

**CONVEGNO «ANTIBIOTICORESISTENZA: UN  
APPROCCIO GLOBALE PER UNA SFIDA MONDIALE»**

***Ozzano Emilia (BO)***

***Aula Messieri - 28 aprile 2017***

***In ricordo di Roberto Mattioli***

# **Monitoraggio AMR nel cane e nel gatto**



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Dati EURISPES Rapporto Italia 2016:

43,33% degli italiani detiene almeno un pet

In Italia circa 60 milioni di animali da compagnia



• **7,5 milioni di gatti**



• **7 milioni di cani**

- 30 milioni di pesci
- 13 milioni di uccelli

- 2 milioni di roditori
  - 1,5 milioni di rettili
-



**Uso degli antimicrobici nei pets**

- MRSA**
- MRSP**
- Enterococchi**
- Enterobacteriaceae**
- Pseudomonas*
- Acinetobacter*
- Campylobacter*
- Clostridium difficile*

Sviluppo e diffusione di batteri resistenti nei pets

batteri pet a pet

Limitazione nelle opzioni terapeutiche di infezioni batteriche nei pets

batteri uomo a pet

**Scambio di batteri resistenti tra pets e uomo**

Limitazione nelle opzioni terapeutiche di infezioni batteriche nell'uomo

Sviluppo e diffusione di batteri resistenti nell'uomo

**Uso degli antimicrobici nell'uomo**

Trasferimento di geni di resistenza tra batteri dei pets

Trasferimento di geni di resistenza tra batteri dei pets e batteri dell'uomo

Trasferimento di geni di resistenza tra batteri dell'uomo

Passaggio batteri resistenti da pet a uomo

Passaggio batteri resistenti da uomo a uomo

Adattato da Guardabassi *et al.*, 2004

## CONSUMO ANTIBIOTICI PETS

ESVAC BY EMA 2010

CASCADE / OFF-LABEL

RICETTA INFORMATIZZATA

VETCOMPASS/SMALL ANIMALS VETERINARY SURVEILLANCE NETWORK SAVSNet

## TEST SENSIBILITA' ANTIBIOTICI SU ISOLATI DA CASI CLINICI

METODOLOGIE STANDARDIZZATE

VetCAST (Veterinary Committee on Antimicrobial Susceptibility Testing)

<5% TERAPIE MIRATE IN TEACHING HOSPITAL –  
ESCHER ET AL. 2011

MECCANISMI MOLECOLARI RESISTENZA

## MONITORAGGIO AMR PICCOLI ANIMALI

network veterinario europeo  
di sorveglianza simile a  
quanto già esiste in umana  
(European Antimicrobial  
Resistance Surveillance  
Network (EARS-Net))

## STRATEGIE DI GESTIONE E LINEE GUIDA

ANCHE A LIVELLO OSPEDALIERO

## IMPATTO IN SANITA' PUBBLICA

AUMENTO CONOSCENZE/INFORMAZIONE

RESEARCH ARTICLE

Open Access

## European multicenter study on antimicrobial resistance in bacteria isolated from companion animal urinary tract infections

