



Problembewusstsein AB – Resistenz aus vet. med. Sicht

Univ. Prof. Tzt. Dr. Josef Köfer

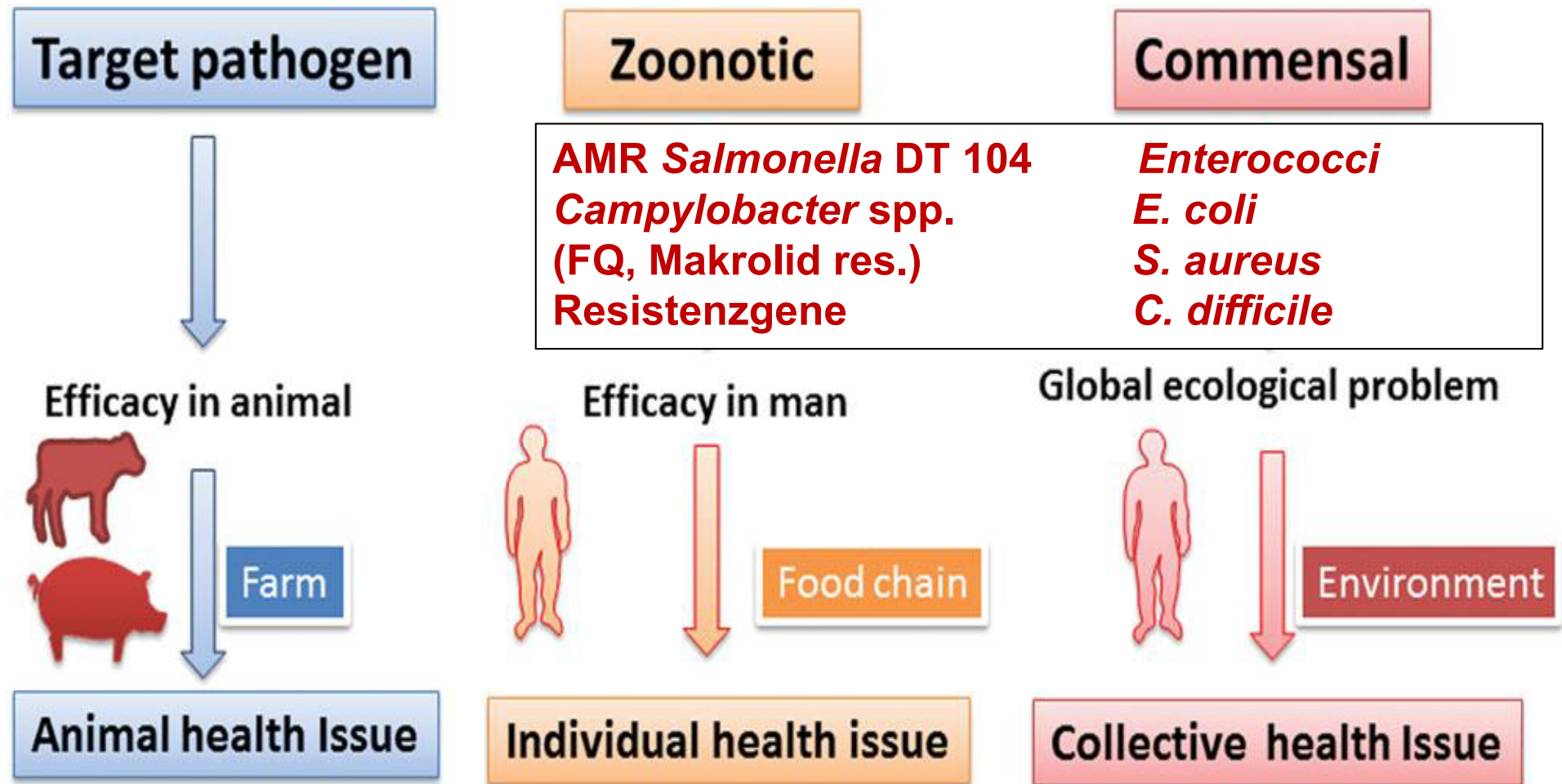
Einfluss von AB – Resistenzen, Gemeinsame Herausforderung für Veterinär- und Humanmedizin

28. März 2017, Steiermarkhof, Graz

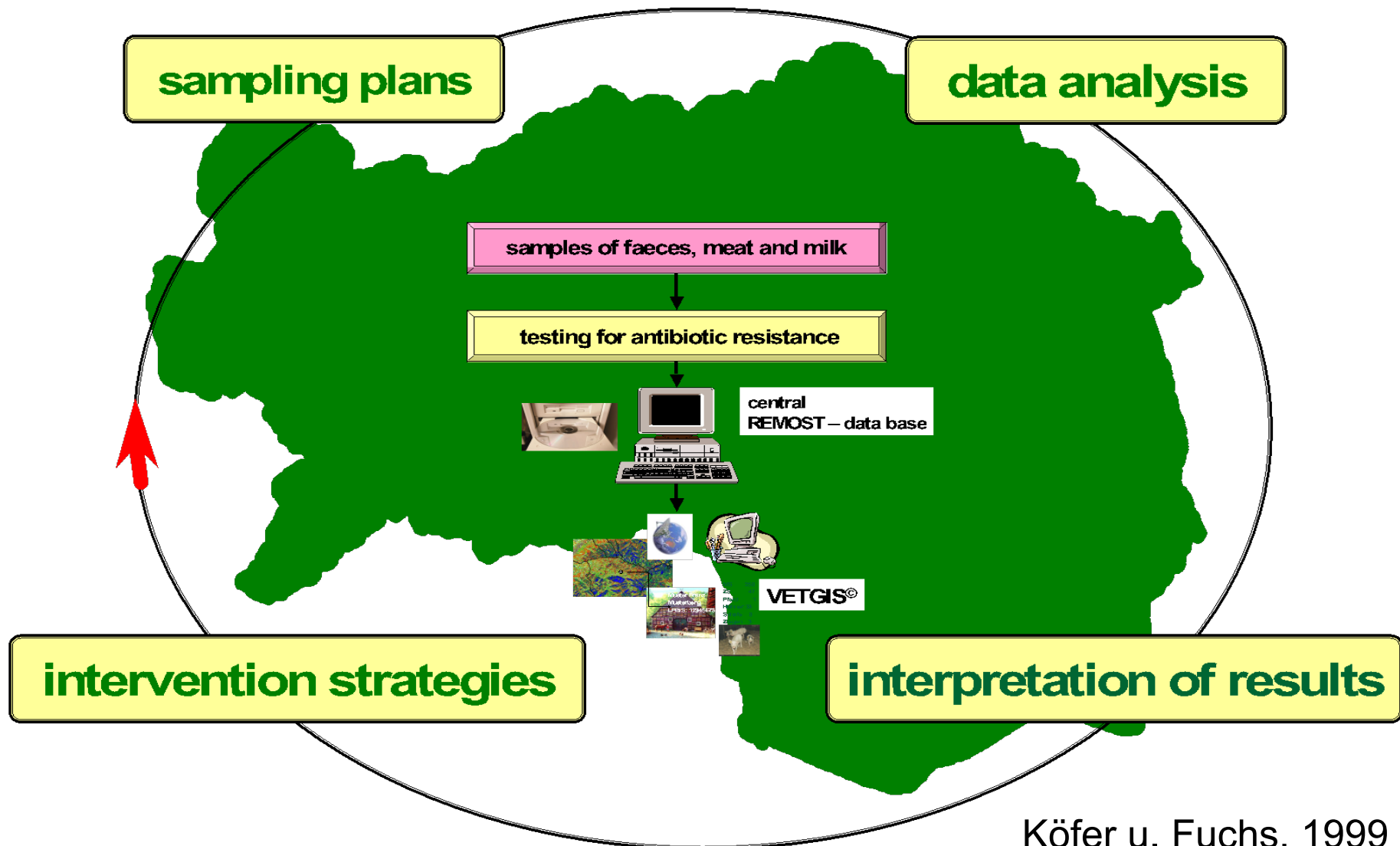
Inhalt

- Material, Methoden, Begriffe
- Glycopeptide/Streptogramine
- Gefahren aus dem Stall ?
MRSA, ESBL
- Antibiotikamengenströme
- Ausblick

AB – Einsatz / Konsequenzen



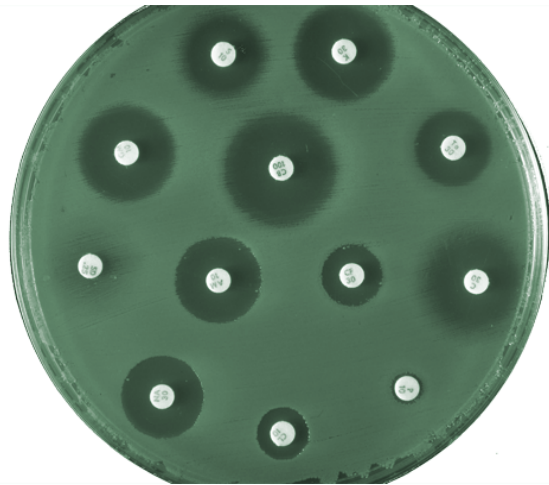
Integrated Monitoring and Control Systems in Food Production



