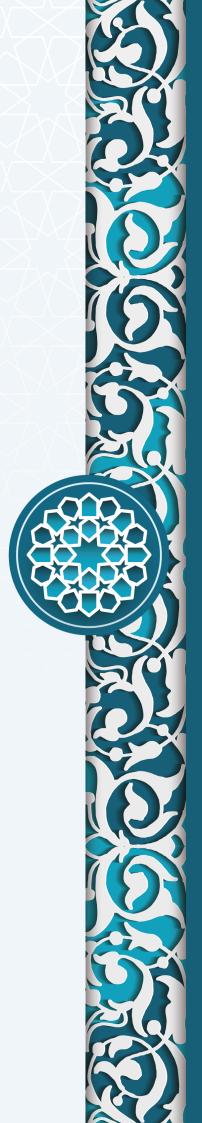
Therapeutic strategies for eliminating biofilm produced by Staphylococcus aureus isolated from catheter by exposing to radiation emitted from radioactive sources (in vitro)

إعداد

Dr.Nebras Rada Mohammed and Noor Ali Salman Ibn sina University of Medical and Pharmaceutical Sciences/ College Dentistry/ Baghdad, Iraq Author: nebrasrada88@ibnsina.edu.iq



الجامعة الإسلامية بمنيسوتا Islamic University of Minnesota I UM المركز الرئيس





Abstract:

Objective: In order to get rid of the biofilm and Staphylococcus aureus bacteria producing the biofilm using different therapeutic.

Study design: Cross-sectional in descriptive study design with case –control in analytical study design.

Backgrounds: S. aureus are gram positive bacteria, nonmotile, non-spore forming, facultative anaerobes with grow on aerobic respiration. Cause several diseases include skin infection, pneumonia, sepsis and bacteremia. It is development by formation biofilm with stages attachment, adhesions, aggregations and biofilm maturation by quorum sensing. There are several strategies anti-biofilm inclusive natural products like plant extracts, photochemical. Sr90 is a radioactive sources isotope of Strontium by nuclear fission, possess different application in medicine and industry. Methodology: Study populations and bacterial isolates with diagnosing S. aureus from catheter and several diseases from patients in Baghdad hospitals in 2024 /2023. Expose the bacteria to radiation emitted from Sr90 radioactive sources in different doses and screening biofilm production before and after exposure to radiation.

Results: The bacteria were biofilm producers before (%100) exposure to radiation and after exposure to radiation. All bacteria lost biofilm production by (%0) when grown on the

medium Congo-Red Agar (CRA) which changed the color of the bacterial colonies to red. This is an indication and evidence of their loss of biofilm production compared to the control before exposure to radiation emitted by Sr90 whose colonies were colored. Results of therapeutic strategies for killing S. aureus by exposing to Sr90 radiation without alaminium of activity 10 µci, percentage of humicide 100 % in doses 3.284235 through 1hr; 6.568471 through 2hr.; 9.852705 through 3hr. Also, exposed to Sr90 radiation to alaminium in activity 10 µci, percentage of humicides 100 % by doses 5-10*5.2076 through 1hr.; 4-10*1.04153 through 2hr. and 3-10*1.56228 through 3hr.

Conclusions: Production of biofilm from S. aureus lower after exposure to Sr90 without aluminum indicated by the color of culture of colony become red compared with control.

Key words: Radiotherapy, rays, bacteria and biofilm.

Introduction

Staphylococci are gram-positive bacteria, non-motile, nonspore forming, facultative anaerobes and implant in aerobic respiration. It is deem to be a major pathogen that settles and infects both hospitalized patients with lowered immunity and healthful immune competent human in the society [1]. Cause several diseases include skin infections, pneumonia and bacteremia [3,2].

Natural outputs to biofilm inhibitory and disrupt activity that significant provenance of biocompatible anti-biofilm factor to evade the put away effectiveness of conventional antimicrobial on person good heal. Like, primary oils of worts [6,5,4], phenolic acids [9,8,7]. It is sturdy curative contra pathogens every after the aurous come of antimicrobial in the mid20-th hundred years [11,10]. The biofilm expansion process in four phases inclusive engagement and coherence, gathering to extracellular matrix with microbial propagation, biofilm constructing with maturation and biofilm stampede with cell disengagement [,12, 13].

