Relationship between Leukotriene B4 (LTB4) and the pathogenicity of Trichomonas vaginalis
Susan Abbas Abdul Al-Ammer1, Liqaa Hameed Hasan1, Kawther A.M. Al-Mussawi2, Huda Abdullah3, Inam Joudah Radhi*1
1Obstetrics and Gynecology in hospital of Kerbala, Iraq.
2Professor, Department of Biology, College of Education for Pure Science, University of Kerbala, Kerbala, Iraq.
3Kerbala, Iraq.
*Corresponding author: Inam Joudah Radhi, Department of chemistry, Collage of Education for Pure Science, University of Kerbala, Kerbala, Iraq, Email: anaam.j@uokerbala.edu.iq

Submitted: 17 January 2023; Accepted: 18 February 2023; Published: 19 March 2023

ABSTRACT
Trichomoniasis, which is brought on by the parasite Trichomonas vaginalis, is the most prevalent non-viral, curable worldwide sexually transmitted disease that affects millions of individuals every year. Numerous immunological and biochemical elements were discovered to be involved in the pathogenesis of T. vaginalis. In our study, we sought to determine how T. vaginalis virulence and leukotriene B4 (LTB4) interact. 350 women (15–65 year) who visited a women's and gynecological hospital in Karbala between November 2021 and June 2022 for this study provided swabs of vagina, samples of urine and samples of blood took place. Routine urine analysis and examination of by direct microscopic to vaginal swabs were used to identify parasites, and its ELISA approach was used to calculate serum (LTB4) levels. Participating women filled out a questionnaire that asked about their age, place of residence, marital status, degree of education, reproductive potential, and use of contraception. As a result, only 100 of the 350 cases tested positive for T. vaginalis, with an age group (15–24 year) having the highest frequency and the age group 65+ having the lowest incidence (15-24). 55 years). By educational background, 18 women (28.57%) were illiterate, 22 (29.33%) in primary education, 42 (28.18%) in secondary education, and 18 (28.57%) in bachelor level. They did not differ significantly from one another (P=0.998). In healthy, non-infected women, the mean and standard deviation of serum (LTB4) were 19.14 and 7.14, respectively, but in patient-infected women, they were (96.41 and 25.9). did. (P 0.001) Significant difference. We can infer from the current study that women who had T. vaginalis infections had significantly higher serum levels of LTB4.

Keywords: Trichomonas vaginalis, Trichomoniasis, Leukotriene B4

INTRODUCTION
A significant worldwide health issue is sexually transmitted infections (STIs). Sexually transmitted illness like Chlamydia, Gonorrhea, Syphilis, genital warts and trichomoniasis are thought to affect 500 million people annually.
Additionally, her HSV2 has infected over 530 million people, and her HPV has infected over 290 million women [1].

Through sexual contact, T. vaginalis is passed from one person to another. The trophozoite of this parasite clings to the mucosal surface of the urogenital canal during its life cycle and divides via longitudinal binary fission. T. vaginalis successfully colonizes the host mucosa by a variety of pathogenic mechanisms, including adhesion, interaction with the vaginal microbiome, evasion and control of the host immune system, production of cytotoxic chemicals and soluble substances, and the initiation of immunological responses [2].

Trichomoniasis, a neglected STI with a prevalence of 110.4 million cases and an incidence of 156 million, is brought on by the flagellated protozoan Trichomonas vaginalis. [3,4]. According to the World Health Organization’s (WHO) most recent estimates, the African region has the largest incidence of trichomoniasis, which is followed by the Americas, Western Pacific, Eastern Mediterranean, Southeast Asia, and finally the European region. Region [5]. Although the majority of cases are asymptomatic, reports of itching, discharge, irritation, and odor persist. Infertility, pelvic inflammatory illness, cervical and prostate cancer, as well as premature birth and low birth weight in babies are all linked to long-term infection with T. vaginalis, which can last for months to year [6,7]. In addition, there is evidence of a reciprocal association between T. vaginalis infection and human immunodeficiency virus (HIV) transmission and acquisition, with people of T. vaginalis infection 1.5 times more likely than healthy individuals to acquire HIV [8].

Leukotrienes (LTs) are lipid mediators produced by the 5-lipoxygenase (5-LO) passageway during the metabolism of arachidonic acid. Leukotriene B4 (LTB4) is created when 5-LO and LTA4 hydrolase act together on LTA4 [9]. Experimental infections caused by microorganisms in vivo and in vitro interact with phagocytes to produce LTs [10]. LTs play essential roles in both innate and adaptive immune responses [11], [12]. Through the modulation of both Th1 and Th2 immune reactions, LTB4 are implicated in the defense against protozoan and helminthic illnesses [13]. Nitric oxide (NO) and cytokine generation during infection is the mechanism through which LTB4's antimicrobial action [14], antiparasitic activity [15], and antifungal activity occur. [16]. The phagocytic and antibacterial capabilities against microorganisms, including parasites, will be reduced in the presence of any abnormality in LTB4 biosynthetic pathways.[14] [16] bacteria [17] and fungi [15].