Resistenzbestimmung

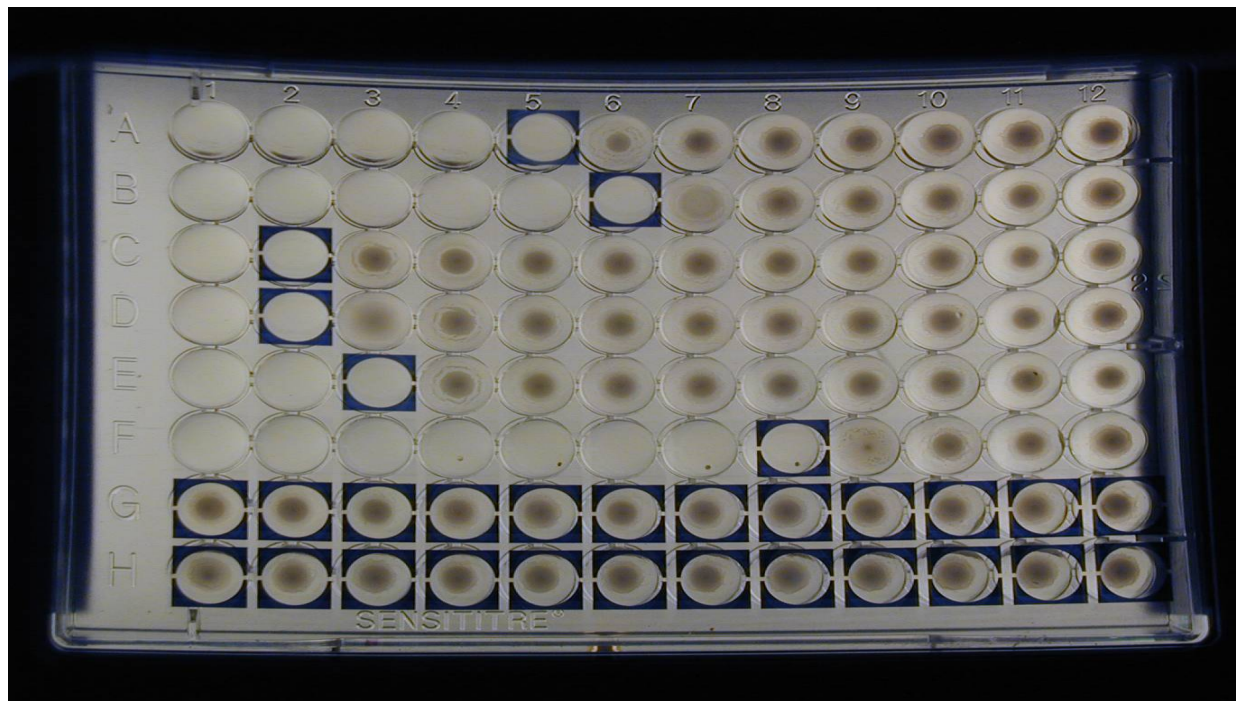
Plattendiskverfahren

semiquantitativ



Sensititre - MHK

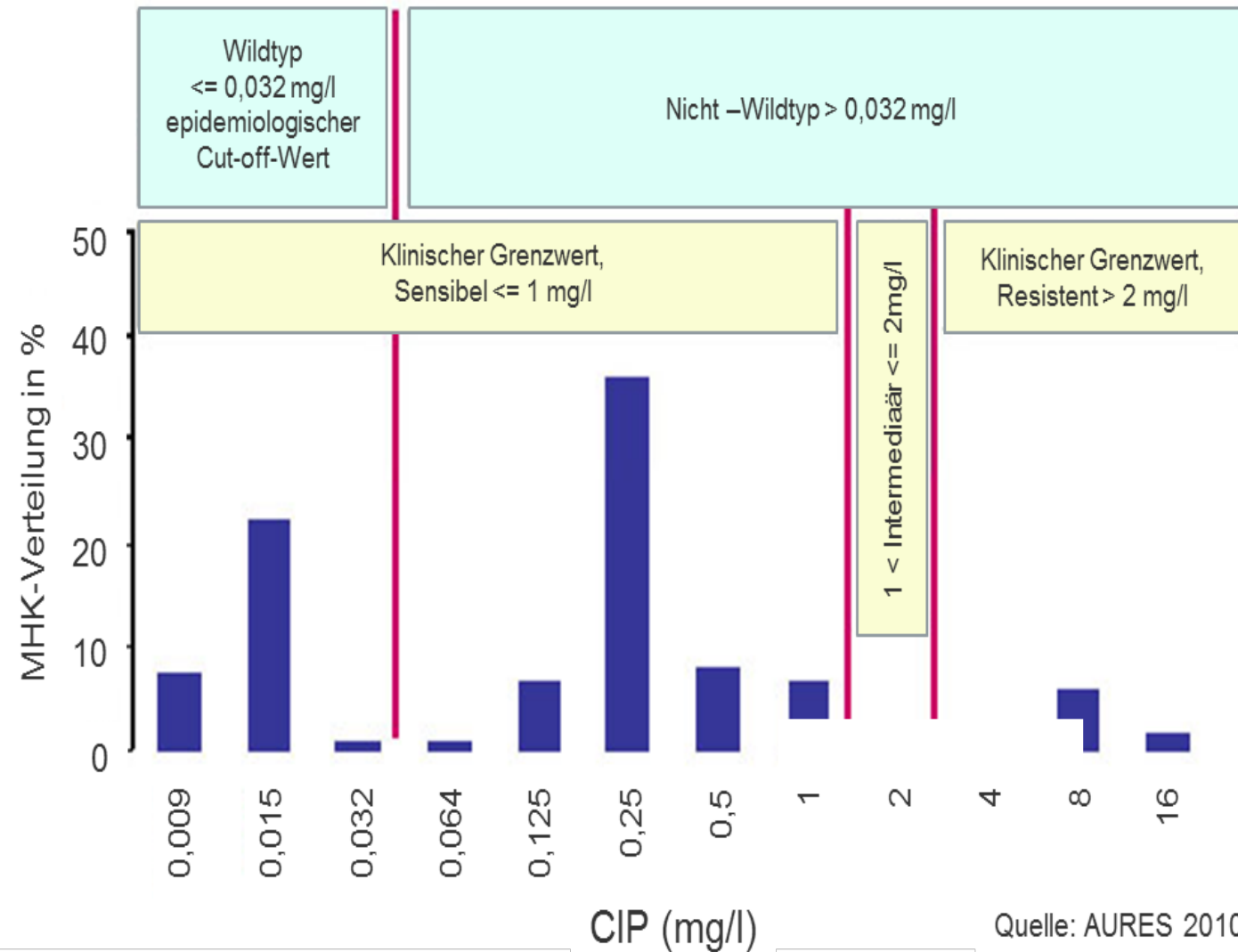
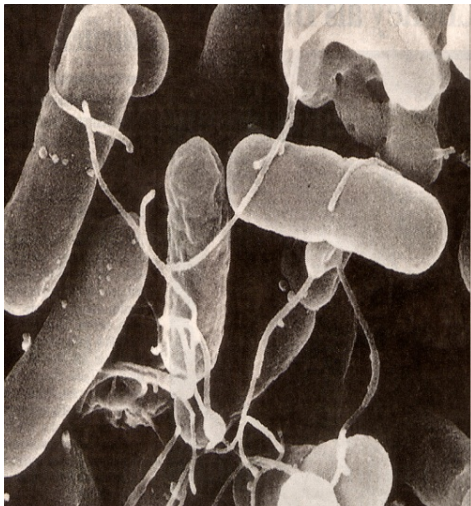
quantitativ



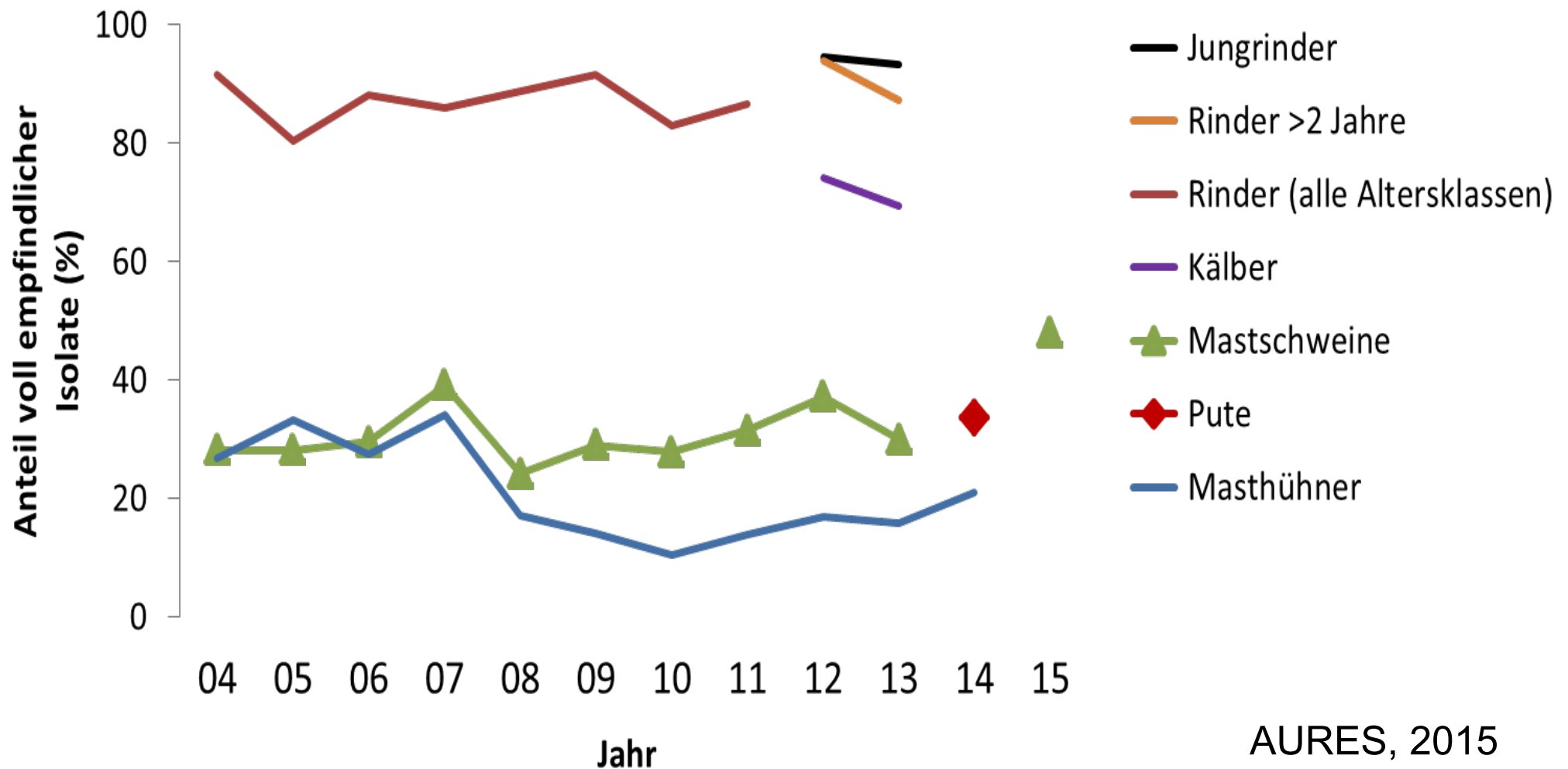
**2015 repräsentative Stichproben von Schlachtschweinen untersucht:
Indikator – *E. coli*, *Salmonella* spp., *E. coli* (AmpC, ESBL,
Carbapenemasen) sowie frisches Schweine- und Rindfleisch**