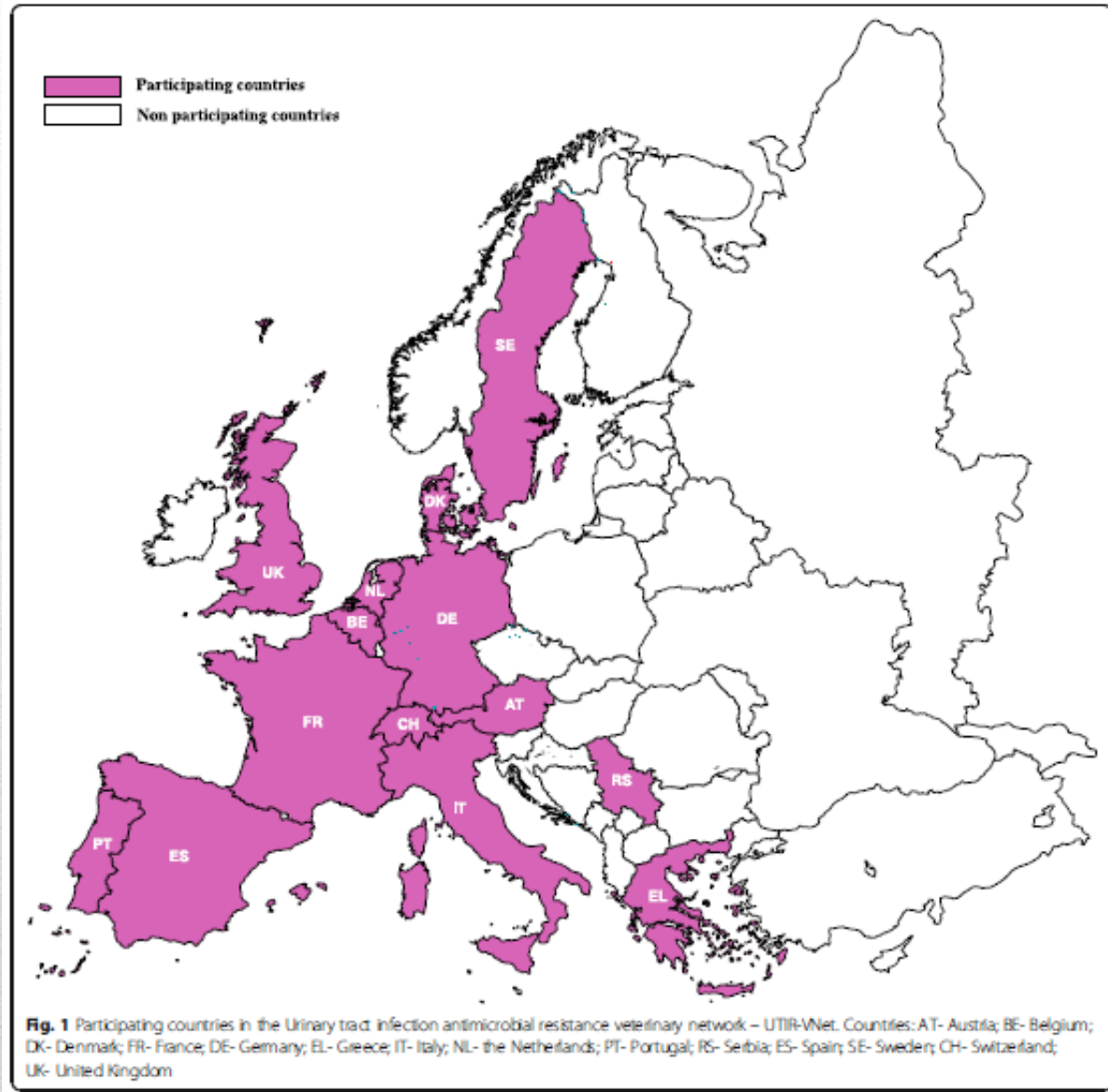


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- ✓ Studio multicentrico retrospettivo realizzato sotto l'ombrello della European Society of Veterinary Nephrology and Urology
  - ✓ Costituzione di un Urinary Tract Infection Resistance-Veterinary Network (UTIR-VNet)
  - ✓ Obiettivo: determinare la frequenza di uropatogeni nel cane e nel gatto e valutare l'occorrenza e l'andamento temporale dell'antibioticoresistenza in un periodo di 6 anni (2008 – 2013)
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## ...UNO STUDIO MULTICENTRICO EUROPEO SU AMR IN PETS

- ✓ 14 paesi europei (16 labs)
- ✓ quasi tutti laboratori universitari di microbiologia clinica veterinaria
- ✓ considerati 22256 isolati in corso di UTI (15097 da cani e 5963 da gatti)



