ORIGINAL ARTICLE

Megrment the Level of SDF1-A in the Serum of Patients with Urinary Tract Infection and Patients with Vaginal Infection

AHMED ABBAS ALI1, HAYDER HAMZAH IBRAHIM2

¹Al Furat Al-Awsat Technical University, College of Health and Medical Techniques, Iraq ²Technical Institute Babil, Al Furat AL-Awsat Technical University, Iraq Correspondence to: Ahmed Abbas Ali

ABSTRACT

Urinary tract infection, Bacterial vaginosis as one of the global threats affecting millions of women and causing some time deaths around the world.

Objective: This study aims to assess levels of SDF1 and IgA in patients with Urinary tract infection, Bacterial vaginosis, and people enjoying good health as a control group.

Methods: Collecting medical information from (180) participants in the Imam AI-Sadiq General Teaching Hospital in Iraq-Babylon, the Gynecology Consultant, according to specific criteria, the subjects were divided into three groups: the control group, the patients (UTI, BV), while the demographic study included age, education, Jobs, and living Laboratory results, signs and symptoms, SDF1-α and IgA levels were assessed by the enzyme-linked immunosorbent assay (ELISA) technique.

Results: The results revealed that there were significant variations in SDF1 and IgA between the UTI group, BV group, and control groups. SDF1 level with the UTI group was (1.206522 ±.0927277 pg/ml) and for patients with BV was(1.213735 ±.0661389 pg/ml) whereas the control group appear was (1.130013±.0496400 pg/ml), IgA level in the UTI patients was (64.252983 ± 7.5946759pg/ml) and patients with BV was (60.441928±6.1457661pg/ml) whereas the control group appears was (48.011745 ±4.7938613 pg/ml)

Conclusion: They can be considered good indicators to give knowledge about the diagnosis of urinary tract infection and bacterial vaginosis, and help doctors to give appropriate medications.

Keywords: Urinary tract infection, bacterial vaginosis, stromal cell-derived factor 1-alpha, Immunoglobulin A.

INTRODUCTION

Urinary tract infections(UTIs) are bacterial infections of the urinary tract and can involve both the lower (cystitis) and upper (pyelonephritis). UTI occurs in females of any age, with the highest prevalence in pregnant and postmenopausal patients.(1),(2).

Women experience lower UTIs (Urethral, Bladder infection) much more frequently than men do. The main reason for this is anatomical variations. In most cases, UTIs begin with per urethral contamination by a uropathogenic living in the gut, followed by urethral colonization and, lastly, pathogen migration to the bladder or kidney. When effective host defense mechanisms are defeated by bacterial virulence mechanisms, infections develop. When uropathogenic organisms climb to the kidneys via the ureters, upper UTIs (Kidney infections) result. (3). (4),(5).

Gram -ve and Gram +ve bacteria, as well as some fungi, are responsible for UTIs. Uropathogenic E. coli is the most frequent cause of UTIs. (6).

UTIs are classified based on (Level of Complicated, Site and Type of Infection, Symptoms of UTI) and the nature of the occurrence. (7),(8),(9).

The most common trigger of unusual vaginal fluid in women of sexually active age is (BV) and the most common cause of vaginal infection in pregnant and non-pregnant women. It is differentiated by a shortage in the number of lactobacilli which generate hydrogen peroxide and an increase in the pH level of the vagina.(10),(11),(12),(13),(14)(15).

SDF-1 is a chemokine protein that would be present in a range of organs and cell types. It's also encoded by the CXCL12 gene, which is found on human chromosome 10q11.1. TNF-induced pro-inflammatory stimuli further influence immune cell activation, migration, and regulation,(16),(17).

UTI and vaginal infection trigger an exciting protective immunity that is in charge of bacterial clearance that is quick and efficient. epithelial cells secretion of the chemokine stromal cell-derived factor 1, triggering the migration and accumulation of immune cells at the site of infection (18), (19),(20),(21),(22).