Klinischer Grenzwert nach CLSI und epidemiologischer Cut-off Wert nach EUCAST

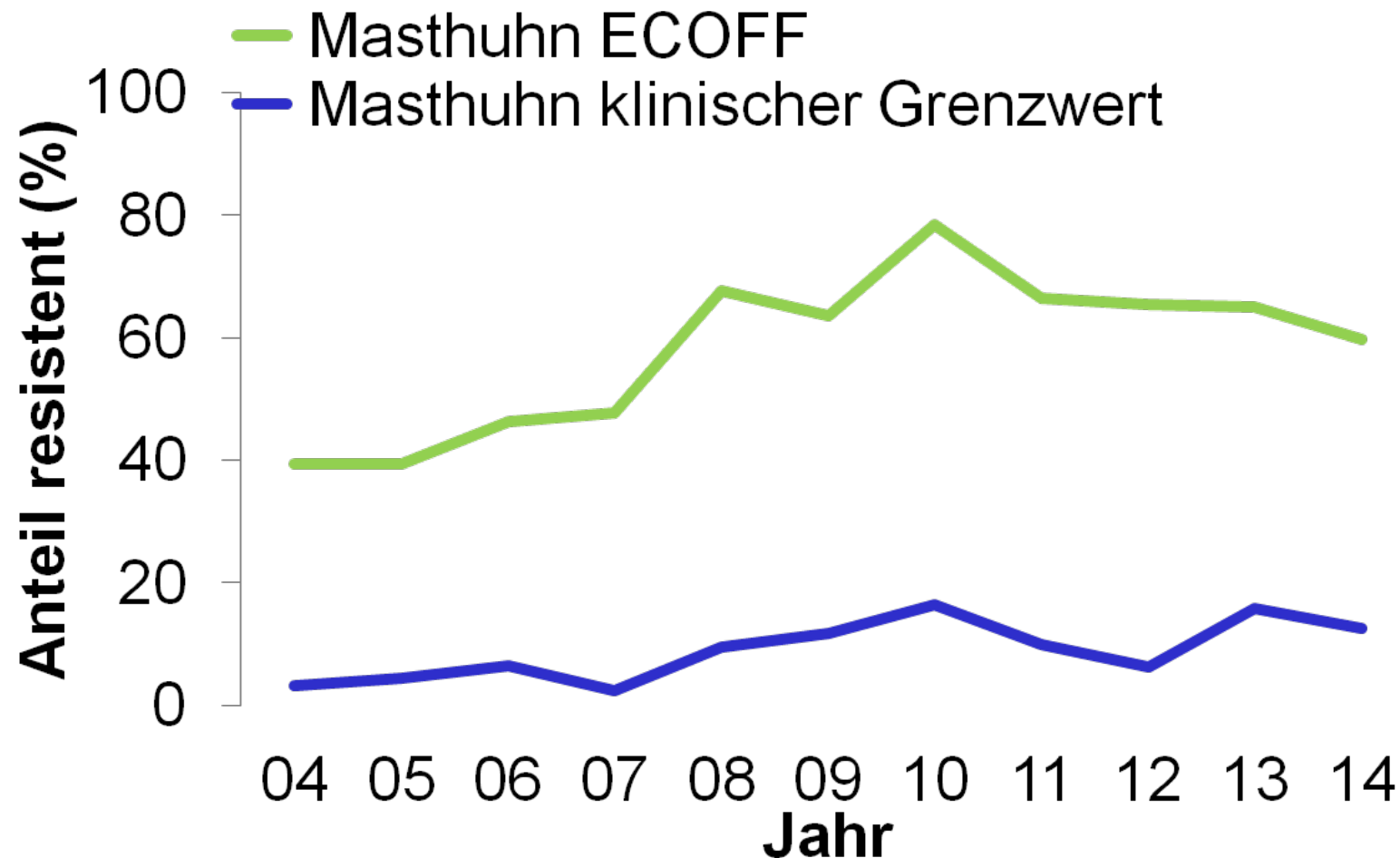
E. coli



Anteil voll empfindlicher Isolate von kommensalen *E. coli* aus verschiedenen Nutztieren, 2004–2015

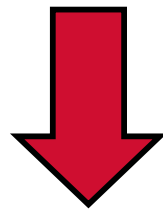


Anteil resistenter Isolate von kommensalen *E. coli* aus Masthühnern, 2004–2015, beurteilt nach ECOFF bzw. CLSI



EU – Council Resolution 1999

„A strategy against the microbial threat“



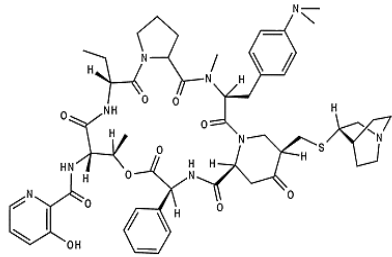
multidisciplinary and cross sectoral strategies to
control AB resistance



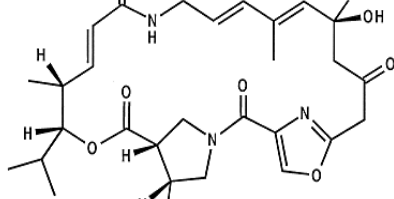
One health strategy !!!

Enterokokken - Resistenz

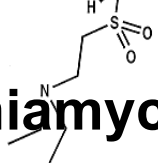
Quinupristin



Dalfopristin



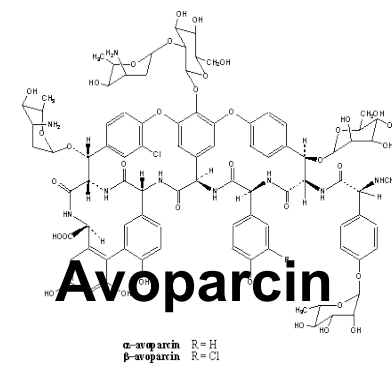
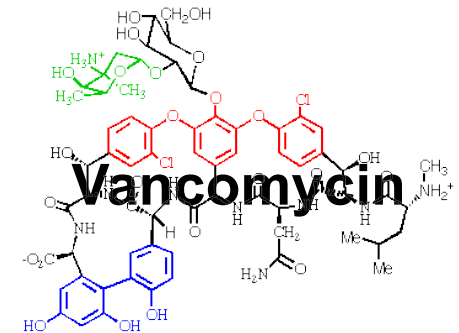
Virginia^Nmycin



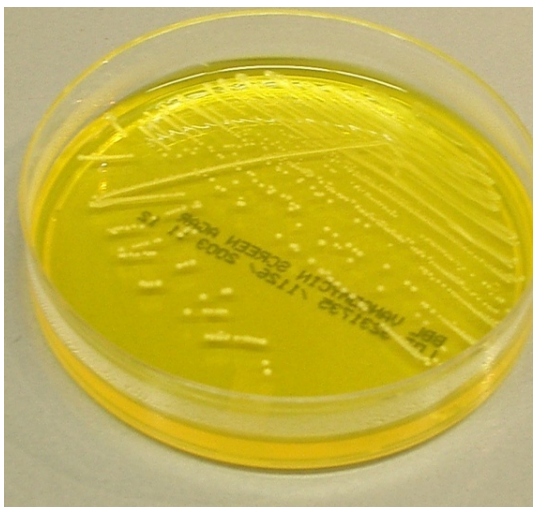
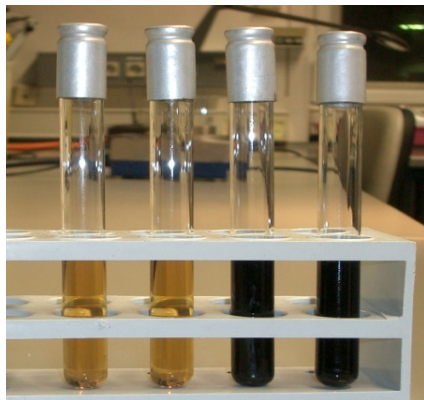
Streptogramins