- ✓ Metodi di identificazione batteri: sistemi API, PCR e MALDI-TOF
  - ✓ 15 antibiotici testati su tutti gli isolati mediante disk diffusion method (CLSI) tranne Svezia (Vet MIC, SVA), Danimarca e UK (COMPAN1F Sensititre Panels, Thermo Fisher), Svizzera e Belgio (VITEK 2 BioMerieux)
  - ✓ Sono stati inclusi i seguenti antimicrobici: amoxicillin clavulanate (AMC), ampicillin (AMP), cefotaxime (CTX), ceftiofur (CVN), ceftazidime (CAZ), cefepime (CEP), cefepime (CPD), ceftiofur (EFT), ciprofloxacin (CIP), enrofloxacin (ENR), gentamicin (CN), marbofloxacin (MAR), oxacillin (OX), penicillin (P) and trimethoprim/sulfamethoxazole (SXT)
  - ✓ Utilizzati i breakpoint stabiliti per l'uomo da CLSI (M100-S25, 2015)
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# FREQUENZA DI UROPATOGENI NEL CANE E NEL GATTO

**Table 2** Uropathogenic bacteria aetiology, single versus mixed infections and cat versus dog as host species

Organism	Overall		Single organism		Mixed infections		P	Dogs		Cats		P
	n	% (95 % CI) <sup>a</sup>	n	% (95 % CI) <sup>a</sup>	n	% (95 % CI) <sup>a</sup>		n	% (95 % CI) <sup>a</sup>	n	% (95 % CI) <sup>a</sup>	
<i>Enterobacter</i> spp.	308	1.38 (1.23–1.54)	244	1.22 (1.07–1.38)	64	2.75 (2.09–3.42)	<b>&lt;0.0001</b>	194	1.21 (1.04–1.38)	114	1.81 (1.48–2.14)	<b>0.0008</b>
<i>Enterococcus</i> spp.	1506	6.77 (6.44–7.10)	1129	5.66 (5.34–5.99)	377	16.22 (14.72–17.72)	<b>&lt;0.0001</b>	745	4.66 (4.34–4.99)	761	12.11 (11.31–12.92)	<b>&lt;0.0001</b>
<i>Escherichia coli</i>	13231	59.45 (58.80–60.09)	12417	62.30 (61.62–62.97)	814	35.03 (33.09–36.97)	<b>&lt;0.0001</b>	9506	59.51 (58.75–60.27)	3725	59.30 (58.08–60.51)	0.7832
<i>Klebsiella</i> spp.	478	2.15 (1.96–2.34)	400	2.01 (1.81–2.20)	78	3.36 (2.62–4.09)	<b>&lt;0.0001</b>	385	2.41 (2.17–2.65)	93	1.48 (1.18–1.78)	<b>&lt;0.0001</b>
<i>Proteus</i> spp.	1992	8.95 (8.58–9.33)	1770	8.88 (8.49–9.28)	222	9.55 (8.36–10.75)	0.2824	1869	11.70 (1.20–1.22)	123	1.96 (1.62–2.30)	<b>&lt;0.0001</b>
<i>Pseudomonas</i> spp.	389	1.75 (1.58–1.92)	315	1.58 (1.41–1.75)	74	3.18 (2.47–3.90)	<b>&lt;0.0001</b>	293	1.83 (1.63–2.04)	96	1.53 (1.22–1.83)	0.1249
<i>Staphylococcus</i> spp.	2893	13.00 (12.56–13.44)	2519	12.64 (12.18–13.10)	374	16.09 (14.60–17.59)	<b>&lt;0.0001</b>	1836	11.49 (11.00–11.99)	1057	16.83 (15.90–17.75)	<b>&lt;0.0001</b>
<i>Streptococcus</i> spp.	802	3.60 (3.36–3.85)	586	2.94 (2.71–3.17)	216	9.29 (8.11–10.47)	<b>&lt;0.0001</b>	675	4.23 (3.91–4.54)	127	2.02 (1.67–2.37)	<b>&lt;0.0001</b>
Other	657	2.95 (2.73–3.17)	552	2.77 (2.54–3.00)	105	4.52 (3.67–5.36)	-	471	2.95 (2.69–3.21)	186	2.96 (2.54–3.38)	-