Immunoglobulin A (IgA) is an antibody which is generated by your immune system and is generated in greater quantities in conjunction with mucosal membranes than all other types of antibodies put together (70%). The most prevalent immunoglobulin in the blood and mucous secretions of the genitourinary tract is IgA. Performs important functions in immune defense against bodily secretion-based microbial invasion.,(23).

MATERIAL AND METHODS

Study Design: A group comparison study design was used, started in September 2022 with 180 participants divided into 3 groups (60 women with UTI and 60 women with BV) and 60 women healthy participants from the Imam AI-Sadiq General Teaching Hospital/ Gynecology Consultant/Babylon /Iraq, the three groups' demographic research criteria were taken (age(15-50Years), education, job, living, and laboratory tests.

Control and Patients: Patients with urinary tract infection and bacterial vaginosis were identified by a specialist in gynecology, according to the patient's signs and symptoms according to the laboratory tests evaluated. The Control group of women who participated in this study matched with patients in age, education, job, living, and laboratory tests, all women in the control groups were checked for signs and symptoms and laboratory tests and were within the normal range and apparently healthy. The medical information was gathered in Imam Al-Sadiq General Teaching Hospital/ Gynecology Consultant/Babylon /Iraq.

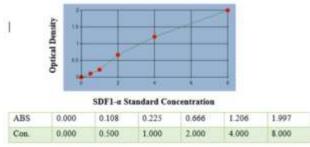


Figure1: Standard Curve for SDF1-a

Samples Collection and ELISA Assay: Blood collection from the vein of approximately (5ml) was obtained from patients and control, let clotting for 15 minutes and centrifugation at (3000xg) for10 minutes to obtain serum stored at -20° C. Determination of Human Stromal Cell Derived Factor 1 α and Immunoglobin A(IgA) Levels by ELISA kit supplied from Melsin Medical Co. applied to

the in vitro quantitative determination of Human Stromal Cell Derived Factor 1 α , IgA concentrations in serum. This ELISA kit used the Sandwich - ELISA principle at 450 nm.

The standard curve for human stromal cell-derived factor 1α and Immunoglobin A were obtained using the ELISA technique (Figures 1, 2).

Y=A[i]*x+B[i], correlation coefficient(R^2) =0.999

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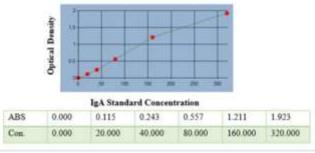


Figure 2: Standard Curve for IgA

Statistical Analysis: All statistical analyses were performed using software package version 28 (SPSS) data were presented as (mean + standard deviation) with 95% confidence intervals.

RESULTS AND DISCUSSIONS:

The medical information (Table 3-1) was collected for the patients and control including:

N=60 each group		UTI		Vaginal		Control		Sig
Mean+ Std.		30.433+9.59		30.283+9.26		34.100+10.50		*P.value
		Freq.	Perc	Freq	Perc.	Freq	Perc.	
Age groups	15-25 Y	21	35.0	20	33.3	19	31.7	0.08
1000000000000	26-36 Y	21	35.0	23	38.3	15	25.0	
	37-47Y	14	23.3	16	26.7	16	26.7	
	48-58 Y	4	6.7	1	1.7	10	16.7	
Education levels	Elementary	8	13.3	11	18.3	9	15.0	*0.001
	Intermediate	14	23.3	14	23.3	19	31.7	
	Academic	11	18.3	13	21.7	13	21.7	
	Institute	9	15.0	6	10.0	7	11.7	
	College	18	30.0	8	13.3	10	16.7	
	Master/PhD	0	0	8	13.3	2	3.3	
Job	Employee	20	33.3	15	25.0	25	41.7	*0.042
	Student	22	36.7	20	33.3	14	23.3	
	Housewife	18	30.0	25	41.7	21	35.0	
Living	Urban	29	48.3	30	50.0	37	61.7	*0.001
	Rural	31	51.7	30	50.0	23	38.3	

*(P.value ≤ 0.05) was significant

The distribution of patients according to 10 years' intervals was shown in table (3-1). It was obvious that the majority of UTI patients were in the age groups (15-25-26-36) years accounting for (35.0% for 15-25 years and,35.0% for 26-36 years), The majority of BV patients were in the age group (26-36) years accounting for (38.3%) in control subjects the highest rate with (15-25) years, this may be attributable to the fact that the age of onset of UTI, BV usually in young women, caused by frequent or recent sexual intercourse (increased sexual activity) is a major risk factor,(24) For UTI and (25),(26),(27) For BV.

