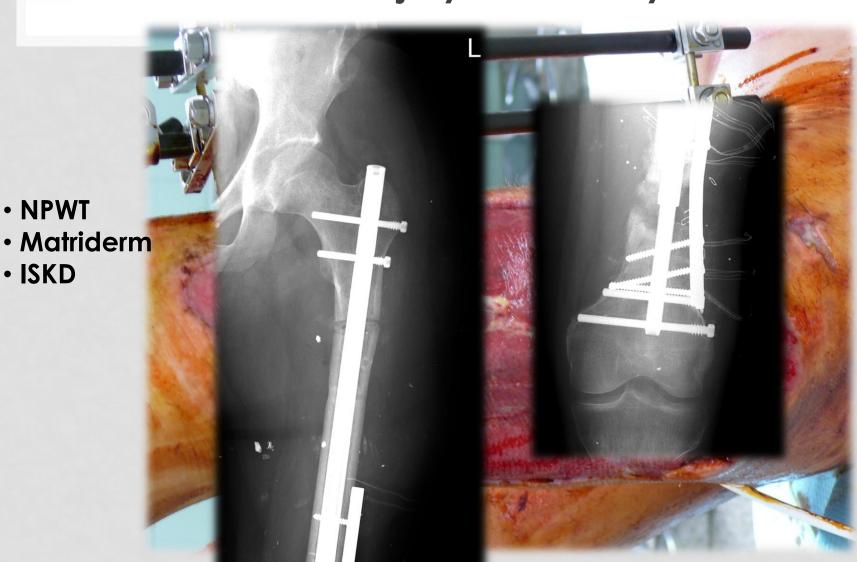


Blast injury caused by IED





Blast injury caused by IED





Blast injury caused by IED: 12 months after











The Lybian experience 2011 - 12





The Lybian experience:

- 10 patients from Lybia, all wounded and pretreated up to three months
- 2/3 lower limb, 1/3 upper limb
- 10/10 with established infection /wound drainage
- > 50% bone defect 50 to 80mm



Cohort- and single - isolation













<u>The Lybian experience – bacteriology:</u>

- bacterial cultures of wounds and surface
- 9/10 positive for MRE
- 3/10 positive for MRSA additionally
- 4/10 Pseudom. Aeruginosa
- 4/10 Acinetob. Baumannii
- 1/10 positive blood-culture



Spectrum of bacteria in combat related wounds

- S. aureus
- MRSA (Methicillin-resistant, "multiresistant")
- MRSE (Multiresistente S. epidermidis specimen)
- VRE
- Pseudomonas aeruginosa
- Klebsiella pneumoniae, Klebsiella pneumoniae (ESBL-pos.)
- Proteus species
- Enterobacter cloacae, Enterobacter cloacae (ESBL-pos.)
- Acinetobacter baumanii
- Enterococcus faecalis
- High-level-Gentamycin-resistenter Enterococcus
- Serratia marcescens, Serratia marcescens (ESBL-pos.)
- •••••





Auftrag
14.11.2011 309231 LabNr Waterial Fragestellung
Waterial In BK-Medium OS re
Enthabme am 14.11.2011 Probeneingang 14.11.2011 13:20: anger

Befundbericht Bakteriologie / Myko

ENDBEFUND

eine Kopie.

BAKTERIOLOGISCHE KULTUR

Keim 1: Klebsiella pneumoniae ssp. pneumoniae

Wichtiger Hinweis:
Es handelt sich um einen ESBL-Lextended-SpektrumB-Lactamase-)produzierenden Stamm. Auch bei anscheinender in-vitro-Empfindlichkeit sollten
B-Lactam-Antibiotika, einschließlich der Cephalosporine der 3. Generation (Cefotaxim, Ceftriaxon,
Cefpodoxim, Ceftazidim) und Monobactame (Aztreonam) nicht für die Therapie der durch diesen Keim
hervorgerufenen Infektionen eingesetzt werden.
Dies gilt auch für Tazobactam, das aber bei
weitreichender Resistenz noch als Reserveoption genutzt werden kann.
ESBL-produzierende Keime können ein ernsthaftes
krankenhaushygienisches Problem darstellen. Ihre
Weiterverbreitung sollte daher durch entsprechende krankenhaushygienische Maßnahmen unbedingt
verhindert werden.
Von Befunden, in denen ein ESBL-Machneis mitgeteilt wird, erhält der hygienebeauftragte

ANTIBIOGRAMM Ampicillin Ampicillin-Sulbactam Amoxicillin-Clavulans R Cefazolin Cefuroxim Cefotaxim -Gentamicin Amikacin Tobramycin Cotrimoxazol Levofloxacin Ciprofloxacin syst.An Ceftazidim Piperacillin Piperacillin-Tazobact Aztreonam Imipenem Meropenem

(S = sensibel, R = resistent, I = intermediar)
Die Angaben über das Resistenzverhalten wurden
als Minimale Hemmstoff-Konzentrationen (MHK)
ermittelt.