<sup>a</sup>95 % CI, 95 % Confidence interval

n – Total number of isolates

P- P value obtained by Fisher exact test when comparing single versus mixed infections and cat versus dog as host. Statistically significant values are highlighted in bold



# OCCORRENZA E ANDAMENTO TEMPORALE AMR

Amoxicillin clavulanate

Third-generation cephalosporins

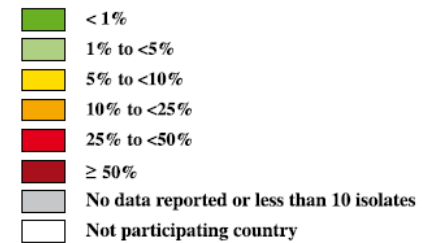
Trimethoprim-sulhamethoxazole



Fluoroquinolones

Gentamicin

Multidrug-resistant



**% AMR *Escherichia coli* 2012-2013**

# OCCORRENZA E ANDAMENTO TEMPORALE AMR

**Table 3** Percentage of resistance in *Escherichia coli* by antimicrobial and country in 2012–2013

Country <sup>a</sup>	AMC		3GC		FLU		CN		SXT		Combined resistance		
	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% MDR (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	% FullS (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>
DK	206	2.88 (0.61–5.16) [c]	208	4.33 (1.41–7.09) [a, c]	208	2.88 (0.61–5.16) [c]	208	1.92 (0.06–3.79) [a, c]	208	8.17 (4.45–11.90) [a, c]	208	2.88 (0.61–5.16) [b]	88.94 (84.68–93.20) [b, c]
IT	69	26.09 (15.73–36.45) [e, f]	69	24.64 (14.47–34.80) [e]	69	31.88 (20.89–42.88) [e]	69	14.49 (6.19–22.80) [e]	69	28.99 (18.28–39.69) [e, f]	69	28.99 (18.28–39.69) [d]	63.77 (52.43–75.11) [d, e, f]

