Assessment of Serum Fatty Acid Binding Protein 4 in Patients with Acne Vulgaris

A.A Ibrahim, A.I.Mustafa, O.S.El-Shimi and A.A.Abd El Salam

Dermatology, Venereology and Andrology Dept., Faculty of Medicine, Benha Univ., Benha, Egypt

E-Mail:ahmed2356@gmail.com

Abstract

Unsaturated fat restricting proteins (FABP) are a gathering of cytosolic lipid transporters which facilitate incendiary and metabolic pathways in cells. The point of this momentum study was to gauge serum levels of FABP4 in patients with skin break out vulgaris and to survey connection between's its level and seriousness of the malady. This investigation included 50 patients with mellow (17 patients), moderate (17 patients) and extreme (16 patients) skin break out vulgaris as indicated by Global Acne Grading System All contemplated patients were exposed to history taking, dermatological assessment and estimation of serum FABP4. The aftereffects of this case–control study showed that the mean serum levels of FABP4 was essentially higher in AV patients. Mean FABP4 levels demonstrated non-noteworthy distinction between various types of severity.FABP4 appears to have a fundamental function in the etiopathogenesis of this fiery skin malady, So it is viewed as a promising objective in treating AV and other provocative infections.

1. Introduction

Normal connections between vulgaris skin break out and its comorbidities fundamentally include the interminable aggravation that rises up out of metabolic tissues (metaflammation), hereditary premise, comparable pathways of safe issues, and bioactive substances orchestrated and discharged by fat tissue. Various distributed examinations have shown the possible function of different adipokines in the advancement of skin break out and its comorbidities [1].

Unsaturated fats, which establish about 15% of lipids in the layer corneum, take an interest in keeping up the penetrability of the epidermis, for instance, by invigorating fermentation of the horny layer. Exploration has demonstrated that deviations in FA organization in keratinocytes might be associated with the pathogenesis of incendiary dermatosis, for example, atopic dermatitis or psoriasis. FA moved to long-chain unsaturated fats (LCFA) are shipped by specific proteins to different tissues, where they are processed, put away or used [2].

Adipocyte unsaturated fat restricting protein (FABP4), likewise named adipocyte protein 2 (aP2), is an adipokine blended and delivered dominantly from adipocytes and, less significantly, created in macrophages and endothelial cells [3].

FABP4, with a sub-atomic mass of 14.6 kDa and 132 amino acids, represents up to 6% of complete cell proteins. The quality encoding this adipokine is situated on chromosome 8q21. Raised degrees of FABP4 are firmly connected with the advancement of stoutness, IR, DM, hypertension and atherosclerosis [4].

Studies in people have indicated that a hereditary variation at the FABP4 locus decreased the danger for atherosclerosis, diabetes and coronary illness, and diminished FABP4 articulation in fat tissue was identified with lower hazard for coronary vein infection (CAD) and DM. A few investigations have demonstrated that expanded circling FABP4 levels are a promising biomarker foreseeing cardiovascular breakdown, stroke, and guess and mortality in patients with end-stage renal infection, CAD or other basic diseases [5].

Another conceivable function of FABP4 in skin inflammation is clear in the effect on metabolic condition, which is unequivocally connected with skin inflammation [6]. FABP4 is an autonomous prescient marker for the improvement of metabolic condition, as was appeared in a 5-year forthcoming investigation and in an investigation of Korean young men [7]. In this way, FABP4 may assume a urgent part in various issues, including heftiness, atherosclerosis, metabolic disorder and IR, which are firmly connected with skin break out. The point of the current examination was to survey serum FABP4 level in patients with skin break out vulgaris.

2. Patient and method

A case-control study was directed on 50 patients experiencing AV (Group A) and 30 evidently sound people of coordinated age and sex as a benchmark (Group B). Patients were selected from the outpatient center of Dermatology and Andrology Department of Benha University Hospitals between September 2018 and March 2019.

The examination was affirmed by the nearby morals board of trustees of Benha Faculty of Medicine. Educated assent was gotten from every person before test assortment.

Inclusion criteria
- Patients ≥ 18 years.
- Both gender.
- Patients with acne who didn’t receive topical or systemic treatment in the last 1 months.

