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Poster Number: 45

Topic Area: Bacterial Disease

Title: Multiple-drug resistant Salmonella enterica serovar Typhimurium monophasic variant (4,[5],12:i:-)

isolated from swine of Santa Catarina state - Brazil

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Santa Catarina (SC) raises approximately 30% of all pigs produced in Brazil, and is responsible for 50% of total pork exports. In recent years, Salmonella 4,[5],12:i:- which is considered to be a serologic variant of S. enterica serovar Typhimurium, has emerged as an important cause of salmonellosis in pigs. Between 2012 and 2016, 62.85% (22/35) of Salmonella strains submitted to a animal diagnostic center were identified as S. 4,[5],12:i:-. Although this data represents laboratory occurrence and not prevalence, it raises an alert due to its clinical relevance. Here we objective to characterize the strain and its in-farm epidemiology. The twenty-two S. 4,[5],12:i:- strains were categorized according to clinical presentation: enteric (isolated from intestinal samples) or septicaemic (isolated from internal organs). One strain (recovered from a septicaemic nursery pig) was submitted for whole genome sequencing (WGS), sequence-typed (ST) and analyzed for the presence of antibiotic resistance traits. Out of the 22 isolates, one was isolated from a suckling pig and was classified as enteric. Eleven isolates were recovered from nursery pigs, being nine enteric and two septicaemic presentations. Ten were isolated from grower/finishers pigs, whereas 8 were classified as enteric and 2 as septicaemic. WGS identified the S. 4,[5],12:i:- strain as ST-19, which was previously associated with serovar Typhimurium isolated from swine. The sequenced strain is predicted to be resistant to multiple drugs (MDR), as acquired resistance genes against aminoglycosides (aac(6')-laa, aadA1, aac(3)-lia), to beta lactam (blaTEM-IB), to fluoroquinolone (qnrE1), to phenicol (floR), to sulphonamide (sul1), to tetracycline (tetA), and to trimethoprim (drfA1). Furthermore, the antibiotic resistance plasmids IncHI2-type (predominant in antibiotic resistant Salmonella), IncFIA-type, IncH12A-type and p0111-type (associated with other Enterobacteriaceae genera) were also detected. Finally, chromosomal point mutations to nalidixic acid and ciprofloxacin (gyrA gene and point mutation p.S83Y) were also identified, typically related to Salmonella isolated from swine. These findings suggest that S. 4,[5],12:i:- is an emerging threat to commercial farms in SC. Its presumptive resistance to multiple antibiotics may have major public health implications. Further analyses will be carried out to compare S. 4,[5],12:i:- strains from outbreaks occurred in Brazil and other countries, to derive directions in regard to pathogen control and disease eradication.