Multiple regulatory suits control for every step of the foremost engagement to the maturation with stampede of biofilm. QS framework is an interior connection framework of bacteria wheresoever the expression of pertinent genes is started during the alterations to the indicative compound arrive a special outset. It encompass diverse indicative transduction path which regulate biofilm formulation malice and movement [14]. In S. aureus QS framework the extension control factor Agr framework responsible for QS [15]. Evolving novel anti-



biofilm agents from natural products like plant extracts and photo chemicals by suppression of microbial cell engagement and cohesion, deactivation of polymer formulation, lowering in the obstetrics of diseases factors and obstruction of QS framework [16]. Strontium90- (Sr90) is a radioactive isotope created via atomic fission with a semi-lifetime of 28.8 years. It kowtow β -dissolution into yttrium90- to a dissolution power of 0.546 MeV. It has implementation in medical and industrial uses [17].

Methodology

Study design

Case-Control study design depending in it research for analytical study design with Cross-Sectional for descriptive study design.

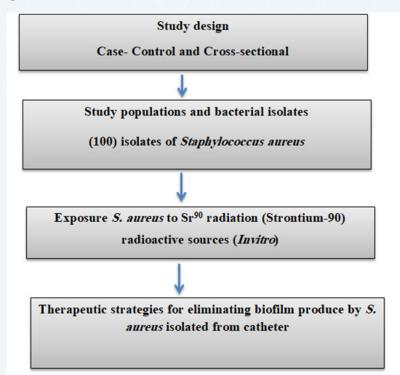


Figure (1): Plan of study design of this explore.

Study populations and Bacterial isolates

A overall of assemblage of specimens (100) S. aureus for patients that were assumptive in Baghdad hospital through 2024/2023 which diagnosis via classical biochemical screening based [18].

Therapeutic strategies for exposure S. aureus to Strontium (Sr90) radioactive sources in different doses

S. aureus grow was completed depending to [19] with many alterations, implanted within Nutrient stock at °37 C for 24 h, subsequently throw away at 5000 rpm until 10 min. The pellet was hanging of sterile naturalistic saline and compare with MacCfrland 0.5, subsequently exhibition 1 ml of hanged to rays released of Sr90 with comparison of dominance (wanting exposure to rays), every round was done in replicate and injected in Trypton soy agar.

The equation of attribution of death:

Therapeutic strategies for eliminate biofilm produced by S. aureus isolated from catheter by exposing to Sr90 Congo Red process, the media formative of BHIB (37 g/ l), sucrose 5g/ l), agar numeral 10) 1 g/ l) and Congo red dye (0.8 gm/ l). Congo red dye was all set intensive liquid sol and autoclaved at 121 oC for 15 minutes. Thereafter, it was add up to autoclaved Brain heart infusion agar with each other with sucrose at 55 oC. Paten were grown with test organism and



brood at 37 oC for 24 to 48 hr. aerobically. Black colonies to a dry crystalline tenacity particular biofilm output [20].

Results and discussions

The disintegration of the extra-polysaccharide is of specific connection for anti-biofilm mensuration. Thus far, different factors have been utilized to eliminate elementary and mingled species biofilms, originally via degenerating auto-produced adhesions, nucleic acids and polysaccharides [21].

)

The results in table (1) and figure (2) show biofilm production from S. aureus bacteria before and after exposure to radiation emitted from the Sr90 radioactive source (Strotium90-). The bacteria were biofilm producers before (%100) exposure to radiation and after exposure to radiation. All bacteria lost biofilm production by (%0) when grown on the medium Congo-Red Agar (CRA) which changed the color of the bacterial colonies to red. This is an indication and evidence of their loss of biofilm production compared to the control before exposure to radiation emitted by Sr90, whose colonies were colored. Dark black on Congo-Red Agar medium. Thus, it was proven that the radiation emitted by Sr90 is very effective and powerful in eliminating the biofilm of bacteria isolated from heart and urinary tract catheterization devices and isolated from patients after their catheterization procedure.