**% AMR *Escherichia coli* 2012–2013**

**Table 7** Temporal trends of antimicrobial resistance in *Escherichia coli* by country

Country <sup>a</sup> (Years)	AMC OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	3GC OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	FLU OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	CN OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	SXT OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	MDR OR <sup>b</sup> (95 % CI) <sup>c</sup> P value	FullS OR <sup>b</sup> (95 % CI) <sup>c</sup> P value
BE (2010–13)	<b>0.787</b> <b>(0.646–0.960)</b> <b>0.0180</b>	- - -	<b>0.749</b> <b>(0.635–0.882)</b> <b>0.0006</b>	<b>0.677</b> <b>(0.507–0.904)</b> <b>0.0081</b>	<b>0.796</b> <b>(0.695–0.912)</b> <b>0.0010</b>	<b>0.529<sup>d</sup></b> <b>(0.393–0.712)</b> <b>&lt;0.0001</b>	<b>1.275<sup>d</sup></b> <b>(1.127–1.442)</b> <b>0.0001</b>
DK (2008–13)	<b>0.698</b> <b>(0.500–0.976)</b> <b>0.0357</b>	0.869 (0.646–1.169) 0.3533	<b>0.742</b> <b>(0.565–0.976)</b> <b>0.0325</b>	0.926 (0.620–1.384) 0.7086	<b>0.793</b> <b>(0.642–0.980)</b> <b>0.0316</b>	0.874 (0.615–1.242) 0.4528	<b>1.396</b> <b>(1.156–1.684)</b> <b>0.0005</b>
FR (2010–13)	0.885 (0.780–1.005) 0.0606	<b>0.859</b> <b>(0.749–0.987)</b> <b>0.0314</b>	<b>0.822</b> <b>(0.727–0.928)</b> <b>0.0016</b>	0.938 (0.734–1.200) 0.6121	0.960 (0.853–1.080) 0.4997	0.901 (0.782–1.037) 0.1448	<b>1.112</b> <b>(1.002–1.233)</b> <b>0.0456</b>
DE (2009–13)	1.029 (0.779–1.358) 0.8424	1.076 (0.805–1.438) 0.6211	1.185 (0.912–1.540) 0.2037	0.856 (0.520–1.409) 0.5397	1.040 (0.831–1.302) 0.7317	1.111 (0.801–1.541) 0.5295	0.941 (0.780–1.136) 0.5281
EL <sup>e</sup> (2009–13)	1.534 (0.851–2.766) 0.1545	1.083 (0.586–2.003) 0.7992	0.924 (0.630–1.355) 0.6855	- - -	0.880 (0.596–1.301) 0.5229	- - -	- - -
IT (2009–2013)	1.175 (0.844–1.637) 0.3391	1.017 (0.749–1.383) 0.9127	0.828 (0.629–1.090) 0.1784	1.007 (0.700–1.449) 0.9686	0.769 (0.582–1.016) 0.0645	1.065 (0.761–1.490) 0.7147	1.248 (0.953–1.634) 0.1076
NL (2008–13)	<b>1.108</b> <b>(1.026–1.197)</b> <b>0.0088</b>	<b>0.465</b> <b>(0.402–0.539)</b> <b>&lt;0.0001</b>	<b>0.916</b> <b>(0.841–0.999)</b> <b>0.0464</b>	0.682 (0.327–1.422) 0.3071	<b>0.917</b> <b>(0.859–0.978)</b> <b>0.0083</b>	<b>0.380<sup>d</sup></b> <b>(0.320–0.450)</b> <b>&lt;0.0001</b>	<b>1.648<sup>d</sup></b> <b>(1.494–1.818)</b> <b>&lt;0.0001</b>
PT (2008–13)	1.139 (0.913–1.419) 0.2482	1.187 (0.945–1.492) 0.1411	1.029 (0.823–1.287) 0.8029	1.222 (0.899–1.660) 0.2010	1.087 (0.867–1.364) 0.4680	1.156 (0.898–1.488) 0.2601	0.797 (0.629–1.010) 0.0608
ES (2010–13)	1.372 (0.855–2.201) 0.1899	1.551 (0.857–2.808) 0.1474	0.801 (0.529–1.214) 0.2965	0.859 (0.467–1.578) 0.6238	0.752 (0.489–1.156) 0.1939	1.237 (0.677–2.258) 0.4897	0.944 (0.570–1.564) 0.8234
SE (2008–13)	0.976 (0.915–1.041) 0.4569	- - -	0.980 (0.827–1.147) 0.8018	<b>0.700</b> <b>(0.562–0.872)</b> <b>0.0015</b>	0.961 (0.892–1.037) 0.3059	<b>0.697</b> <b>(0.493–0.985)</b> <b>0.0407</b>	1.035 (0.965–1.110) 0.3341
CH (2008–13)	1.143 (0.905–1.445) 0.2621	1.116 (0.861–1.447) 0.4067	1.007 (0.841–1.205) 0.9426	<b>1.493</b> <b>(1.009–2.208)</b> <b>0.0451</b>	1.080 (0.901–1.294) 0.4050	1.189 (0.920–1.536) 0.1863	1.025 (0.877–1.197) 0.7594
UK (2008–13)	1.075 (0.857–1.357) 0.5194	1.106 (0.873–1.400) 0.4041	0.945 (0.739–1.208) 0.6511	1.306 (0.792–2.155) 0.2952	0.972 (0.797–1.185) 0.7778	1.355 (0.954–1.925) 0.0892	1.154 (0.950–1.401) 0.1492