Exclusion criteria
- Patients under any dietary restrictions.
- Patients who received topical or systemic treatment in the last two months.
- Patients with dyslipidemia.
- Obese patients.
- Patients with inflammatory disorders.
All patients were subjected to the followings

1. Full history taking including
   - Personal history: name, age, sex, occupation, residence, special habits of medical importance and marital status.
   - Present history: duration of AV, relation to diet, relation to stress, relation to sun exposure, previous treatment, as well as history of other skin diseases.
   - Family history of AV and post acne scarring.
   - Past history: history of medications (type, dose and duration), contraceptive or hormones, associated autoimmune diseases.

2. Clinical examination
   - Full general examination to prohibit related fundamental maladies, for example, diabetes mellitus and other provocative sicknesses.
   - The weight file (BMI) was determined as weight/(height)^2 (kg/m^2)
   - Detailed dermatological examination; to assess clinical variation, seriousness of AV and presence of skin break out scars.
   - Diagnosis of AV was made based on the patient's history and the regular clinical highlights of non-fiery injuries including shut (whiteheads) and open (clogged pores) comedones, and provocative sores including papules, pustules, knobs as well as blisters on the face, chest, shoulders and upper back.
   - Patients were isolated by GAGS into three gatherings, mild (17 patients), moderate (17 patients) and serious (16 patients). The total severity score was nderived from the summation of six regional subscores. Each was derived by multiplying the factor for each region; (factor for forehead and each cheek was 2, chin and nose was 1 for each and chest and upper back was 3 for each); by the most heavily weighted lesion within each region (1 for comedones, 2 for papules, 3 for pustules and 4 for nodules). A score of (1-18) is considered mild; (19-30) moderate; (31-38) severe; and > 39 very severe.

3. Laboratory investigations
   All studied subjects were tested for serum level of FABP4.

3. Results
   There were no significant differences between both groups as regard age (P value = 0.073), gender (P value = 0.948), smoking (P value = 0.628) and BMI (P value = 0.720).

   Of the cases group, 26 patients (52%) experienced sudden onset while 24 patients (48%) experienced gradual onset. As regard disease course, 23 patients (46%) showed progressive course, 21 patients (42%) showed fluctuating course and 6 patients (12%) showed stationary course. Median disease duration was 1 year and ranged from 2 weeks to 10 years.

   The most frequent affected site was face (100.0%) followed by back (44.0%), chest (36.0%) and the least frequent was shoulder (14.0%).

   Of the cases group, 3 patients (6.0%) had scars. 44 patients (88.0%) showed positive relation to stress. None of the cases group showed positive relation to sun exposure or diet.

   6 patients (12.0%) of cases group showed positive history of treatment. Of them, one patient (16.7%) received only topical treatment and 5 patients (83.3%) received topical and systemic treatment. 24 patients (48.0%) of cases group had positive family history. None of cases had positive history of systemic disease.

   Mean serum FABP4 level was significantly higher in cases (7.64) compared to controls (3.74) (P value was <0.001).

   ROC analysis was done for FABP4 in diagnosis of acne vulgaris. It showed significant Area Under Curve (AUC) of 0.906 which indicating excellent performance in diagnosing acne vulgaris (P value <0.001). 95% confidence interval ranged from 0.844 to 0.967. Best cutoff point for differentiation between cases and controls was > 6.38 at which sensitivity and specificity were 68.0%, 100.0% respectively.

   There were no significant correlations between FABP4 and age, BMI and duration (P values were 0.065, 0.57 and 0.334 respectively).

   Mean FABP4 showed non-significant difference between different forms of severity (P value was 0.069).

   Mean FABP4 showed non-significant difference between males and females (P value was 0.816).

   Mean FABP4 showed non-significant difference as regard different sites affected including back (P value = 0.097), Chest (P value = 0.312) and shoulder (P value = 0.166).

   Mean FABP4 showed non-significant difference between patients with positive relation to stress and those without (P value was 0.053).

   Mean FABP4 showed non-significant difference as regard history of treatment and family history (P values were 0.530 and 0.193 respectively).

   Multivariate logistic regression analysis was done for prediction of acne vulgaris. FABP4 was a significant predictor with Odds ratio of 2.606 and 95% confidence interval ranging from 1.684 to 4.032 (P value was <0.001).

   17 patients (34.0%) showed mild form severity. The same for moderate form while 16 patients (32.0%) of patients showed severe form of the disease.
4. Discussion

As indicated by the aftereffects of the current investigation, there was a measurably huge increment in serum FABP4 level among cases contrasted with control gathering.