Table (1): Biofilm production from S. aureus before and after radiation



الجامعة الإسلامية بمنيسوتا العامينيينية المركز الرئيسيات

No.	Biofilm production before radiation	Percentage	Biofilm production after radiation	Percentage
1	+	100%	-	0%
2	+	100%	-	0%
3	+	100%	-	0%
4	+	100%	-	0%
5	+	100%	-	0%
6	+	100%	-	0%
7	+	100%	-	0%
8	+	100%	-	0%
9	+	100%	-	0%
10	+	100%	-	0%
11	+	100%	-	0%
12	+	100%	-	0%
13	+	100%	-	0%
14	+	100%	-	0%
15	+	100%	-	0%
16	+	100%	-	0%
17	+	100%	-	0%
18	+	100%	-	0%
19	+	100%	-	0%
20	+	100%	-	0%
21	+	100%	-	0%
22	+	100%	-	0%
23	+	100%	-	0%
Control (++++) Black colony		(+) : Positive production/ Black colony (-) : Negative production/ Red colony		

🖸 ium.usa.online@gmail.com 🚯 w w w . i u m . e d u . s o

 $\left(C \right)$

6

🔇 + 967 730053135 🚯 http://ium.usa.online

A previous study by [22] display Nerolidol was locate to repress S. aureus biofilm via more seventy percentage at concentricity extending of one to four mg/ml [131].

A preceding study by [23] exhibit Alkaloids, aromatic acids are naturalistic components that anti-biofilm efficiency contra S. aureus like, the alkaloid sinomenine able essentially up regulate agrA and down-regulate icaA level.

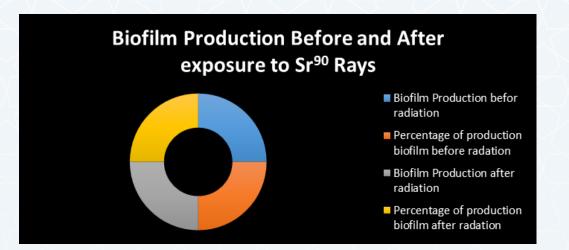


Figure (2): Biofilm production from S. aureus before and after exposure to Sr90 radiation.

A antecedent study by [24] revelation restricted therapeutic chosen for S. aureus recognizing efficient incorporation supplies an substitutional for infection therapy. Like, aside from possess considerable biological effectiveness, curcuminfounded metal complicated induce the bioavailability of curcumin. At the concentricity of one-hundred μ M, curcumin suppressed S. aureus biofilm formulation fifty-six percentage whilst oxovanadium complicated of curcumin has a safely sturdy impact eithy-two percentage that might stay in the interaction together impact of complicated technicality have suppression and antibacterial technicality.

۲۰۰۲ ۲۰۱۲ الجامعة الإسلامية بمنيسوتا المحرب المثلينيين المركب المركب المركب المركب المركب المركب



A-

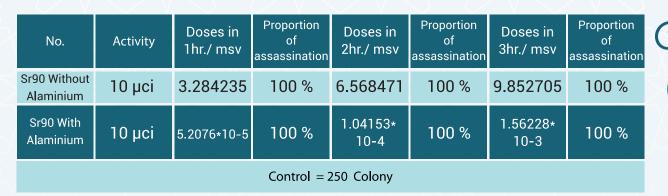
Figure (3): A- S. aureus production biofilm before exposure to Sr90 radioactive sources. B- S. aureus production biofilm after exposure to Sr90 radioactive sources.

A past study by [25] show the extractor of methanol from Hemidesmus indicus root that interaction together the anti-biofilm effectiveness of AML and CL contra Methicillin Resistance S. aureus. Furthermore, in the renal and liver of Methicillin Resistance S. aureus -infective mouse, the aggregates safely minimized microbial carriage, sickness effectiveness effect and gram-positive place.