Our earlier research demonstrated that T. vaginalis LTB4 release might draw in and stimulate neutrophils. [18]. We also discovered elevated amounts of LTB4 in vaginal discharges from patients with symptomatic trichomoniasis in a preliminary research. [19]. LTB4 also causes a chemotactic response and a number of other reactions, including aggregation, degranulation, and oxidative metabolism, all of which aid in the eradication of invader species [20]. Since neutrophils are known to produce LTB4 and interact with T. vaginalis during the inflammatory reaction, [21]. A sizable concentration of LTB4 is conceivably expected to liberate and heighten host defenses against T. vaginalis. So, in this investigation, we looked into how humoral immunity might control the formation of LTB4 by neutrophils in response of stimulation of T. vaginalis.

MATERIALS AND METHODS
In the current search, swabs of vagina, samples of urine, and samples of blood were collected from 350 women who attended the women's obstetrics and gynecologist hospital in Karbala province between November 2021 and June 2022. The women's ages ranged from 15 to 65. The parasite was found using a comprehensive urine examination and a direct microscopic examination of vaginal swabs, while the serum (LTB4) level was determined using the ELISA approach. The participant women were asked a series of questions on their age, place of residence, marital status, degree of education, fertility, and use of contraceptives.

The samples of venous blood were drawn into anticoagulant-free gel tubes, allowed to clot for...
15 minutes at room temperature, and then centrifuged for 10 minutes at 3000 rpm to obtain serum. Serum LTB4 levels were measured using the ELISA method, which uses enzyme-linked immune sorbent assays. The presence of the parasite T. vaginalis was determined using both a general urine test and a direct microscopic inspection of the vaginal swabs.

**The Statistical Tests**
The SPSS version 25 software was used to analyze the statements. For variances, the t-test as well as Chi square were used, P 0.05 was regarded like respectable.

**The Results**
Only 100 (28.57%) of the 350 women in this study tested positive for the parasite T. vaginalis, while 250 (71.42%) tested negative, as shown in the table (1).

**TABLE 1**: shows the number and percentage of positive cases based on microscopic analysis.

<table>
<thead>
<tr>
<th>Total No. of examined samples</th>
<th>Positive Samples(+)</th>
<th>Negative Samples(-)</th>
</tr>
</thead>
<tbody>
<tr>
<td>350</td>
<td>100</td>
<td>250</td>
</tr>
<tr>
<td>No.</td>
<td>%</td>
<td>No.</td>
</tr>
<tr>
<td>28.57</td>
<td>71.42</td>
<td></td>
</tr>
</tbody>
</table>

The results in table (2) showed The relationship between positive samples with the level of fertility, and the table showed that the number of positive samples was 28/12 (35.71%) in infertile cases, while the table showed that the number of samples in cases without infertility was 90/322 (27.95%). There are no significant differences (P=0.383).

**TABLE 2**: Positive samples distributed according to fertility

<table>
<thead>
<tr>
<th>P-Value</th>
<th>Positive Cases(+)</th>
<th>Total Number</th>
<th>Fertility</th>
</tr>
</thead>
<tbody>
<tr>
<td>%</td>
<td>No.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>0.383*</td>
<td>35.71</td>
<td>10</td>
<td>28</td>
</tr>
<tr>
<td>27.95</td>
<td>90</td>
<td>322</td>
<td>No</td>
</tr>
</tbody>
</table>

As for the relationship between infection with vaginal trichomoniasis and cases of abortion, the table below indicates the incidence of infection, which was (34.09%) for non-abortion women and (12.35%) for aborted women. The current study found significant differences at the level of likelihood (P > 0.05). There are significant differences at the level of probability 0.05(P=0).

**TABLE 3**: Distribution of parasite infection according to abortions

<table>
<thead>
<tr>
<th>P-Value</th>
<th>Positive Cases(+)</th>
<th>Total Number</th>
<th>Abortions(Abs.)</th>
</tr>
</thead>
<tbody>
<tr>
<td>%</td>
<td>No.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>0*</td>
<td>12.35</td>
<td>11</td>
<td>89</td>
</tr>
<tr>
<td>34.09</td>
<td>89</td>
<td>261</td>
<td>No</td>
</tr>
</tbody>
</table>

According to the results in table (4), there was a highly significant difference (P 0.001) between the mean and standard deviation of serum LTB4 levels in the healthy women group (19.147.14) and the infected patient women group (96.4125.9).
**TABLE 4:** The Mean ±SD in healthy, LTB4 levels and infected women