Glycopeptides



Resistances of *Enterococci* to Vancomycin in livestock production



	cattle	pig	broiler
analysed faecal samples	208	206	205
faecal samples containing vancomycin-resistant enterococci	46 (22.1%)	48 (23.3%)	158 (77.1%)
faecal samples containing <i>vanA</i> (<i>E. faecium</i>)	1 (0.5%)	0	86 (42.0%)
faecal samples containing <i>vanC1</i>	9 (4.3%)	27 (13.1%)	71 (34.6%)
faecal samples containing <i>vanC2/3</i>	37 (17.8%)	22 (10.7%)	4 (2.0%)



Available online at www.sciencedirect.com

SCIENCE @ DIRECT®

Diagnostic Microbiology and Infectious Disease 53 (2005) 17–21

DIAGNOSTIC
MICROBIOLOGY
AND INFECTIOUS
DISEASE

www.elsevier.com/locate/diagmicrobio

Identification of Glycopeptide-resistant enterococci by VITEK 2 system and conventional and real-time polymerase chain reaction[☆]

Alexandra Eisner^a, Gregor Gorkiewicz^a, Gebhard Feierl^a, Eva Leitner^a, Josef Köfer^b, Harald H. Kessler^{a,*}, Egon Marth^a

^aInstitute of Hygiene, Medical University of Graz A-8010, Austria

^bDepartment of Veterinary Administration in Styria, Graz A-8010, Austria

Received 24 February 2005; accepted 13 April 2005

Abstract

Glycopeptide-resistant enterococci (GRE) are important causes of nosocomial infection. Rapid detection of GRE is essential for the implementation of appropriate control measures. In this study, we compared the reliability of 3 different methods, VITEK 2 automated system (bioMérieux), conventional polymerase chain reaction (m-PCR) protocol, and a real-time PCR protocol performed on the LightCycler (Roche) in a routine microbiology laboratory. Species identification and glycopeptide resistance were determined for 100 isolates with different glycopeptide resistance phenotypes. With the VITEK 2 system, 39% of isolates showed resistance to vancomycin. Resistance to vancomycin was detected in all isolates; however, discrepancies occurred between the different methods. Real-time PCR protocols proved to be suitable for detecting clinically relevant GRE; 90% of isolates showed resistance to vancomycin with conventional m-PCR and 100% of *vanA* and *vanB* isolates with the real-time PCR protocol. Real-time PCR assays are superior tools for identification of GRE in clinical samples.

© 2005 Elsevier Inc. All rights reserved.



Europe. Rapid detection of GRE in this study, we compared the reliability of 3 different methods, VITEK 2 automated system (bioMérieux), conventional polymerase chain reaction (m-PCR) protocol, and a real-time PCR protocol performed on the LightCycler (Roche) in a routine microbiology laboratory. Species identification and glycopeptide resistance were determined for 100 isolates with different glycopeptide resistance phenotypes. With the VITEK 2 system, 39% of isolates showed resistance to vancomycin. Resistance to vancomycin was detected in all isolates; however, discrepancies occurred between the different methods. Real-time PCR protocols proved to be suitable for detecting clinically relevant GRE; 90% of isolates showed resistance to vancomycin with conventional m-PCR and 100% of *vanA* and *vanB* isolates with the real-time PCR protocol. Real-time PCR assays are superior tools for identification of GRE in clinical samples.

APPLIED AND ENVIRONMENTAL MICROBIOLOGY, Oct. 2005, p. 6407–6409
0099-2240/05/\$08.00+0 doi:10.1128/AEM.71.10.6407–6409.2005
Copyright © 2005, American Society for Microbiology. All Rights Reserved.

Vol. 71, No. 10

High Prevalence of VanA-Type Vancomycin-Resistant Enterococci in Austrian Poultry

Alexandra Eisner,^{1,*} Gebhard Feierl,¹ Gregor Gorkiewicz,¹ Franz Dieber,² Harald H. Kessler,¹ Egon Marth,¹ and Josef Köfer²

¹Institute of Hygiene, Medical University of Graz,¹ and Department of Veterinary Administration in Styria,² A-8010 Graz, Austria

Received 23 December 2004/Accepted 10 May 2005

Fecal samples from humans and food-producing animals were analyzed for the presence of vancomycin-resistant enterococci (VRE). The VRE carriage rate in humans was 6%, and there was a predominance of VanC-type resistance. *Enterococcus faecium* with *vanA*-mediated resistance was frequent in broiler chickens (42%) but rare in cattle and pig samples.

Gefahr aus dem Stall

Antibiotikaresistenz



Gefahr aus dem Stall

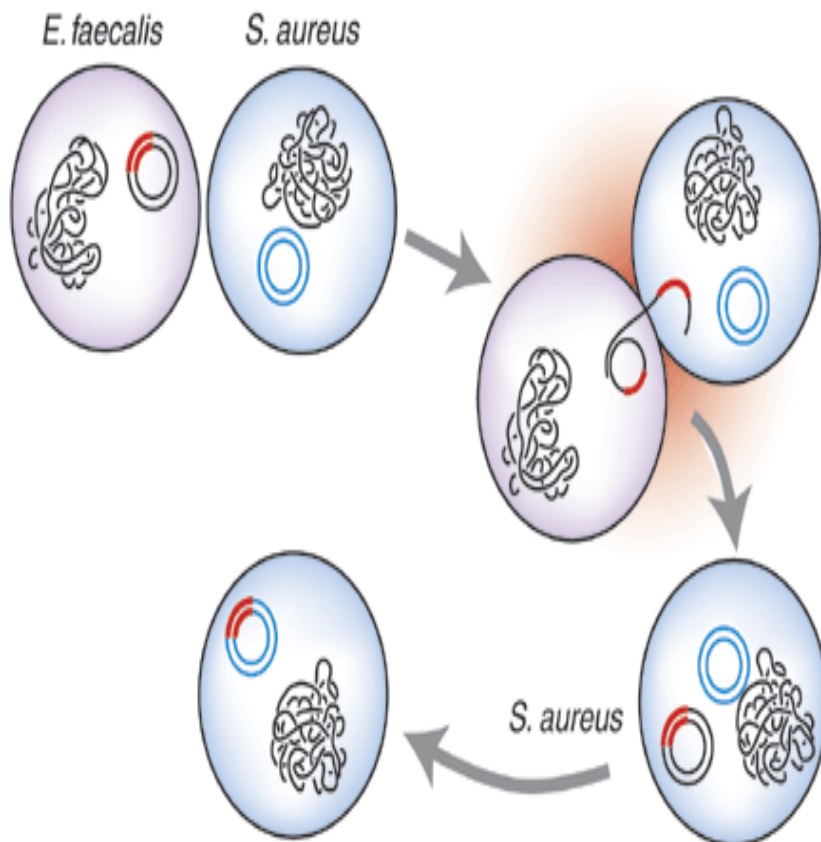
Ein gegen Antibiotika vielfach resistenter Bakterienstamm verbreitet sich unter Haustieren und befällt auch den Menschen

Von FOCUS-Korrespondentin Kerstin Schweighöfer

Den Verdacht hegte die niederländische Schweinezüchterin Ine van den Heuvel schon lange: Könnte es sein, dass sich ihre Familie im Tierstall immer wieder mit einem gefürchteten Bakterienstamm infizierte? „Wir versuchten alles, um die Keime loszuwerden“, erzählt die dreifache Mutter aus Nistelrode. „Doch wir waren innerhalb kürzester Zeit wieder befallen.“



Landwirte können sich im Stall mit gefährlichen Keimen anstecken



MRSA – EFSA Studie (2008)



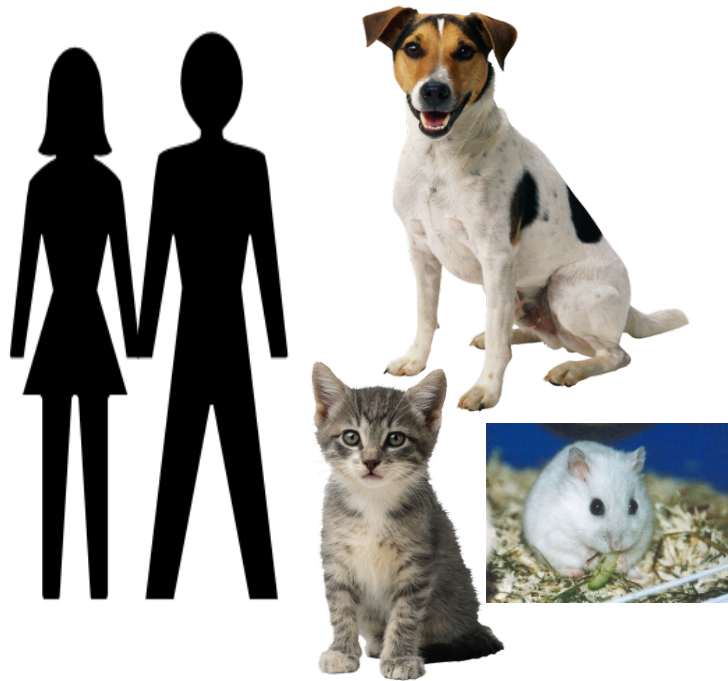
Prevalence of MRSA positive breeding holdings

MRSA & Community trades of pigs



Quelle: EFSA, 2009

■ Übertragung von MRSA ?



„These results suggest that companion animals of MRSA – infected patients can be culture-positive for MRSA, representing a potential source of infection or reinfection for humans“.