Results of therapeutic strategies in table (2) for killing S. aureus exposed to Sr90 radiation wanting alaminium in activity 10 μ ci, percentage of humicides 100 % through doses 3.284235 through 1 hr.; 6.568471 in 2 hr.; 9.852705 in 3 hr. Also, exposed to Sr90 radiation wanting alaminium of activity 10 μ ci, percentage of humicide 100 % in doses 3.284235

through 1hr; 6.568471 through 2 hr.; 9.852705 through 3 hr. Also, exposed to Sr90 radiation to alaminium in activity 10 µci, percentage of humicides %100 via doses 5-10*5.2076 through 1hr.; 4-10*1.04153 through 2 hr. and 3-10*1.56228 through 3 hr. as shown in table (2).

Table (2): Proportion of homicides of S. aureus exposure to Sr90 radioactive sources.



A anterior study by [26] demonstration a fatal impact on Pseudomonas through displayed to rays on various times through (1,2,3) hr. with diverse doses by cobalt with impact 1 µci and 10 µci in the existence and non-attendance of aluminium with the use of strontium in the existence and non-attendance of aluminium. The killing rate of P. aeruginosa by cobalt wanting aluminium was %78 with in the existence of aluminium %100. In rapprochement, the death average of Pseudomonas with efficient wanting aluminium was %100; in the existence of aluminium %98, the death average of Pseudomonas wanting aluminium was %83 and in the existence of aluminium %96 rapprochement with control.

Conclusions

Production of biofilm from S. aureus lower after exposure to Sr90 without alminium indicate by the color of culture become red compared with control is black colony.

Ethical approval

All examination protocols were confirmed by the College of Ibn Sina University of Medical and Pharmaceuticals Sciences. All screening was achieved following the confirmed guidelines.

Financial support and sponsorship

There was no financial disclosure.

Conflicts of interest

The authors declare that there are no conflicts of interest.

Acknowledgment about author

Researcher Dr. Nebras Rada Mohammed PhD. in Biotechnology with a Genetic Engineering, Molecular Genetics and Protein Engineering, a scientist, expert, researcher, creator, inventor, writer, written and author, editor-in-chief of the Journal of Articles and Inventions in the American Goidi Journal, teaching, lecturer at the University College of Al-Turath University college, a Bachelor>s degree in Microbiology and a Master>s degree in Molecular Biology in Microbiology from Al-Mustansiriya University, an arbitrator, international resident and consultant In medical laboratories, an expert in medical laboratories and a holder of the title of a scientist project, an arbitrator, a distinguished publisher, a silver supporter of



scientific platforms, a chairman of a committee in a scientific society, receiving accolades from international intellectual property, the Best Arab Woman Award 2020, also the Best Community Personality Award, the Best Research Award 2019, also the Best Research Award 2020 and an American Award For the invention of 2020 by the American Goidi the World Investment Commission in America, holds the title of the best distinguished inventor in the world by the World Investment Commission in America and holds the first places in the world for inventions presented in the world from the American Goidi, the world investment commission in America. The Edison Prize, The Pascal Prize, The creativity award, the scientific medal and the Everest medal for innovation, creativity for inventions from USA.