According to the educational levels, for the UTI group, most of the had a college level (30%), for the BV group, most of them had an intermediate level (23.3%), and for the control group, most of them had intermediate level (31.7%),(28).

For the jobs, for the UTI group, most of them were students (36.7%), BV group most of them were housewives 41.7%, and 56.7% respectively. Finally, 41% of the control were employees (29).

For the living area, 51.7% of the UTI group was living in a rural area, 50% of the BV group was living in an urban area and the other half was in a rural area. 58.3% and control were living in urban areas respectively.

Table (3-2) SDF1 Alfa comparison among study groups

Groups	No.	Mean	Std. Deviation	Std. Error	Lower Bound	Upper Bound	"P.value	
UTI 60	1.206522	.0927277	.0119711	1.182568	1.230476	0.001		
BV	60	1.213735	.0661389	.0085385	1.196650	1.230820		
Control	.60	1.130013	.0496400	.0064085	1.117190	1.142837		

results showed that the concentrations of SDF-1 α in healthy controls as well as in patients with UTI (1.206522 ±.0927277 pg/ml) and patients with BV (1.213735 ±.0661389 pg/ml) were significantly greater (P.value = 0.001) than that of normal controls (1.130013±.0496400 pg/ml).

SDF-1 α is one of the chemokines responsible for attracting and accumulating T lymphocytes (30),(31), monocytes as well as responsible for the activation, adhesion, and migration of neutrophil leukocytes to inflammatory sites (21) , (22). Elevated production of SDF-1 has been shown to be associated with some of infectious and inflammatory diseases (32),(33) ,(34). (16) that says SDF-1 is secreted shortly after infection of UTI and most significantly elevated during infection of UTI also elevated in the Reproductive system .

Table (3-3) IgA comparison among study groups

Groups	No.	Mean	Std. Deviation	Std. Error	Lower Bound	Upper Bound	*P,value	
UTI	60	64.252983	7.5946759	.9804684	62.291071	66.214896	0.001	
BV	60	60.441928	6 1457661	7934150	58 854309	62.029548		
Control	.60	48.011745	4.7938613	6188848	46.773369	49.250131		

The study's findings of women with UTI have the elevation of IgA (64.252983 \pm 7.5946759pg/ml) and patients with BV (60.441928 \pm 6.1457661pg/ml) compared with those in normal controls (48.011745 \pm 4.7938613 pg/ml), according to the statistical test, the comparison among means reveals that there is highly statistically significant difference between them (0.001).

Immunoglobulin A (IgA) is the most abundant type of antibody 70% in the body compared with other types of immunoglobin, it serves to protect the mucosal tissues from microbial invasion by preventing bacterial adherence to per urethral epithelia and uroepithelia linings of the genitourinary tracts (23).

Conclusion

SDF1-Alpha and IgA were used as diagnostic tools to reveal the Urinary tract infection and Bacterial vaginosis. Elevated levels of SDF1 and IgA act to protect the mucosal tissues from microbial invasion by preventing bacterial adherence to per urethral epithelia and uroepithelia linings of the genitourinary tracts and attracting and accumulating, activation, adhesion, and migration of immune cells to infection/inflammatory sites.

Recommendation: Compare between the gene of stromal cellderived factor 1 alpha in women with UTI and women with BV by polymerase chain reaction.

Ethical Approval: The study was carried out after obtaining the approvals of the patient and the Iraqi Ministry of Health. **Study Conflict:** there are no studies conflicts.