Jorge Pinto Ferreira et al., PLOSone



**Methicillin-resistant *Staphylococcus aureus* :
a new zoonotic agent ?**

Burkhard Springer, Ulrike Orendi, Peter Much et al.,
Wiener Klinische Wochenschrift, 2009;121(3-4):86-90.

MRSA in Wundabstrichen - 479 *Staph. aureus*- Isolate

Hund (201):  63 % **MRSA**

Katze (140):  46 % **MRSA**

Pferd (138):  41 % **MRSA**

Vincze et al., PLOSone

Gefahr aus dem Stall

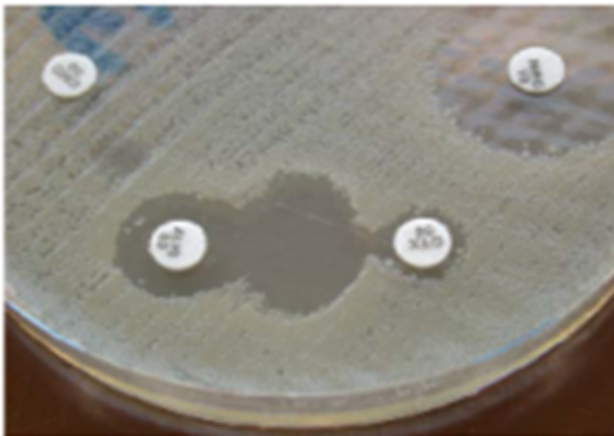
ORIGINAL ARTICLE

10.1111/j.1469-0691.2011.03497.x

Dutch patients, retail chicken meat and poultry share the same ESBL genes, plasmids and strains

M. A. Leverstein-van Hall^{1,2}, C. M. Dierikx³, J. Cohen Stuart¹, G. M. Voets¹, M. P. van den Munckhof¹,
A. van Esse¹, J. M. A. van de Sande-Bruinsma², J. Scharinga¹, M. J. M. Bonten^{1,5}
and D. J. M. de Boer¹

1) Department of
Health and the
4) SALTO, P
6) Department



Utrecht, Utrecht, 2) Centre for Infectious Disease Control, National Institute for Public
Microbiology and TSEs, Central Veterinary Institute of Wageningen UR, Lelystad,
3) Department of Health Sciences and Primary Care, University Medical Centre, Utrecht and
4) Veterinary Medicine, Utrecht University, Utrecht, the Netherlands

Charakterisierung *E. coli* – β Lactamasen

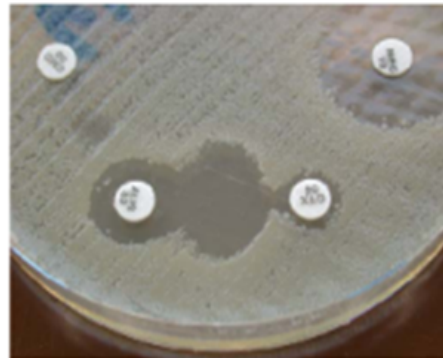


N = 177



N = 34

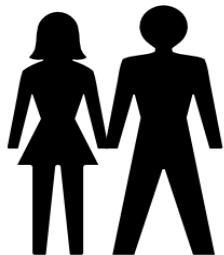
CTX-M-1



Übertragung von ESBL *E. coli* über Nahrungskette möglich

Anpassung von *E. coli* mit CTX-M-15 Genen an humanes Milieu (HWI)

Begrenztes Reservoir von *E. coli* mit CTX-M-15 ESBL bei Nutztieren



N = 40

CTX-M-9

CTX-M-15

CAVE: Selektion CTX-M-1 tragender *E. coli* durch β -Laktamantibiotika bzw. Co-Selektion durch FQ oder Tetracycline

AT – Erzeugung von Masthähnchen

Großelterntierküken (1 Tag alt)

GB

Cephalosporin



Großelterntierherden

GB

kein Einsatz



Bruteier für Elterntierherden

AT

kein Einsatz



Eintagsküken od. Jungtiere

AT

kein Einsatz



Masthähnchen

AT

kein Einsatz



**30 % ESBL *E. coli* > 50 % Resistenz
bei 3. Gen. Cephalosporinen**

Quelle: RESCH 2010, AGES

Datenerfassung von AB-Mengenströmen



Datenerfassung von AB - Mengenströmen

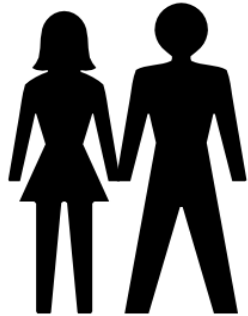
→ § 6 (3) ZoonG

Zur Absicherung der Resistenzüberwachung **hat** die Bundesministerin für Gesundheit und Frauen im Einvernehmen mit dem Bundesminister für Land- und Forstwirtschaft, Umwelt und Wasserwirtschaft

Systeme zur Überwachung von AB- Mengenströmen

durch Verordnung **festzulegen**.

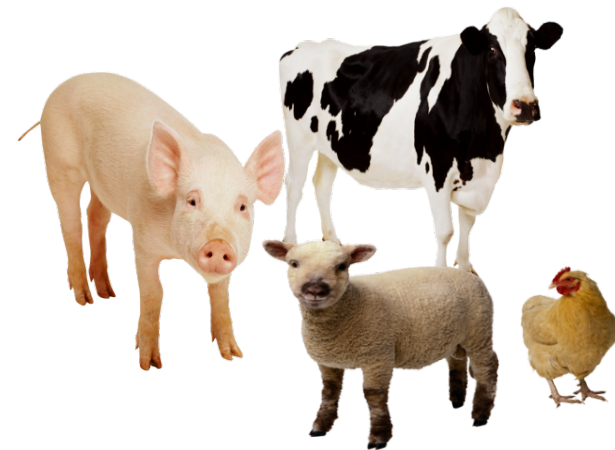
AB-Mengenströme „top – down“ 2015



8.5 Mio.

70 t AB pro 1 Mio GVE (äquivalent)

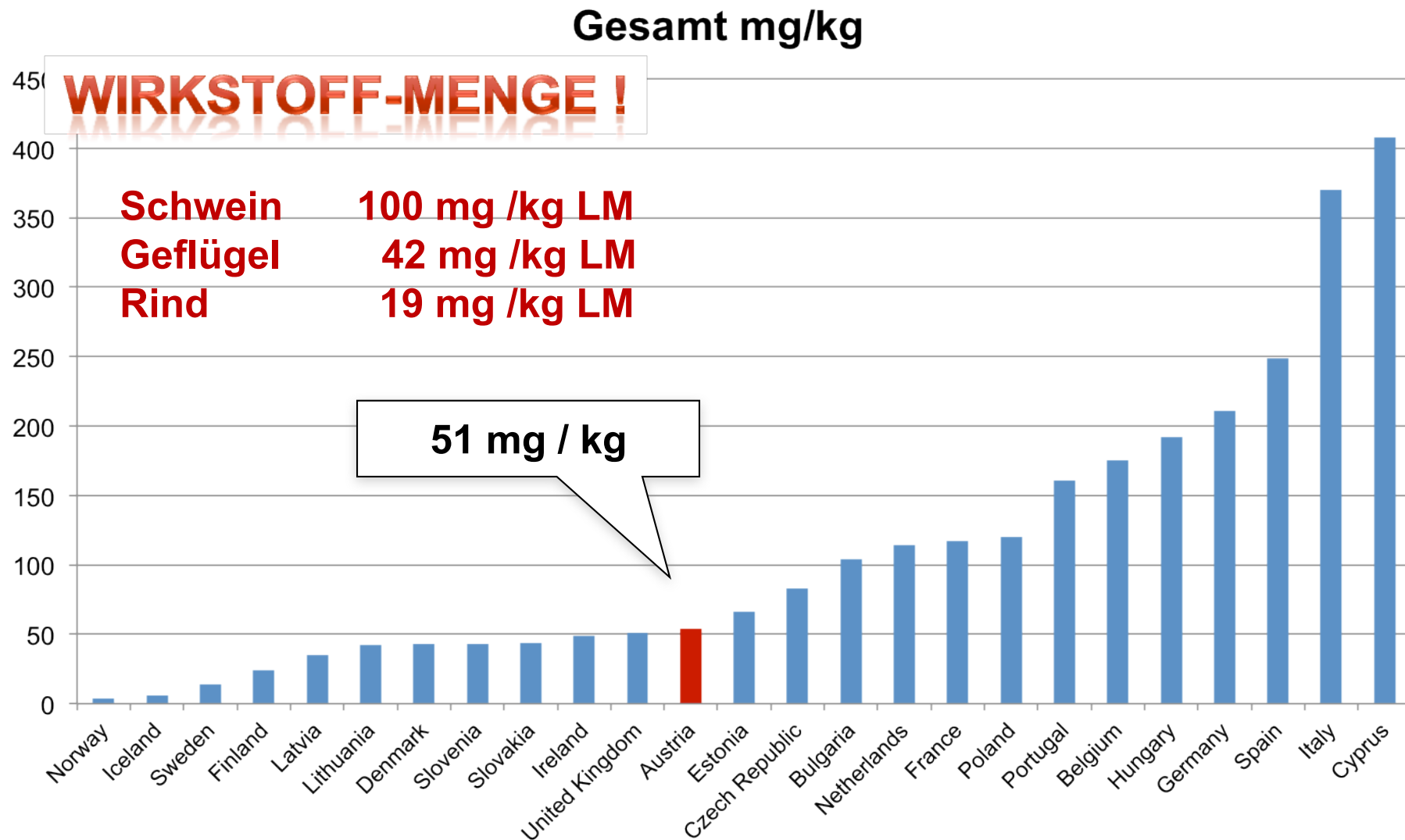
Σ 48 t AB für 2,0 Mio GVE
~ 24 t pro 1 Mio. GVE





**Antibiotika-
vertriebsmengen
sind in der
Tiermedizin
gesunken**

3rd ESVAC report 2015



„Highest Priority Critically Important Antimicrobials“ ?

