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Evaluation of synergistic activity of lactoferrin with antibiotics against drug resistant bacterial pathogens

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The multidrug-resistant (MDR) bacterial infections have escalated as one of the world's utmost health issues. The progress of novel antibiotics has declined over the last half century. The aim of this study was to determine the effect of lactoferrin (human, bovine and camel) on minimum inhibitory concentrations (MICs) of important antibiotics in use against drug resistant bacterial pathogens encountered in the region of Hail, Kingdom of Saudi Arabia (KSA). Totally 147 clinical bacterial isolates were successfully isolated. Pathogens included were: Methicillin resistant *Staphylococcus aureus* (MRSA)-30 isolates; methicillin resistant coagulase negative *Staphylococcus*-30 isolates; extended-spectrum beta-lactamase (ESBL) producing *Enterobacteriaceae*-40 isolates; fluoroquinolone resistant gram negative pathogens-30 isolates; multidrug resistant *Pseudomonas* species- 05 isolates; carbapenem resistant gram negative pathogens- 05 isolates; AmpC beta-lactamase producing gram negative pathogens- 05 isolates; vancomycin resistant enterococci (VRE)- 02 isolates. The methods employed were MALDI-TOF for identification, MicroScan WalkAway system for identification, susceptibility testing and LF synergism testing. PCR-Sanger sequencing was done (before and after exposure to LF synergism) to study the molecular biology aspects of the study. In our study, the reproducible synergism effect of LF with antibiotics was found to be remarkable. To specify the phenotypic effects of LF in synergism with antibiotics: the isolates producing ESBL had turned non-ESBL; quinolone resistant isolates had turned susceptible; MRSA had turned MSSA (methicillin susceptible) and VRE had turned susceptible. The molecular biological study suggests changes only in the gene expression after the exposure to LF compounds. The results of this study demonstrated similar effect with comparable results for the LF tested from three different sources (human, bovine and camel). The outcome knowledge of the study would certainly help the Ministry of Health (MOH) in planning the LF adjuvant treatment methods for locally faced drug resistant pathogens causing different infections.

No LF		Microbiology Report		With hLF		Microbiology Report	
HAIL UNIVERSITY - MDRPTU		HAIL UNIVERSITY - MDRPTU		HAIL UNIVERSITY - MDRPTU		HAIL UNIVERSITY - MDRPTU	
Name		Specimen	MB1402700	Name		Specimen	MB1402700hLF
Patient ID	MB1402700	Source	Dr Godfred	Patient ID	MB1402700hLF	Source	Dr Godfred
Date of Birth		Ward of No.	Dr Godfred, Molecular	Date of Birth		Ward of No.	Dr Godfred, Molecular
Att. Phys.				Att. Phys.			
1 Enterococcus faecium				1 Enterococcus faecium			
1 E. faecium				1 E. faecium			
Drug	MIC (mg/L)	Drug	MIC (mg/L)	Drug	MIC (mg/L)	Drug	MIC (mg/L)
Amoxicillin	>384	Amoxicillin	>384	Amoxicillin	>384	Amoxicillin	>384
Ampicillin	>16 R	Ampicillin	>16 R	Ampicillin	>16 S	Ampicillin	>16 S
Ciprofloxacin	2 I	Ciprofloxacin	2 I	Ciprofloxacin	>16 S	Ciprofloxacin	>16 S
Gentamycin	>32	Gentamycin	>32	Gentamycin	>32	Gentamycin	>32
Daptomycin	4 S	Daptomycin	4 S	Daptomycin	4 S	Daptomycin	4 S
Erythromycin	>4 R	Erythromycin	>4 R	Erythromycin	>4 R	Erythromycin	>4 R
Fosfomycin	>32	Fosfomycin	>32	Fosfomycin	>32	Fosfomycin	>32
Fusidic Acid	>12	Fusidic Acid	>12	Fusidic Acid	>12	Fusidic Acid	>12
Gen. Synergy	>500	Gen. Synergy	>500	Gen. Synergy	>500	Gen. Synergy	>500
Gentamicin	>16	Gentamicin	>16	Gentamicin	>16	Gentamicin	>16
Linezolid	2 S	Linezolid	2 S	Linezolid	2 S	Linezolid	2 S
Moxifloxacin	>16 S	Moxifloxacin	>16 S	Moxifloxacin	>16 S	Moxifloxacin	>16 S
Mupirocin	1	Mupirocin	1	Mupirocin	1	Mupirocin	1
Nitrofurantoin	>16	Nitrofurantoin	>16	Nitrofurantoin	>16	Nitrofurantoin	>16
Oxacillin	>16	Oxacillin	>16	Oxacillin	>16	Oxacillin	>16
Penicillin	>16 R	Penicillin	>16 R	Penicillin	>16 R	Penicillin	>16 R
Rifampin	>16 S	Rifampin	>16 S	Rifampin	>16 S	Rifampin	>16 S
Strep. Synergy	>1000	Strep. Synergy	>1000	Strep. Synergy	>1000	Strep. Synergy	>1000
Synercid	>16 S	Synercid	>16 S	Synercid	>16 S	Synercid	>16 S
Tecoplanin	>16 S	Tecoplanin	>16 S	Tecoplanin	>16 S	Tecoplanin	>16 S
Tetracycline	>16 R	Tetracycline	>16 R	Tetracycline	>16 R	Tetracycline	>16 R
Trimeth/Sulfa	>16 S	Trimeth/Sulfa	>16 S	Trimeth/Sulfa	>16 S	Trimeth/Sulfa	>16 S
Vancomycin	>16 R	Vancomycin	>16 R	Vancomycin	>16 S	Vancomycin	>16 S

Figure 1: Demonstration of synergistic effect of hLF (human lactoferrin) on vancomycin resistant *Enterococcus faecium* (VRE).

Biography

Godfred A Menezes is currently working at RAK College of Medical Sciences (RAKCOMS), RAKMHSU, UAE. He was an Assistant Professor and Scientist at Hail University, Saudi Arabia; In-charge/Scientist of Central Research Laboratory (CRL) and; Assistant Professor of Microbiology at Sree Balaji Medical College & Hospital, Chennai, India for three years. He has also worked as a Scientist in the Department of Medical Microbiology and Infectious Diseases, Netherlands. He has been worked extensively on molecular characterization of antimicrobial resistance in clinical bacterial pathogens. He has been a faculty of several medical institutes and also a Para-Medical Institute.

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