Department Trauma surgery and Orthopedics, Reconstructive,
Hand and Plastic surgery, Burn medicine













reconstruction of bone defect acc. to MASQUELET's method with cancellaus bone graft augmented by platelet rich plasma





one-step revision: relapsing infection thereafter







Infection, sepsis, ESBL:





Patient's bacteria cultures

	wound	skin	blood culture
MRSA		+	
P. aeroginosa	+	+	
A. baumanii	+		
H-L-G-R Enterococcus		+	
ESBL-Klebsielle	++	+	+
P. mirabilis		+	
ESBL-Serratia	++	+	+
S. haemolyticus		+	
C. freundii		+	
E. coli		+	



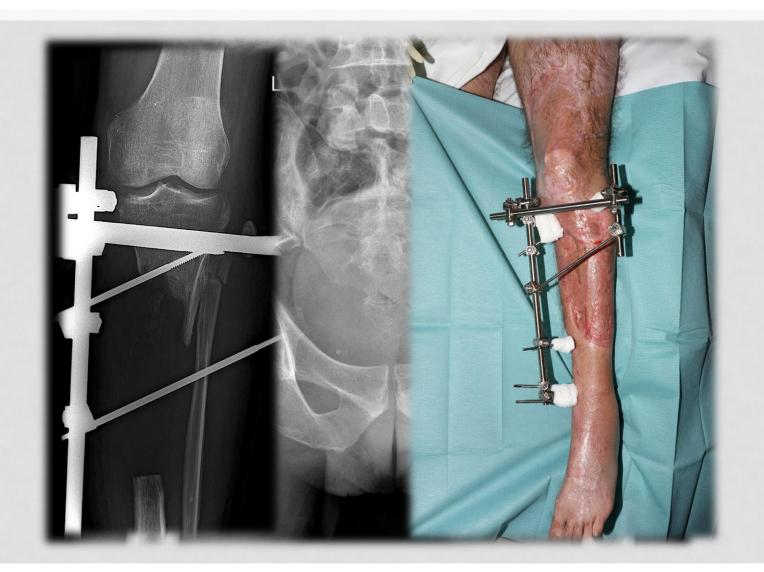
...leaving the internal Titanium fixator in situ: 4 weeks post







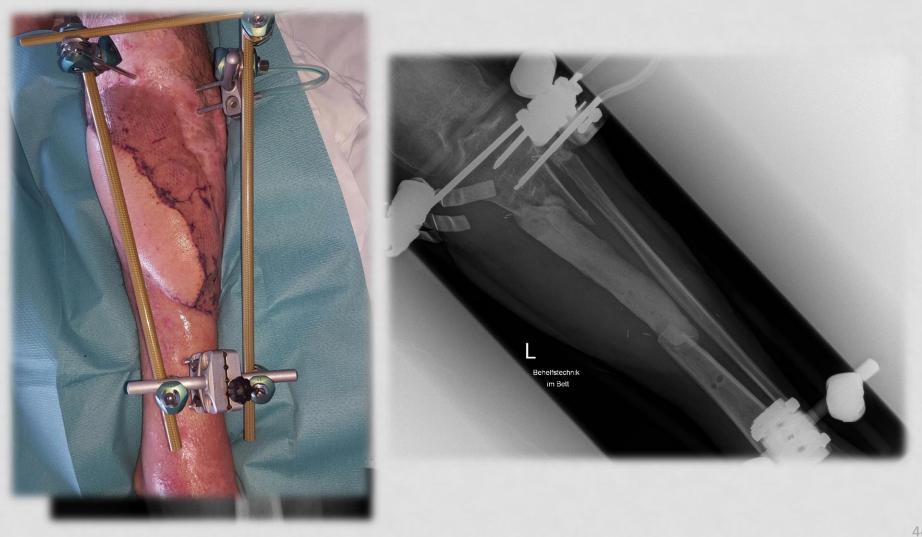
Ukrainian patient, multiple mortar wounds, 3- + 4-MRGN, MRSA





Department Trauma surgery and Orthopedics, Reconstructive Hand and Plastic surgery, Burn medicine

