## Antimicrobial resistance trends of Staphylococcus aureus isolated from bovine intramammary infections from 1990 to the present time in Argentina

Staphylococcus aureus is one of the most prevalent major mastitis pathogens in dairy herds worldwide [1]. Antimicrobial therapy is one of the bases of S. aureus mastitis control programmes, both for subclinical cases at drying off and clinical cases during lactation [1]. However, cure rates following antibiotic therapy are variable since several factors associated with host, pathogen and treatment regimen affect the probability of cure of S. aureus intramammary infection (IMI) [2]. Among pathogen factors, antibiotic resistance is an obvious reason for treatment failure, although selection of antibiotics based on in vitro susceptibility testing does not assure therapeutic success [3]. Despite this drawback, most authors agree that antibiotic susceptibility testing should precede antibiotic treatment, mainly in case of subclinical mastitis [2]. In addition, antimicrobial susceptibility testing is important for monitoring the spread of resistant strains among bacterial populations. Both determination of minimum inhibitory concentration (MIC) and disk diffusion test (DD), which is the most widely applied method in routine veterinary laboratories due to its simplicity and low cost, have been used for performing antimicrobial susceptibility surveys worldwide [4, 5]

The aim of this study was to review published research in Argentina on antimicrobial susceptibility of S. aureus isolated from bovine IMI. Searching for the survey included the words "mastitis", "bovine", "Staphylococcus aureus", "coagulase-positive Staphylococcus", "antimicrobial" "susceptibility" "resistance". Since previous reports indicated that the majority of coagulasepositive Staphylococci isolated from bovine milk are S. aureus [1] studies which characterized the isolates either as S. aureus or coagulase-positive Staphylococci were considered. The inclusion criteria were: studies

performed in Argentina which evaluated antimicrobial susceptibility of S. aureus isolated from mammary secretion quarter and composite samples of cows with subclinical and clinical mastitis using MIC determination or DD test published in peer reviewed journals. Scopus, PubMed, and Academic Google databases were searched for scientific papers unrestricted by language and published from 1990 to 2018. A total of nine scientific publications which reported antimicrobial susceptibility of S. aureus isolated from bovine mastitis in Argentina is included (Table 1). Only those antibiotics included in at least three studies are shown in the table.

## **DISCUSSION**

There is a growing concern about overuse and misuse of antimicrobial products for treating and preventing infectious diseases in cattle due to its contribution to the emergence and spread of antimicrobial resistant organisms. These organisms represent a great threat to human and animal health, and to the world ecosystem [15]. Bovine mastitis is the most frequent reason for treating both lactating and nonlactating dairy cattle [16, 17] There are no studies in Argentina and limited studies worldwide which compare resistance patterns before and after antibiotic usage throughout years using consistent procedures to evaluate the emergence of resistance due to antibiotic usage [16]. However, several studies have described occurrence of S. aureus resistance to antibiotics over time [5]. The antibiotics that were more consistently evaluated in Argentina were the beta-lactams and those of the macrolide-lyncosamide class. Penicillin is considered a first choice antibiotic for treating bovine mastitis. Resistance to penicillin has varied among studies showing the highest percentages during the first two decades (1990-2000) and lower percentages in studies conducted in the current decade. Only in one study, published in 2001, oxacillinresistant coagulase-positive Staphylococci were detected, indicating that it could have been an isolated finding. However, since methicillin-resistant *S. aureus* have been detected in several countries [18], continuous surveillance is needed for early detection of emergence of this type of resistance.

Antibiotics from macrolide-lyncosamide class are frequently used in Argentina for treating bovine mastitis [17]. Erythromycin is the most commonly tested macrolide as representative of this group, using approved human interpretive criteria [19]. Resistance to erythromycin was low in most studies. Only in one study in the last decade did it exceed 20%, which is higher than percentages reported both in Argentina and other countries [5]. Pirlimycin was commercially available in Argentina during the 90s, and has recently been introduced again to the veterinary market. Susceptibility to this antibiotic was reported in studies published at the beginning of the 2000 decade using veterinary-specific interpretive criteria [19], showing variable resistance among studies and percentages comparable with previous reports in other countries [5].

In conclusion, although information is limited, there is no apparent emergence or progression of S. aureus resistance to the most commonly used antibiotics for treating bovine mastitis in Argentina. This finding is in accord with previous reports which included studies conducted in different countries [5, 16]. However, there is a need to extend these studies to other bovine mastitis pathogens, using a harmonized approach to allow measuring resistance trends over time [15]. This information, together with increasing knowledge about antibiotic usage in dairy farms in Argentina [17] will allow us to evaluate and propose actions for a more responsible and prudent use of antibiotics in our dairy farms.

Antibiotic	% Resistant								
Penicillin	14.81	77.5	55.89	40.3	47.6	48.4	14.3	28.12	33.85
Oxacillin	-	-	2.94	0	0	0	-	0	0
Erythromycin	-	-	5.6	11.6	2	2.1	22.2	3.12	7.69
Pirlimycin	-	-	14.71	7.7	4	-	-	-	-
N	33	79	34	206	101	95	63	96	65
Reference	1	2	3	4	5	6	7	8	9

Table 1 - Antimicrobial resistance of Staphylococcus aureus isolated from bovine mastitis in Argentina ([6, 7, 8, 9, 10, 11, 12, 13, 14].