References

- Harris, L.G.; S.J. Foster and R.G. Richards (2002). An introduction to Staphylococcus aureus and techniques for identifying and quantifying S. aureus adhesins in relation to biomaterials: review. European Cells and Materials. Vol.4.2002(pages60-39). ISSN : 2262-1473.
- Cheung, G.Y.; Bae, J.S. and Otto, M. (2021). Pathogenicity and virulence of Staphylococcus aureus. Virulence.69-547:(1)12. https://doi.org/2150559/10.1080 4.2021.1878688.
- Van Hal, S.J.; Jensen, S.O.; Vaska, V.L.; Espedido, B.A.; Paterson, D.L. and Gosbell, I.B. (2012). Predictors of mortality in Staphylococcus aureus bacteremia. Clinical microbiology reviews. 86-362:(2)25. https://doi. org/10.1128/CMR.11-05022
- Zhao, X.C.; Liu, Z.H.; Liu, Z.J.; Meng, R.Z.; Shi, C. and Chen, X.R. (2018). Phenotype and RNA-seq-based transcriptome profiling of Staphylococcus aureus biofilms in response to tea tree oil Microb Pathog, 123, pp. 10.1016 ,313-304/j. micpath.2018.07.027 (https://doi.org/10.1016/j. micpath.2018.07.027)
- Sharifi, A.; Mohammadzadeh, A.; Zahraei Salehi, T. and Mahmoodi, P. (2018). Antibacterial, antibiofilm and antiquorum sensing effects of Thymus daenensis and Satureja hortensis essential oils against Staphylococcus aureus isolates J Appl Microbiol, 2) 124), pp. ,388-379 10.1111/jam.13639 (https://doi.org/10.1111/jam.13639).
 - L.N. Silva, ; Da Hora, G.C.A.; Soares, T.A.; Bojer, M.S.; Ingmer,

H. and Macedo, A.J. Myricetin (2017). Protects Galleria mellonella against Staphylococcus aureus infection and inhibits multiple virulence factors Sci Rep, 1) 7). p. ,2823 10.1038/s1-02712-017-41598 (https://doi.org/10.1038/s1-02712-017-41598)

- Mu, D.; Xiang, H.; Dong, H.S.; Wang, D.C. and Wang, T.D. (2018). Isovitexin, a potential candidate inhibitor of Sortase A of Staphylococcus aureus USA300 J Microbiol Biotechnol, 9) 28), pp. 10.4014 ,1432-1426/jmb.1802.02014 (https:// doi.org/10.4014/jmb.1802.02014)
- Mu, D.; Luan, Y.X.; Wang, L.; Gao, Z.Y.; Yang, P.P. and Jing, S.S. (2020). The combination of salvianolic acid A with latamoxef completely protects mice against lethal pneumonia caused by methicillin-resistant Staphylococcus aureus Emerg Microb Infect, 1) 9), pp. 22221751.2020.1711817/10.1080 ,179-169 (https://doi. org/22221751.2020.1711817/10.1080)
- Qin, N.; Tan, X.J.; Jiao, Y.M.; Liu, L.; Zhao, W.S. and Yang, S.(2014). RNA-Seq-based transcriptome analysis of methicillin-resistant Staphylococcus aureus biofilm inhibition by ursolic acid and resveratrol Sci Rep, 4, p. 10.1038 ,5467/srep05467 (https://doi.org/10.1038/ srep05467).
- Merghni, A.; Noumi, E.; Hadded, O.; Dridi, N.; Panwar, H. and Ceylan, O. (2018). Assessment of the anti-biofilm and antiquorum sensing activities of Eucalyptus globulus essential oil and its main component -1,8cineole against methicillin-resistant Staphylococcus aureus strains Microb Pathog, 2018) 118), pp. 80-74.

Natural products as platforms to overcome antibiotic resistance (2017). Chem Rev, 19) 117), pp. 12474-12415. (https://doi.org/10.1021/acs.chemrev.7b00283).

Cheng, C. S.; Jiang, T.; Zhang, D.W.; Wang, H.Y.; T. Fang, C.C. Li Attachment characteristics and kinetics of biofilm formation by Staphylococcus aureus on ready-to-eat cooked beef contact surfaces J Food Sci, 2023) (6) 88), pp. 3841.16592-1750/10.1111 ,2610-2595 (https://doi. org/3841.16592-1750/10.1111).

Konduri, R.; Saiabhilash, C.R. and Shivaji, S. (2021). Biofilmforming potential of ocular fluid Staphylococcus aureus and Staphylococcus epidermidis on ex vivo human corneas from attachment to dispersal phase Microorganisms, 9 6)), p. 1124.