# OCCORRENZA E ANDAMENTO TEMPORALE AMR

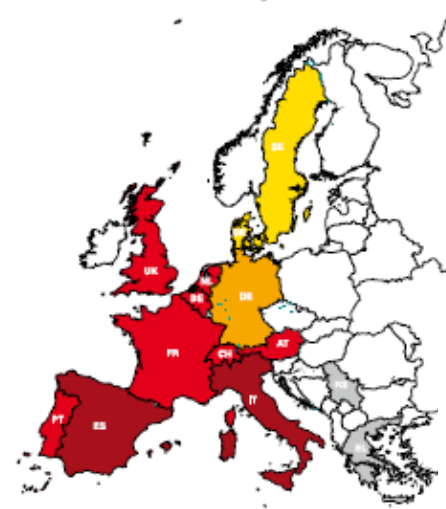
Amoxicillin clavulanate



Third-generation cephalosporins



Trimethoprim-sulfamethoxazole



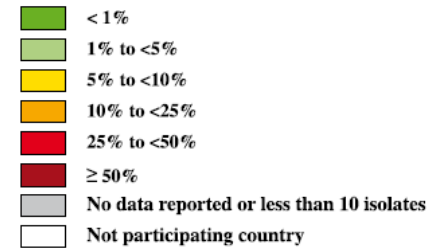
Fluoroquinolones



Gentamicin



Multidrug-resista



**% AMR *Proteus* spp. 2012-2013**

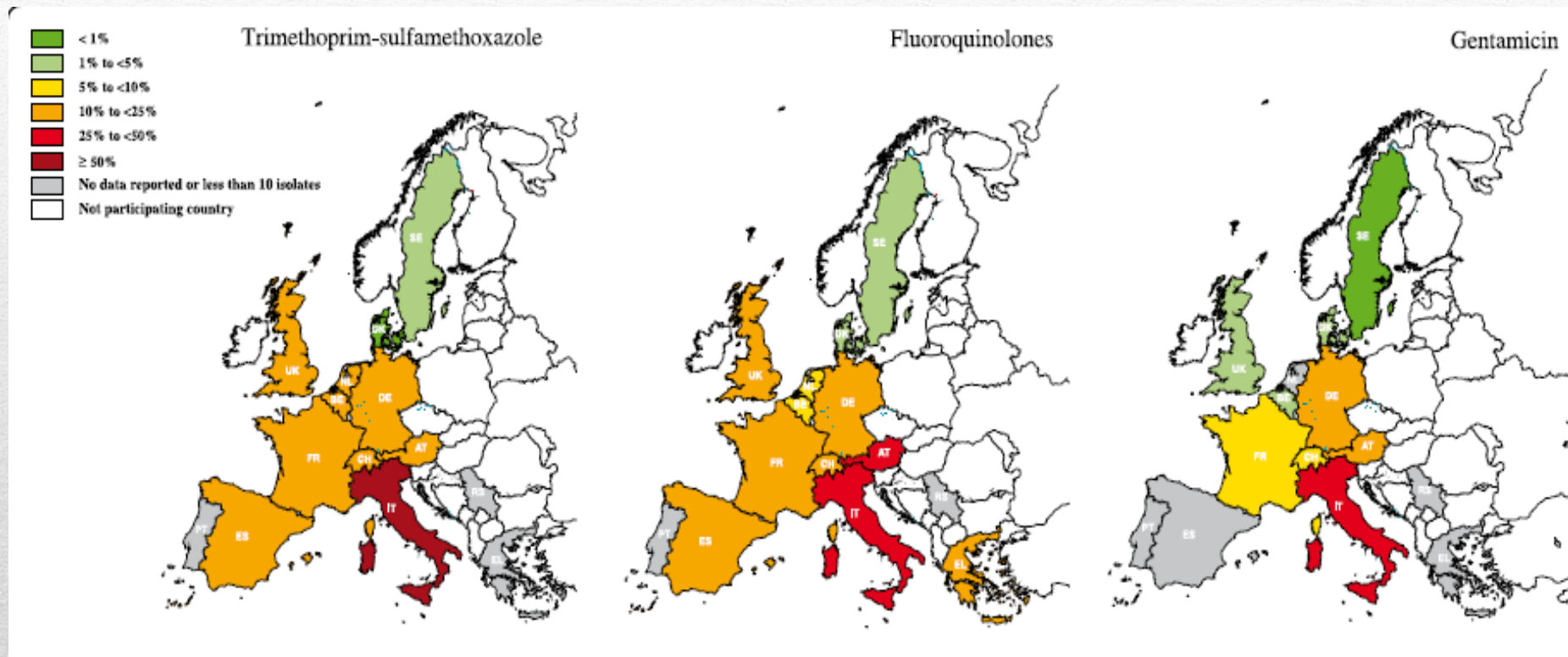
# OCCORRENZA E ANDAMENTO TEMPORALE AMR

**Table 4** Percentage of resistance in *Proteus* spp. by antimicrobial and country in 2012-2013

Country <sup>a</sup>	AMC		3GC		FLU		CN		SXT		Combined resistance		
	n	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	n	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	n	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	n	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	n	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	n	% MDR (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	% FullS (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>