- **Fluoroquinolones**
- **3rd and 4th generation Cephalosporins**
- **Macrolides**
- Glycopeptides

have been categorized as being of highest priority for risk management among antimicrobials.

- Carbapenems
- Lipopeptides
- Oxazolidinones

currently have no veterinary equivalent. WHO recommends that these classes as well as any new class of antimicrobial developed for human therapy should not be used in animals, plants, or in aquaculture.

Critically Important Antimicrobials for Human Medicine

3rd Revision 2011



HPCIA – verkaufte Wirkstoffmengen 2015

Makrolide 3,9 t

Fluorchinolone 0,5 t

**3. u. 4. Generation
Cephalosporine** 0,2 t

Summe 4,6 t



„AB“ beim Geflügel - Totalerhebung

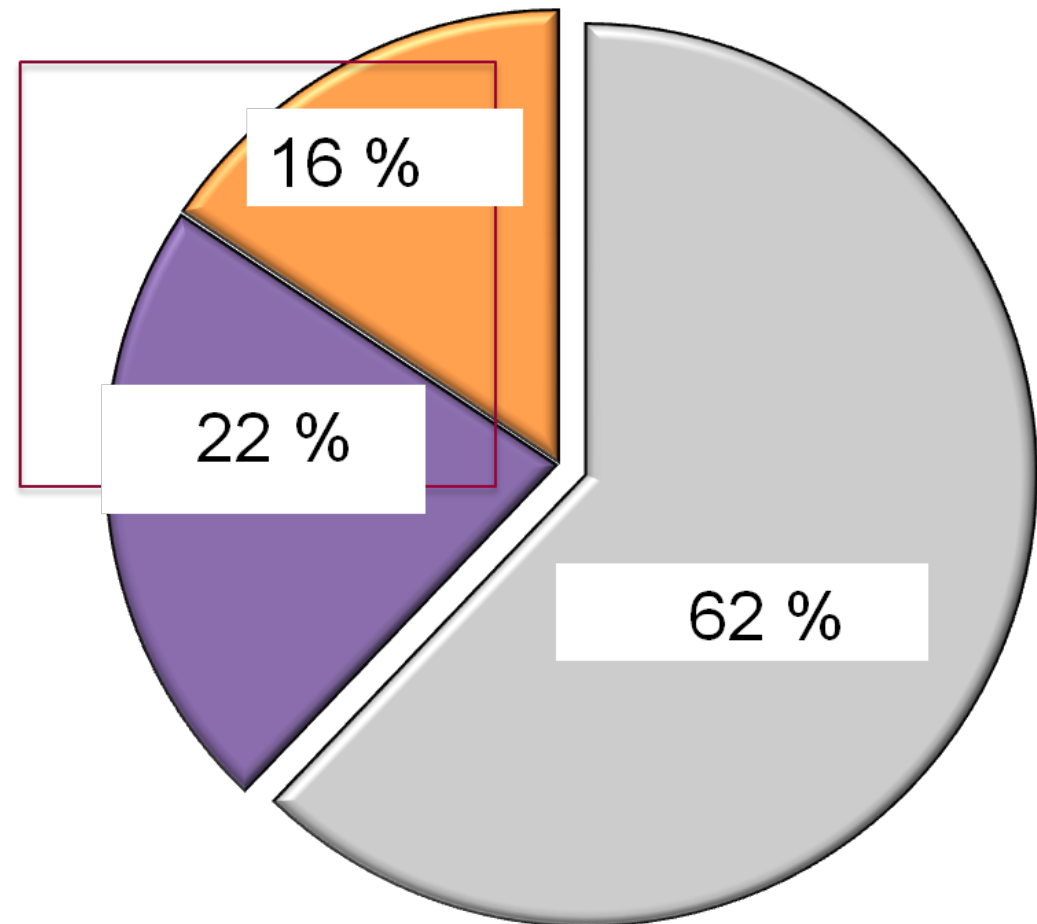


**Gesamt 42 mg/kg
LM für AT**

AURES, 2015

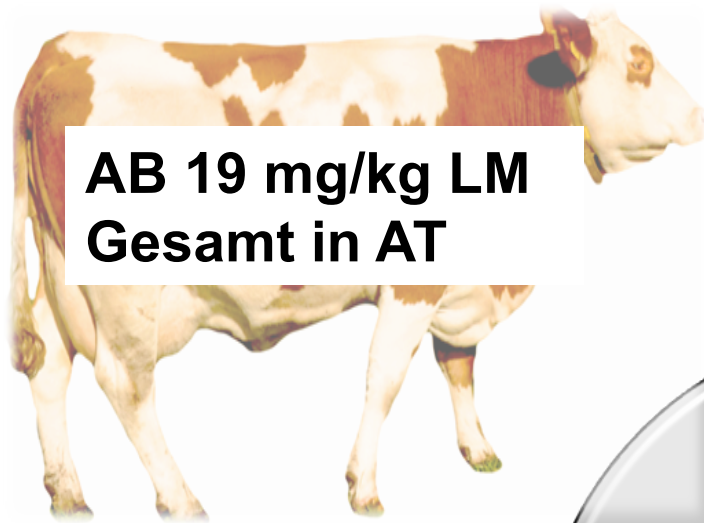
38% „HPCIA´s“

- ◆ Makrolide
- ◆ Fluorchinolone
- ◆ Andere Wirkstoffe



Obritzhauser et al., 2016

„AB“ bei Milchrindern - (Stichprobe: n = 14.858)

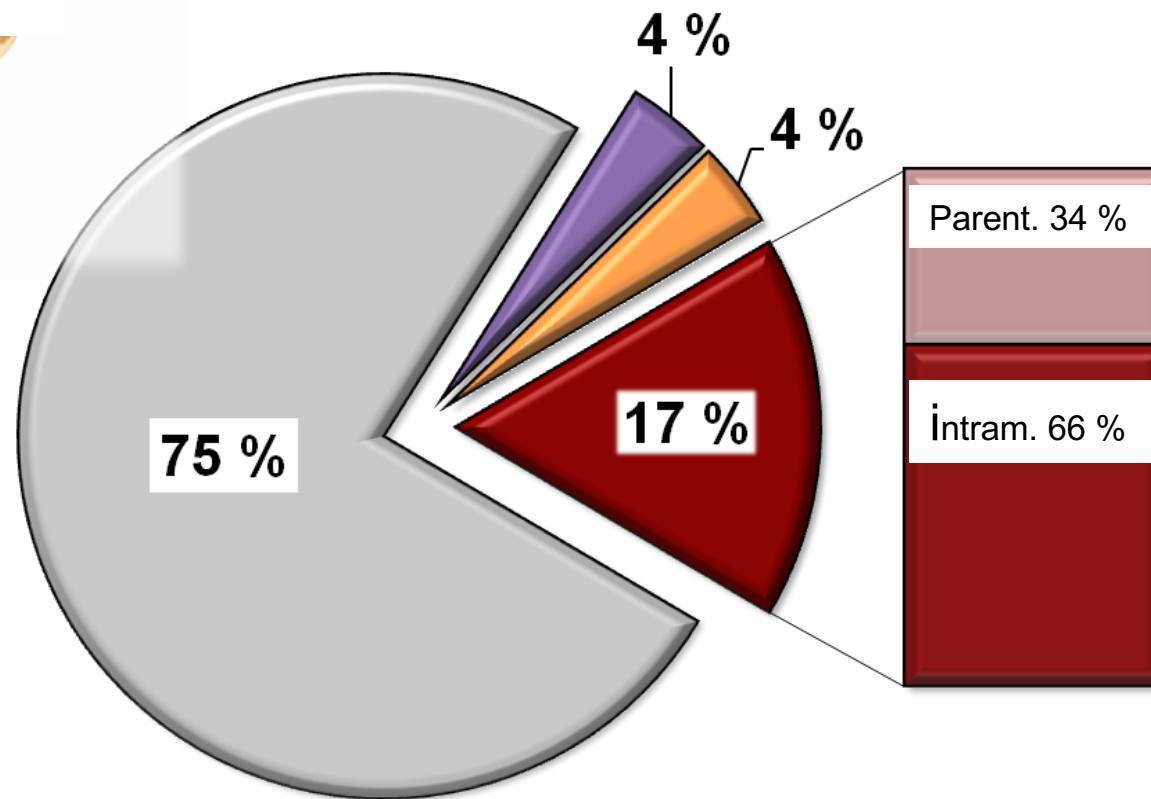


**AB 19 mg/kg LM
Gesamt in AT**

AURES, 2015

- ◆ Makrolide
- ◆ Fluorchinolone
- ◆ Cephalosporine
d. 3.u.4. Gen.
- ◆ Andere Wirkstoffe