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## REFERENCES

- Barkema, H.W., Schukken, Y.H. & Zadoks, R.N. (2006) The role of cow, pathogen, and treatment regimen in the therapeutic success of bovine Staphylococcus aureus mastitis. J. Dairy Sci. 89:1877–1895
- Calvinho, L.F., Vitulich, C.A., Zurbriggen, M.A., de Canavesio, V., Tarabla, H.D. (1991a) Prevalence of udder pathogens in dairy herds from the Santa Fe dairy area (In Spanish). Therios 18:188–96.
- Calvinho, L.F., Delgado, A.R., Vitulich, C.A., Occhi, H.L., de Canavesio, V., Zurbriggen, M.A., Tarabla, H.D. (1991b) *In vitro* susceptibility to antimicrobials of udder pathogens isolated from clínicas mastitis from dairy herds of Santa Fe dairy área. (In Spanish) Vet. Arg. 8:677–680.
- Calvinho, L.F., Toselli, F.G., Weinmann, W.R., Canavesio, V.R., Neder, V.E., Igusquiza, I.A. (2002) Antimicrobial susceptibility of coagulase-positive Staphylococcus isolated from bovine mastitis in the Central dairy area of Argentina. (In Spanish). Rev. Arg. Microbiol. 34:171–175.
- CLSI (Clinical and Laboratory Standards Institute). (2013) Performance standards for antimicrobial disk and dilution susceptibility tests for bacteria isolated from animals. Clinical Laboratory Standards Institute, Wayne, PA, USA. 2013; Vol. 28, No.

- 8. 4th ed. Approved standard, VET01-A4.
- Erskine, R.J., Walker, R.D., Bolin, C.A., Barlett, P.C., White, D.G. (2002) Trends in antibacterial susceptibility of mastitis pathogens during a seven-year period. J. Dairy Sci. 85:1111–1118
- Erskine, R., Cullor, J., Schaellibaum, M., Yancey, B., Zecconi, A. (2004) Bovine mastitis pathogens and trends in resistance to antibacterial drugs. National Mastitis Council Research Committee Report. In: NMC Annual Meeting Proceedings. Pg. 400–414.
- Gentilini, E., Denamiel, G., Llorente, P., Godaly, S., Rebuelto, M., DeGregorio, O. (2000) Antimicrobial Susceptibility of Staphylococcus aureus Isolated from Bovine Mastitis in Argentina. J Dairy Sci 83:1224–1227
- González Pereyra, V., Pol, M., Pastorino, F., Herrero, A. (2015) Quantification of antimicrobial usage in dairy cows and preweaned calves in Argentina. Prev. Vet. Med. 122:273–279.
- Hoe, F.G., Ruegg, P.L. (2005) Relationship between antimicrobial susceptibility of clinical mastitis pathogens and treatment outcome in cows. J.A.V. M. A. 227:1461–1468
- 11. Huber, H., Koller, S., Giezendanner, N., Stephan, R., Zweifel. C. (2010) Prevalence and characteristics of meticillin-resistant *Staphylococcus aureus* in humans in contact with farm animals, in livestock, and in food of animal origin, Switzerland, 2009. Euro Surveill. 2010;15:1–4.
- 12. Micheo, C., Amand de Mendieta, V., Soriano, C., Tabera, A., Stefano, A., Casasnovas, G., Purrán, P., Corradeti, A., Carabajal, S. (2001) Study of in vitro susceptibility of bacterial strains isolated from bovine mastitis in Mar y Sierras dairy area. (In

- Spanish). Vet. Arg. 18:588-597.
- Neder, V.E., Araujo, L., Gianre, V.R., Calvinho, L.F. (2016) Antibiotic susceptibility of Staphylococcus aureus isolated from bovine mastitis in dairy farms from the central dairy area of Argentina (In Spanish). REDVET Rev. Electrón. vet. Vol. 17 № 9 - http://www.veterinaria.org/revistas/redvet/n09 0916 html
- 14. Office International des Épizooties. (2016) The OIE Strategy on Antimicrobial Resistance and the Prudent Use of Antimicrobials. www.oie.int/antimicrobial-resistance.
- Oliver, S.P., Murinda, S.E. (2012) Antimicrobial resistance of mastitis pathogens. Vet. Clin. Food Anim.28:165–185.
- 16. Pellegrino, M.S., Frola, I.D., Odierno, L.M., Bogni, C.I. (2011) Mastitis bovina: resistencia a antibióticos de cepas de *Staphylococcus au*reus aisladas de leche. (In Spanish). REDVET Rev. electrón.vet. http://www.veterinaria.org/revistas/redvet Volumen 12 N° 7 - http://www.veterinaria.org/ revistas/redvet/n070711.html
- Russi, N.B., Bantar, C., Calvinho, L.F. (2008) Antimicrobial susceptibility of Staphylococcus aureus causing bovine mastitis in Argentine dairy herds. Rev. Arg. Microbiol. 40:116–119.
- Srednik, M.E., Abate, S., Gentilini, E.R. (2016) Antibiotic susceptibility of *Staphylococci* isolated from milk samples obtained from bovine mastitis (In Spanish). In Vet. 18:39–44.
- Zecconi, A., Calvinho, L., Fox, L. (2006) Staphylococcus aureus intramammary infections. Bulletin of the International Dairy Federation. 408/2006. Pp 39. FIL-IDF. ISSN 0250–5118