<table>
<thead>
<tr>
<th>The Groups</th>
<th>Mean ± SD</th>
</tr>
</thead>
<tbody>
<tr>
<td>The Healthy women</td>
<td>19.14±7.14</td>
</tr>
<tr>
<td>The Patients</td>
<td>96.41±25.9</td>
</tr>
<tr>
<td>Calculated P-Value</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

**FIGURE 1:** Display of the ELISA-generated LTB4 concentration estimate curve

**DISCUSSION**

The epidemiology of T. vaginalis infection takes into account factors including age, living place, socioeconomic status, pedagogics, marital status, form of contraception used, existence as well as type of vaginal secretions, medicines taken, and history of other STDs. Numerous risk factors may have an impact on infections [22].

According to the study's findings, there are noticeable distinctions between infection and abortion, as the highest infection rate was recorded among women who did not suffer from abortion, amounting to 34.09%, compared to 12.35% for aborted women. 2008. And Al-Kazragee (2013) recorded the highest infection rate among women with normal pregnancies (non-abortions), amounting to 5.82%, compared to aborted women, 4.67%, and the results of this study were inconsistent with some studies, such as the study conducted by Al-Ziyadi (2004) in Najaf, when an infection was recorded High in aborted women, with a rate of 19.76%, compared to women with normal pregnancies, 19.4% also agreed with Al-Khatawy (2012) . The highest rate of infection was recorded among women with normal pregnancies, at a rate of 10%, which is higher than that recorded for aborted women by 6.67%. It was found that the parasite Trichomoniasis is able to pass into the Fallopian tubes, carrying many pathogenic organisms that cause infections and infertility (Lujan et al, 2022).

As for the occurrence of miscarriage, it is attributed to many reasons, including those that are embryonic causes. It is possible that infection of the placenta leads to inflammation of the membranes surrounding the fetus and the release of the prostaglandin hormone, or miscarriage may occur as a result of the early activity of the uterus (Terzic & Aimagambetova et al, 2021)

The current study's findings also showed that there were notable disparities in the distribution of parasite infection between pregnant and non-pregnant women, with the highest infection rate among non-pregnant women being 35.71% and the infection rate among pregnant women being 27.95%. The rate of infection among non-pregnant women was 25.79% higher than that of pregnant women at 13.69%, also in agreement with Al-Saeedi (2016). The rate of parasite infection among non-pregnant women was more than that of pregnant women. The reason for the increase in the infection rate in non-pregnant
women is attributed to the abnormal conditions of the vagina, as well as the person’s health level, and that pregnant women may not have the effectiveness for the occurrence of the disease, and the occurrence of infection in those women may be attributed to the change that occurs to the environment of the vagina from the conditions of acidity that change Animal growth from Moniliasis to Trichomoniasis. The appearance of the infection in pregnant women may be due to the hormonal changes that occur during pregnancy and the enlargement that occurs in the vaginal epithelium, as well as the high level of glycogen and estrogen, and this helps in providing a suitable medium for parasite growth and reproduction (Studd, 2008).

According to Dahab et al. (2012) and Al-Ibrahim (2008), non-pregnant women had the greatest infection rates, 5.6% and 13.3%, respectively. This result supports the need for routine clinical examinations for the early detection and treatment of these diseases in women. Pregnant women are advised to visit health units regularly for a pregnancy check-up (Adeoye & Akende, 2007).

According to the results of the current investigation, there was a highly significant difference in the level of LTB4 between the trichomoniasis infected women and the healthy control women. The percentage of the LTB4 immune criterion in the serum of infected women was (19.14 7.14) compared to control women. This is consistent with a rise in leukotriene as a result of parasite infection, which led to the development of immunity in the lining tissues. The vagina contains a large number of monocytes and thus produces large amounts of LTB4. The findings of this study are consistent with a study by Eida et al. (2015), which found that women with symptoms of T. vaginalis infection had higher levels of the leukotriene LTB4. LTB4 is a leukotriene stimulating factor that T. vaginalis secretes. Nemati et al. (2018) show that through the interaction of the leukotriene LTB4 with the leukotriene receptors BLT of host cells, T. vaginalis has the capacity to communicate directly with immune cells, This modification of the host immune response and activation of neutrophils is the result of this communication. As of Min et al.(2017), These results also correlate well with previous animal studies where leukotrienes LTB4 has been demonstrated to be increased in animals infected with other nematodes and other parasites such as. Entamoeba histolytica (Jimenez et al., 2021).

REFERENCES


J Popul Ther Clin Pharmacol Vol 30(5):e526–e531; 19 March 2023. This article is distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 International License. ©2021 Muslim OT et al. e530