DK	31	0 [a, d]	31	0 [a, b]	31	0 [e, f]	31	0 [a, b]	31	6.45 (0.00–15.10) [c]	31	0 [a, b]	93.55 (44.41–84.90) [c]
IT	12	0 [a, b, c]	12	8.33 (0.00–23.97) [a, b, c]	12	41.67 (13.77–69.56) [a, d, g]	12	8.33 (0.00–23.97) [a, b, c]	12	66.67 (39.99–93.34) [b, e]	12	8.33 (0.00–23.97) [a, b, c]	16.67 (0.00–37.75) [d]

**% AMR *Proteus* spp. 2012-2013**

# OCCORRENZA E ANDAMENTO TEMPORALE AMR



**% AMR *Staphylococcus* spp. 2012-2013**

# OCCORRENZA E ANDAMENTO TEMPORALE AMR

**Table 6** Percentage of resistance in *Staphylococcus* spp. by antimicrobial and country in 2012–2013

Country <sup>a</sup>	FLU		CN		SXT	
	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>	<i>n</i>	% R (95 % CI) <sup>b</sup> [Stat. Dif.] <sup>c</sup>
DK	51	1.96 (0.00–5.77) [c, e, f]	51	3.92 (0.00–9.25) [b]	51	0 – [d]
IT	19	42.11 (19.90–64.31) [a]	19	26.32 (6.52–46.12) [a]	19	63.16 (41.47–84.85) [e]