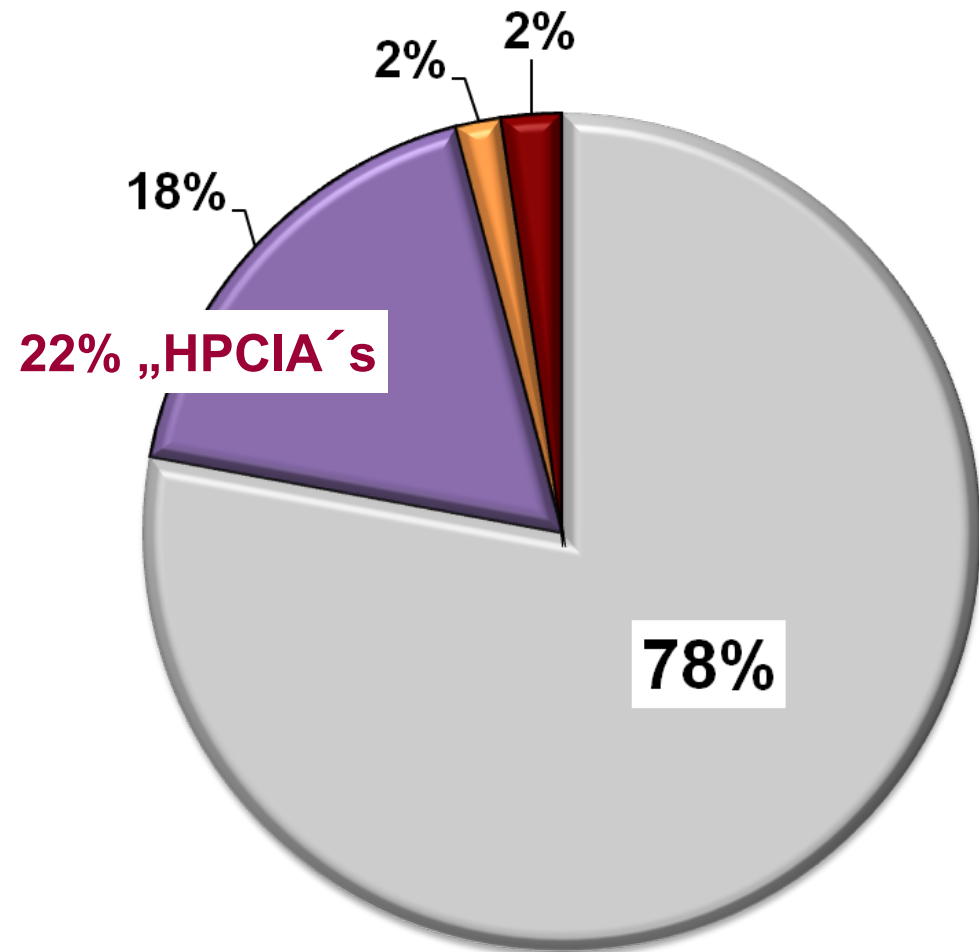
25 % „HPCIA´s“



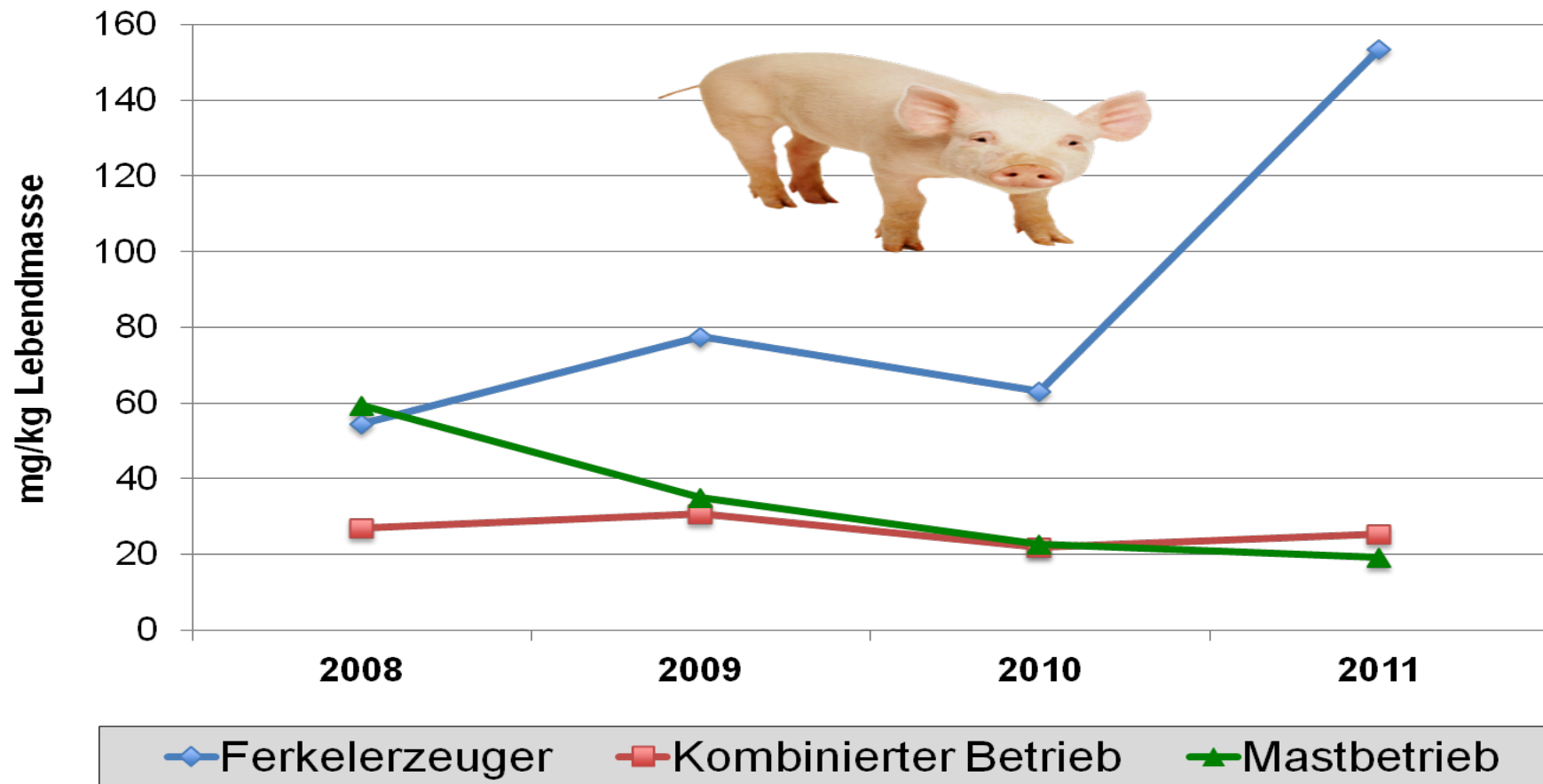
Obritzhauser, et al., 2015

„AB“ bei Schweinen (Stichprobe: n= 105.267 GVE)

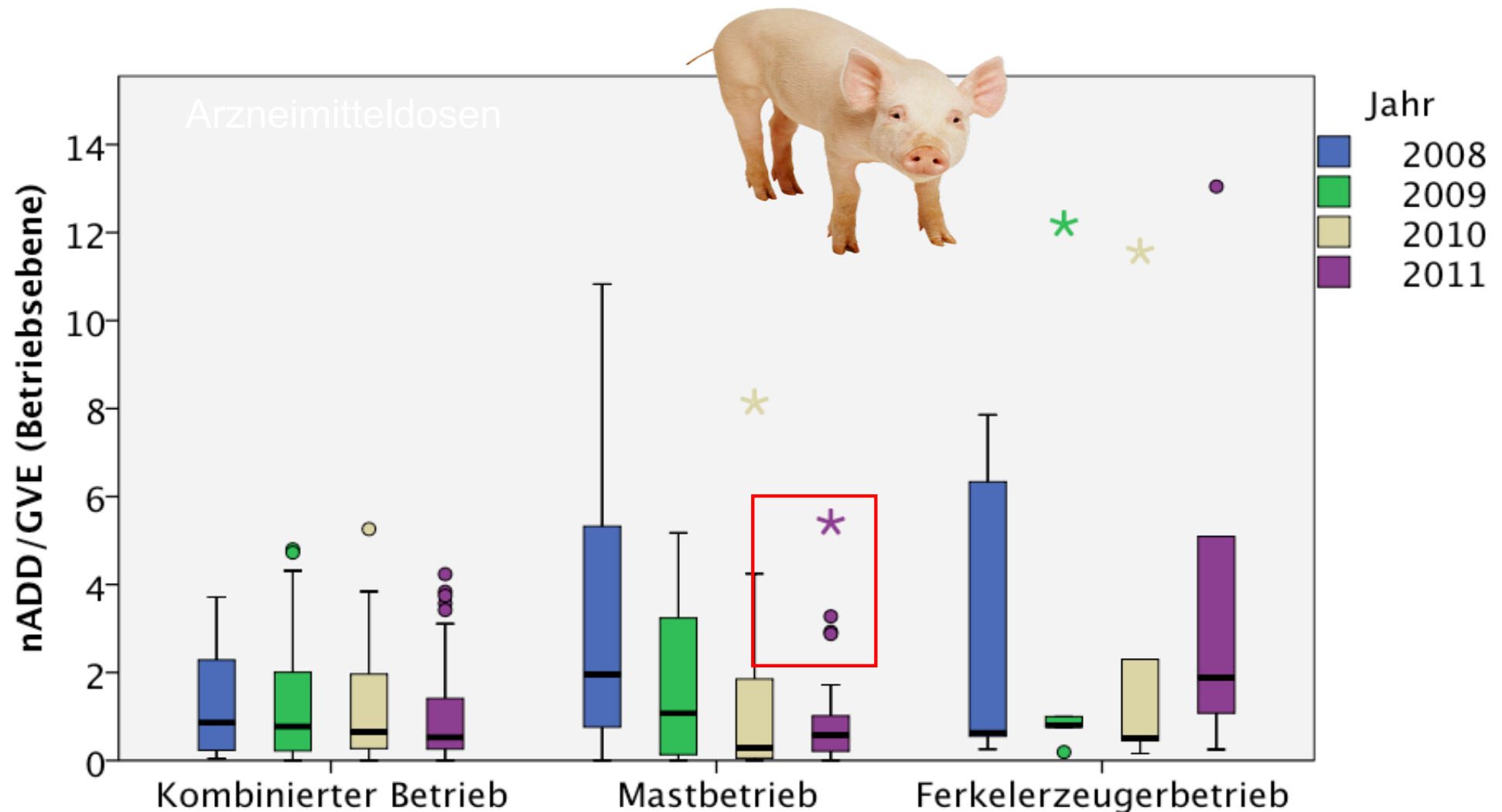
- ◆ Makrolide
- ◆ Fluorchinolone
- ◆ Cephalosporine 3. und 4. Generation
- ◆ Andere Wirkstoffe