Ukrainian Patient, multiple mortar wounds, 3- + 4-MRGN, MRSA





Ukrainian Patient, multiple mortar wounds, 3- + 4-MRGN, MRSA



PATIENT A

Lokalisation	Wundinfektion
28.06.2017 Wunde (Oberarm links außen und innen)	E.cloacae (ESBL)
05.07.2017 Intraoperative Abstriche Markraum und Frakturregion (Humerus links)	E. cloacae (ESBL) A. baumannii
10.07.2017 Intraoperative Abstriche Wunde (Oberarm links außen)	E.cloacae (ESBL)
17.07.2017 Intraoperative Abstriche Wunde (Oberarm links außen Pinstelle)	A. baumannii (4MRGN)
02.08.2017 Intraoperative Abstriche Wunde (Oberarm links außen Pinstelle	A. baumannii (4MRGN)
18.08.2017 Punktat	Kein Keimnachweis

Kontamination in Ganzkörperabstrichserien

E. cloacae (ESBL)

K. pneumoniae (ESBL)

A. baumannii

K. pneumoniae (ESBL) (4MRGN)

A. baumannii (4MRGN)

PATIENT B

Lokalisation	Wundinfektion
28.06.2017 Wunde (Oberschenkel rechts, tief)	A. baumannii (3MRGN)
30.06.2017 Wunde (Oberschenkel rechts, tief)	A. baumannii (3MRGN und Gentamicin resistent)
03.07.2017 Wunde (Oberschenkel rechts, tief)	A. baumannii (3MRGN Carbapenem sensibel)
17.07.2017 Wunde (Oberschenkel rechts, tief)	A. baumannii, S. lugdunensis, E. faecalis
02.08.2017 Verschlossene Wundverhältnisse, reizfrei	Keimfrei

Kontamination in Ganzkörperabstrichserien

K. pneumoniae (ESBL)

PATIENT C

Lokalisation	Wundinfektion
28.06.2017 Wunde (Hüfte rechts)	Staphylokokkus hominis (Vancomycin sensibel)
04.07.2017 Femur Markraum (rechts)	S. epidermidis (Linezolid sensibel)
04.08.2017 Femur Markraum, Kapsel, Pfannengrund (rechts	E. coli (ESBL) (3MRGN) (nur Tigecyclin und Carbapeneme sensibel)
23.08.2017 PE Zugang Hüfte und Kapsel (rechts)	A. baumannii (4MRGN) (Gentamicin sensibel)
06.09.2017 PE Zugang Hüfte (rechts)	E. coli (ESBL, 3MRGN)
22.09.2017 PE Zugang Hüfte und Wunde Hüfte (rechts)	E. coli (ESBL, 3MRGN) S. Aureus)
05.10.2017 Redondrainage	E. coli (ESBL, 3MRGN)

Kontamination in Ganzkörperabstrichserien

A. baumannii,

K. pneumoniae (ESBL) (3MRGN)

K. pneumoniae (ESBL) (4MRGN)

K. pneumoniae (ESBL) (3MRGN) (Tigecyclin resistent)

E. coli (ESBL) (3MRGN)

E. cloacae (ESBL) (VRE)

PATIENT D

Lokalisation	Wundinfektion
28.06.2017 Wunde (Schulter links, tief)	S. hämolyticus, S. epidermidis
04.07.2017 Wunde (Schulter links, tief)	S. epidermidis (Clindamycin resistent, Gentamicin resistent)
28.07.2017 Primärer Wundverschluss	S. epidermidis (Clindamycin resistent, Gentamicin resistent) Ohne weitere Konsequenz und Wundheilungsstörung

Kontamination in Ganzkörperabstrichserien

K. pneumoniae (ESBL)



Surgical management I

- consequent debridement
- appropriate stabilization / immobilization
- no additional damage by searching for foreign bodies, as long as not responsible for complications
- irrigation no jet-lavage



Surgical management II

- programmated re-do's acc. to development
- NPWT recommended esp. after gross contamination and extended soft tissue damage
- topical use of antiseptic agents (e. g. polyhexanide)
- only viable tissue will heal
- no change to internal fixation methods unless proof of absence of infection is given



management of antibiotics

- flanking procedure
- acc. to resistance
- time is of an essence
- colonization is no target
- recurrent culture controls time scheduled
- "off label" use to be considered



Orthop/Trauma: results since 2005

- no additional amputation necessary
- all infections controlled, no relapse
- colonization can be eradicated is it a must?
- no outbreak of specific bacteria
- combat related infections are highly demanding



Take home messages (Orthop/Trauma):

- identify the gram-negative MRE as a global problem
- initial and repeated bacteriological screening
- strict isolation of patients, rigorous hygienics
- the surgical skills only ensure infection control and healing
- antibiotics only on demand and after proof