REFERENCES

- Czajkowski K, Broś-Konopielko M, Teliga-Czajkowska J. Urinary tract infection in women. Prz Menopauzalny. 2021;20(1):40–7.
- Almukhtar SH. Urinary Tract Infection Among Women Aged (18-40) Years Old in Kirkuk City, Iraq. Open Nurs J. 2019;12(1):248–54.
- 3. Consequences H, Improvement P, Kit T, Information P. In theClinic

Osteoporosis. 2013;

- Beahm NP, Nicolle LE, Bursey A, Smyth DJ, Tsuyuki RT. The assessment and management of urinary tract infections in adults: Guidelines for pharmacists. Can Pharm J. 2017;150(5):298–305.
- Geerlings SE. Clinical Presentations and Epidemiology of Urinary Tract Infections. Microbiol Spectr. 2016;4(5):27–40.
- Totsika M, Gomes Moriel D, Idris A, A. Rogers B, J. Wurpel D, Phan M-D, et al. Uropathogenic Escherichia coli Mediated Urinary Tract Infection. Curr Drug Targets. 2012;13(11):1386–99.
- Smelov V, Naber K, Bjerklund Johansen TE. Improved Classification of Urinary Tract Infection: Future Considerations. Eur Urol Suppl [Internet]. 2016;15(4):71–80. Available from: http://dx.doi.org/10.1016/j.eursup.2016.04.002
- Gajdács M, Ábrók M, Lázár A, Burián K. Urinary tract infections in elderly patients: A 10-year study on their epidemiology and antibiotic resistance based on the who access, watch, reserve (aware) classification. Antibiotics. 2021;10(9).
- 9. Angeles L. I /. 1995;196:1-8.
- Russo R, Karadja E, De Seta F. Evidence-based mixture containing Lactobacillus strains and lactoferrin to prevent recurrent bacterial vaginosis: A double blind, placebo controlled, randomised clinical trial. Benef Microbes. 2019;10(1):19–26.
- Achondou AE, Fumoloh FF, Aseneck AC, Awah AR, Utokoro AM. Prevalence of bacterial vaginosis among sexually active women attending the CDC central clinic tiko, South West Region, Cameroon. African J Infect Dis. 2016;10(2):96–101.
- Bhavana AM, Kumari PHP, Mohan N, Chandrasekhar V, Vijayalakshmi P, Manasa RV. Bacterial vaginosis and antibacterial susceptibility pattern of asymptomatic urinary tract infection in pregnant women at a tertiary care hospital, Visakhaptn, India. Iran J Microbiol. 2019;11(6):488–95.
- French AL, Adeyemi OM, Agniel DM, Evans CT, Yin MT, Anastos K, et al. Bacterial Vaginosis and Vitamin D Deficiency Among Nonpregnant HIV-Infected and Uninfected Women in the United States. J Womens Health (Larchmt) [Internet]. 2011;4(4):36–40. Available from: http://www.ncbi.nlm.nih.gov/pubmed/21876923
- Mendling W, Shazly MA El, Zhang L. The Role of Lactic Acid in the Management of Bacterial Vaginosis: A Systematic Literature Review. Futur Pharmacol. 2022;2(3):198–213.
- Joyisa N, Moodley D, Nkosi T, Talakgale R, Sebitloane M, Naidoo M, et al. Asymptomatic bacterial vaginosis in pregnancy and missed opportunities for treatment: A cross-sectional observational study. Infect Dis Obstet Gynecol. 2019;2019.
- Isaacson B, Hadad T, Glasner A, Gur C, Granot Z, Bachrach G, et al. Stromal Cell-Derived Factor 1 Mediates Immune Cell Attraction upon Urinary Tract Infection. Cell Rep [Internet]. 2017;20(1):40–7. Available from: http://dx.doi.org/10.1016/j.celrep.2017.06.034
- Mosleh AL-Kafajy MN, Abdul-Razzaq MS. Study of gene polymorphism for stromal cell-derived factor 1 (SDF1/XCL12) and its elevation in plasma of women of reproductive age with pelvic inflammatory diseases in Babylon province. Ann Trop Med Public Heal. 2020;23(138).
- Abraham SN, Miao Y. The nature of immune responses to urinary tract infections. Nat Rev Immunol [Internet]. 2015;15(10):655–63. Available from: http://dx.doi.org/10.1038/nri3887
- Choi HW, Bowen SE, Miao Y, Chan CY, Miao EA, Abrink M, et al. Loss of Bladder Epithelium Induced by Cytolytic Mast Cell Granules.