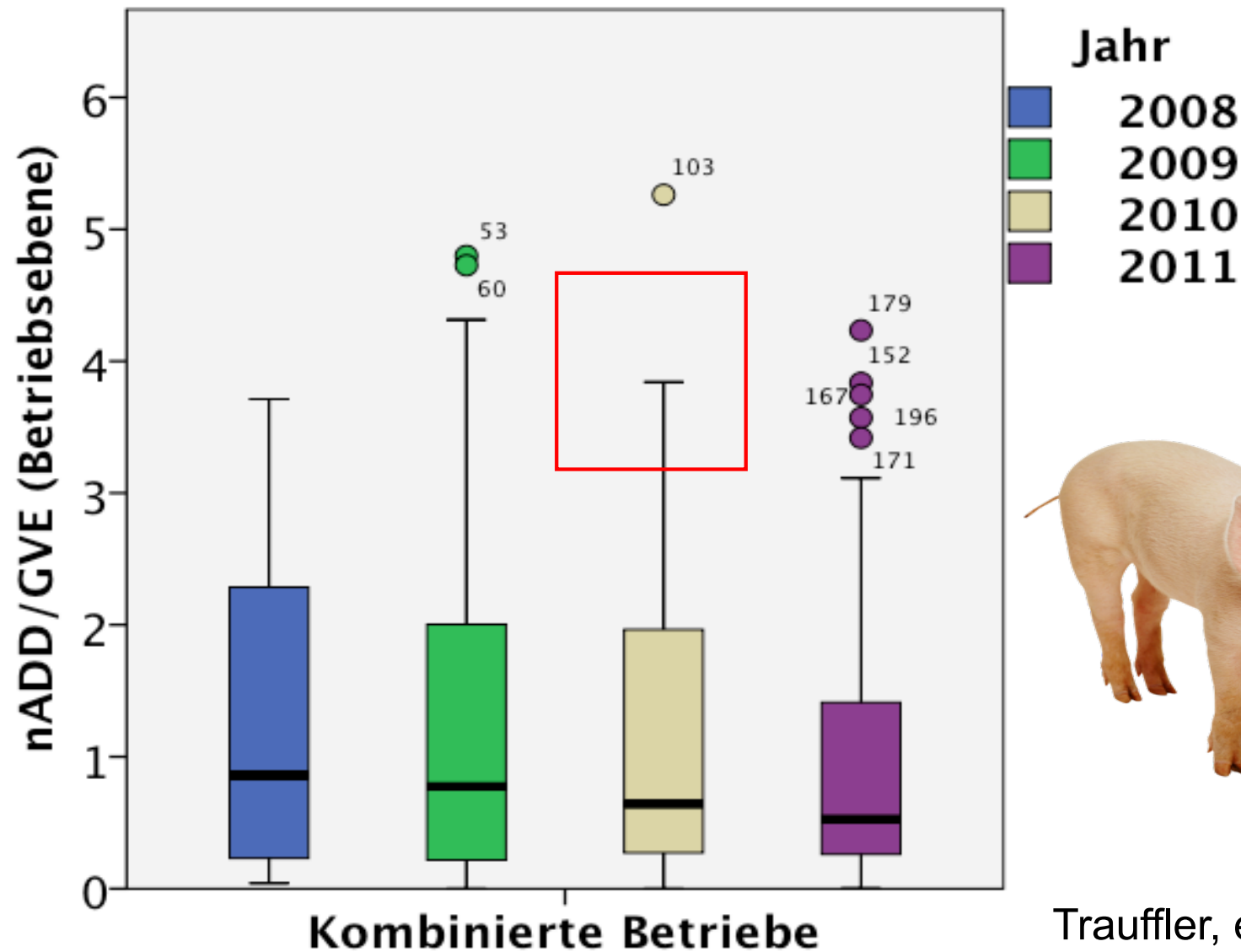
Antibiotikaeinsatz je Betriebsform in [mg/kg Lebendmasse] n= 256.916



AB-Einsatz (Arzneimitteldosen) pro Betriebsform n = 256.916 Schweine



AB – Einsatz (Dosis) – Betriebsform; n = 256.916



Trauffler, et al., 2015

AB Leitlinien



GZ: BMG-74330/0007-II/B/12/2013

Wien, am 24. Juli 2013

KUNDMACHUNG

zu § 20 Abs. 3 Tierärztegesetz – BGBl. 1975/16

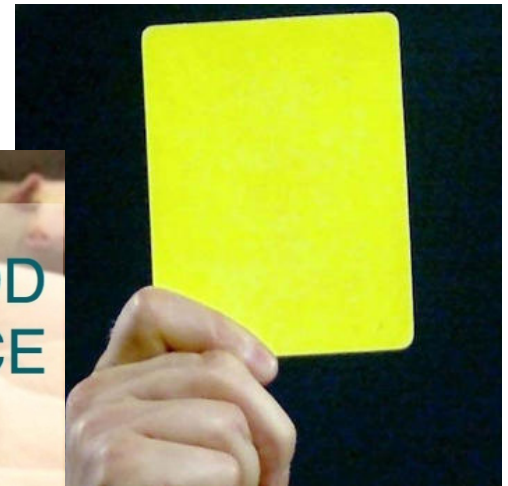
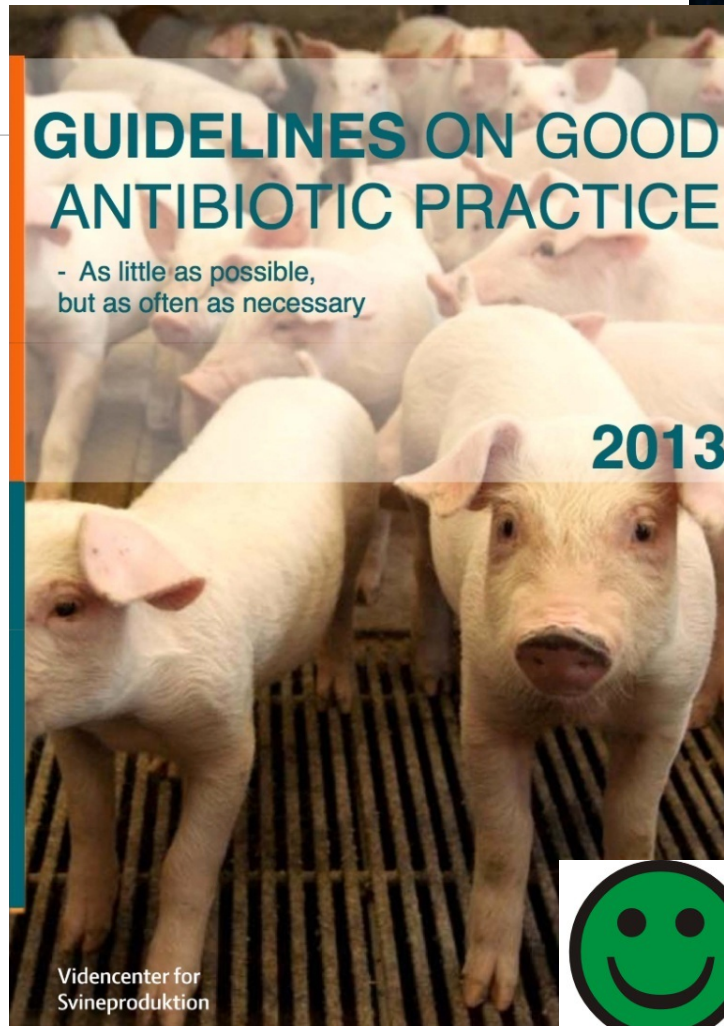
in der jeweils geltenden Fassung

**Leitlinien
für den sorgfältigen Umgang mit
antibakteriell wirksamen Tierarzneimitteln**

Diese Kundmachung tritt mit 1. September 2013 in Kraft

Für den Bundesminister

Hon.-Prof. Dr. Aigner



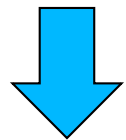
Ausblick

 AB – Rückgang 2010 – 2015 um 22 %

 QGV - „Poultry health data“

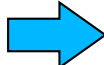
 X – Projekt Schwein

 ADDA Projekt - Rind (Elektronisches Stallbuch)



AB Reduktion um 30 % incl. HPCIA

„a must“  „bottom up“ Erfassung (n ADD/Jahr/Kategorie)

„a must“  „TGD – Alternativen“

Weiterführende aktuelle Informationen





Vielen Dank für Ihre Aufmerksamkeit!

Institut für Öffentliches Veterinärwesen
Veterinärmedizinische Universität Wien

Institut für Öffentliches Veterinärwesen
Veterinärmedizinische Universität Wien