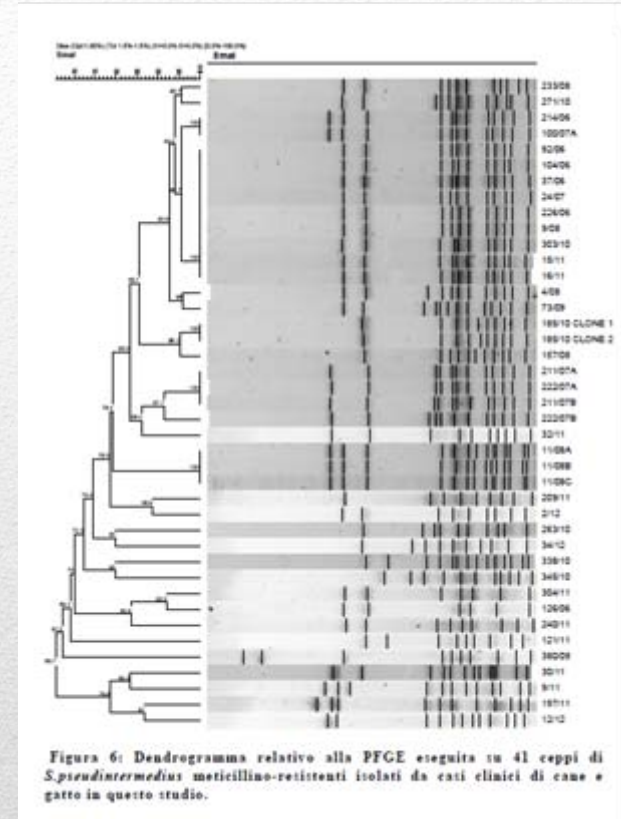
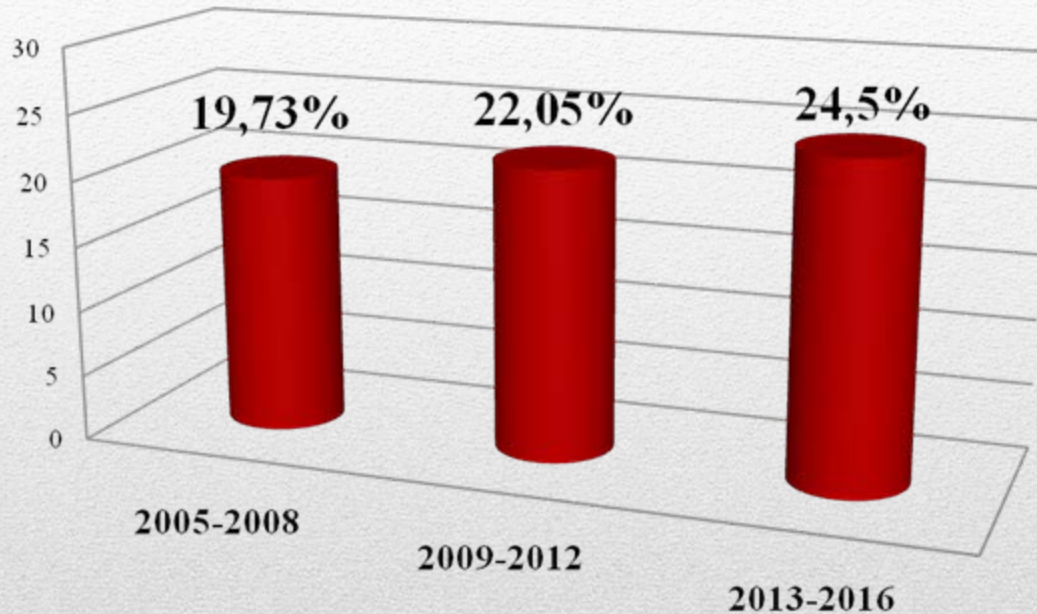
**% AMR *Staphylococcus* spp. 2012-2013**

## **Staphylococcus spp. e meticillino-resistenza nei diversi paesi nel 2012-2013**

Country <sup>a</sup>	Methicillin-resistance within each group			
	MRSA		MRSP	
	n tested	R % (95 % CI) <sup>b</sup>	n tested	R % (95 % CI) <sup>b</sup>
AT	5	1R/4S -	15	33.33 (9.48-57.19)
BE	18	5.56 (0-16.14)	64	12.50 (4.40-20.60)
DK	1	1S -	27	0 -
FR	20	40.0 (18.53-61.47)	0	- -
DE	0	- -	0	- -
EL	-	- -	1	1R -
IT	-	- -	18	50.0 (26.90-73.10)
NL	0	- -	174	10.92 (6.29-15.55)
PT	1	1R -	4	2R/2S -
RS	-	- -	-	- -
ES	-	- -	-	- -
SE	28	0 -	174	1.15 (0-2.73)
CH	2	0R/2S -	20	10.00 (0-23.15)
UK	3	1R/2S -	12	8.33 (0-23.97)



**% MRSP su 373 ceppi di *S. pseudintermedius* isolati in corso di casi clinici in cani e gatti 2005-2016 (dati DIMEVET)**



**Alla MLST l'85% dei 113 ceppi di MRSP sono risultati appartenere al sequence type (ST) 71**

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**GRAZIE PER L'ATTENZIONE**

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