Immunity [Internet]. 2016;45(6):1258–69. Available from: http://dx.doi.org/10.1016/j.immuni.2016.11.003

- Schiwon M, Weisheit C, Franken L, Gutweiler S, Dixit A, Meyer-Schwesinger C, et al. Crosstalk between sentinel and helper macrophages permits neutrophil migration into infected uroepithelium. Cell. 2014;156(3):456–68.
- Tecchio C, Cassatella MA. Neutrophil-derived chemokines on the road to immunity. Semin Immunol [Internet]. 2016;28(2):119–28. Available from: http://dx.doi.org/10.1016/j.smim.2016.04.003
- David BA, Kubes P. Exploring the complex role of chemokines and chemoattractants in vivo on leukocyte dynamics. Immunol Rev. 2019;289(1):9–30.
- Corthésy B. Multi-faceted functions of secretory IgA at mucosal surfaces. Front Immunol. 2013;4(JUL):1–11.
- Dubois B, Kobelt G, Berg J, Capsa D, Gannedahl M. New insights into the burden and costs of multiple sclerosis in Europe: Results for Belgium. Mult Scler J. 2017;23(2_suppl):29–40.
- Chow K, Wooten D, Burke L, Edi R, Morris SR. Impact of (Recurrent) Bacterial Vaginosis on Quality of Life and the Need for Accessible Alternative Treatments. 2022;1–9.
- Reiter S, Kellogg Spadt S. Bacterial vaginosis: a primer for clinicians. Postgrad Med [Internet]. 2019;131(1):8–18. Available from: https://doi.org/10.1080/00325481.2019.1546534
- Madhivanan P, Krupp K, Chandrasekaran V, Karat C, Arun A, Cohen C, et al. Prevalence and Correlates of Bacterial Vaginosis Among Young Women of Reproductive Age in Mysore, India. Indian J Med Microbiol [Internet]. 2008;26(2):132–7. Available from: https://doi.org/10.1016/S0255-0857(21)01928-9
- Lelie-van der Zande R, Koster ES, Teichert M, Bouvy ML. Womens' self-management skills for prevention and treatment of recurring urinary tract infection. Int J Clin Pract. 2021;75(8):1–7.
- Javaheri Tehrani F, Nikpour S, Haji Kazemi EA, Sanaie N, Shariat Panahi SA. The effect of education based on health belief model on health beliefs of women with urinary tract infection. Int J community based Nurs midwifery [Internet]. 2014;2(1):2–11. Available from: http://www.ncbi.nlm.nih.gov/pubmed/25349840%0Ahttp://www.pubm edcentral.nih.gov/articlerender.fcgi?artid=PMC4201186
- Janssens R, Mortier A, Boff D, Vanheule V, Gouwy M, Franck C, et al. Natural nitration of CXCL12 reduces its signaling capacity and chemotactic activity in vitro and abrogates intra-articular lymphocyte recruitment in vivo. Oncotarget. 2016;7(38):62439–59.
- Nerviani A, Pitzalis C. Role of chemokines in ectopic lymphoid structures formation in autoimmunity and cancer. J Leukoc Biol. 2018;104(2):333–41.
- Shao S, Cai W, Sheng J, Yin L. Role of SDF-1 and Wnt signaling pathway in the myocardial fibrosis of hypertensive rats. Am J Transl Res. 2015;7(8):1345–56.
- Palma J, Tokarz-Deptuła B, Deptuła J, Deptuła W. Natural antibodies – Facts known and unknown. Cent Eur J Immunol. 2018;43(4):466– 75.
- Chalin A, Lefevre B, Devisme C, Pronier C, Carrière V, Thibault V, et al. Serum CXCL10, CXCL11, CXCL12, and CXCL14 chemokine patterns in patients with acute liver injury. Cytokine [Internet]. 2018;111(May):500–4. Available from: https://doi.org/10.1016/j.cyto.2018.05.029