- Peng, Q.; Tang, X.H.; Dong, W.Y.; Sun, N. and Yuan, W.C. (2022). A review of biofilm formation of Staphylococcus aureus and its regulation mechanism. Antibiotics, 1) 12), p. 12.
- Pei, J.J.; Huang, Y.G.; Ren, T.; Guo, Y.D.; Dang, J. and Tao, Y.D. (2022). The antibacterial activity mode of action of plantaricin YKX against Staphylococcus aureus. Molecules, 13) 27), p. 10.3390 ,4280/molecules27134280.
- Guo, X.Q.; Mei Aloe vera, N. (2016). A review of toxicity and adverse clinical effects. J Environ Sci Health C Environ Carcinog Ecotoxicol Rev, 2016) (2) 34), pp. 96-77.
- Meng, W.; Huang, W.J.; Kondev, F.G.; Audi, G. and Naimi, S. (2021). «The AME 2020 atomic mass evaluation (II). Tables, graphs and references». Chinese Physics C. :(3) 45 030003.



- Forbes, B.A.; Sahm, D.F. and Weissfeld, A.S. (2007). Baily and Scott>s Diagnostic Microbiology.11thedition. Mosby, Inc. Baltimore, USA. 398-384.
 - Ismail, M.C.H.; Waleed, S.; Jabbar, F. and Ibrahim, K. (2012). Effect of Diode Lazer (805) nm on alpha –toxin production and antibiotic sensitivity of Staphylococcus aureus. Iraqi J. Sci Vol 53. No. 2. Pp 759-755.
- Arciola, C.R.; Baldassarri, L. and Montanaro, L. (2001) Presence of icaA and icaD genes and slime production in a collection of Staphylococcal strains from catheter associated infections. Journal of clinical microbiology. 2156-2151:(6)39.
- Yuan, G.J.; Li, P.Y.; Xu, X.J.; Li, P.B.; Zhong, Q.W. and He, S. (2020). Azalomycin F5a eradicates Staphylococcus aureus biofilm by rapidly penetrating and subsequently inducing cell lysis Int J Mol Sci, 3) 21), p. 862.
- de Moura, D.F.; Rocha, T.A.; de Melo Barros, D.; da Silva, M.M.; Dos Santos Santana, M. and Neta, B.M. (2021). Evaluation of the antioxidant, antibacterial, and antibiofilm activity of the sesquiterpene nerolidol Arch Microbiol, 203 7)), pp. 4311-4303.
- Yum, S.-J.; Jeong, H.G. and Kim, S.M. (2023). Anti-biofilm effects of sinomenine against Staphylococcus aureus Food Sci Biotechnol, 1) 32), pp. 90-83.
- Katsipis, G.; Tsalouxidou, V; Halevas, E.; Geromichalou, E.; Geromichalos, G. and Pantazaki, A.A. (2021). In vitro and in silico evaluation of the inhibitory effect of a curcuminbased oxovanadium (IV) complex on alkaline phosphatase activity and bacterial biofilm formation Appl Microbiol

Biotechnol, 1) 105), pp. 168-147.

- Sannat, C.; Hirpurkar, S.D.; Shakya, S.; Dutta, G.K.; Roy, M. and Jolhe, D.K. (2022). Methanolic extract of Hemidesmus indicus root augments the antibacterial and antibiofilm activity of amoxicillin and clindamycin against methicillinresistant Staphylococcus aureus of bovine origin. Lett Appl Microbiol, 6) 75), pp. 1589-1579.
- Mhana, W. J. (2022). The Impact of Cobalt and Strontium Radiation on Pseudomonas aeruginosa. Al-Mustansiriyah Journal of Science. ISSN: 635-1814X (print), ISSN:-2521 3520 (online) Volume 33, Issue 3.

-Alziyyadi H, Obaid.S.Hanan, almusawi AMohammed,AL-JamaL S,(2023). E-learning after the Corona pandemic - a strategic necessity and a development alternative INTERNATIONAL MINNESOTA JOURNAL OF ACADEMIC STUDIES, (VOL,1),(ISSUE,3), PP.37-15.



الجـامعة الإسـلامية بمنيسوتا Islamic University of Minnesota المـركــز الـرئــيــســـي IV M