As AV is considered as one of the incendiary sicknesses [8], the aftereffects of this investigation come in concurrence with comparative examinations performed on other provocative illnesses that affirm this idea. For instance, these outcome concurred with the examination led by Baran [9]; they announced that Serum FABP4 levels were altogether expanded in patients with psoriasis, showing that this protein might be a marker of psoriasis and an autonomous indicator of the danger of comorbidities or confusions in psoriatic patients.

As far as we could possibly know, no distributed examinations were found to think about the outcomes between AV patients concerning serum FABP4 levels.

The function of FABP4 in AV can be clarified in a few different ways, AV is described by interminable irritation of pilosebaceous unit [10], and has been recognized as an individual from the group of metabolic infections, for example, weight, T2DM, and disease [11] since heftiness influences sebaceous organs and sebum creation. Hence we could bring up a skin inflammation stoutness relationship. Skin inflammation is unmistakably bothered by stoutness related issues, for example, hyperandrogenism and hirsutism. Likewise as an indication of seborrhea-acne-hirsutism-androgenetic alopecia (SAHA) and hyperandrogenism-insulin obstruction acanthosis nigricans (HAIR-A) disorder. Additionally Previous reports recommended that FABP4 can disturb the seriousness of provocative related malady by expanding the degrees of chemokines, for example, the tumor rof factor-α (TNF-α) [12].

Rising proof has shown that serum FABP4 levels are decidedly connected with markers of the metabolic disorder and that an expansion in serum levels of FABP4 at standard predicts the danger for metabolic horribleness and mortality. These discoveries propose that circling FABP4 got from adipocytes is a helpful biomarker to gauge flow status of metabolic ailments and foresee their occurrence later on [13].

Xu [14] gave a clinical proof showing that expanded coursing FABP4 levels are related with stoutness, insulin obstruction, type 2 diabetes, hypertension, heart brokenness, and atherosclerosis, Also IR play arole in skin inflammation pathogenesis because of increment insulin like development factor1 whichstimulate 5α-reductase, adrenal and gondal androgen synthesis,androgen receptor signal transduction, sebocytes expansion and lipogensis [15].

Fig (1) Serum FABP4 according to disease severity.

Fig (2) ROC analysis of FABP4 in diagnosing acne vulgaris.
and the comedogenic impacts of IGF-1 and high androgen levels are believed to be liable for the acne [16].

In spite of the fact that the function of FABP4 in metabolic illnesses and irritation is valued, its inclusion in the pathogenesis of skin inflammation has not been very much explained. Be that as it may, a line of proof recommends the pathogenic function of FABP4 in AV. For instance, previous reports recommended that FABP4 can irritate the seriousness of provocative related infection by expanding the degrees of chemokines, for example, the tumor rot factor-α (TNF-α) [12], TNFα likewise assumes a significant part in numerous phases in the pathogenesis of AV, and is considered as one of the proinflammatory cytokines that is fundamentally significant in the regulation of MMP enactment in the dermis during skin break out aggravation [17].

Likewise, FABP4 can reversibly tie to hydrophobic ligands, for example, soaked and unsaturated long-chain unsaturated fats (FA) [18], which are viewed as the primary creation of sebum [17]. Sebum overproduction in turns causes improvement of skin break out lesions [17].

Expanded degrees of FABP4 are likewise portrayed for other interminable provocative conditions. FABP4 was additionally answered to be raised in serum of psoriasis vulgaris patients. Interestingly, this illness imparts a few highlights to skin inflammation. For instance, TNF α is one of the focal cytokines in the pathophysiology of psoriasis. It is likewise assumed a significant function in numerous phases in the pathogenesis of AV [19].

In this investigation ROC examination was accomplished for FABP4 in determination of skin break out vulgaris. It demonstrated noteworthy Area Under Curve (AUC) of 0.906 which showing magnificent execution in diagnosing skin inflammation vulgaris (P esteem <0.001). 95% certainty stretch extended from 0.844 to 0.967. Best cutoff point for separation among cases and controls was > 6.38 at which affectability and explicitness were 68.0%, 100.0% separately.

Steady with the past discoveries, it isn’t amazing that this examination discovered expanded serum levels of FABP4 in AV, which is because of the presence of these exorbitant fiery cells in AV which additionally a significant wellspring of lipocalin 2 creation [20].

All in all, FABP4 appears to have a basic part in the etiopathogenesis of provocative skin infections including AV. So it is viewed as a promising objective in treating incendiary maladies [21].

5. Conclusion

From the results of the present work, it was concluded that; serum FABP4 levels are higher among AV patients but there was insignificant difference between different forms of severity, and it seems to have an essential role in the etiopathogenesis of inflammatory skin disease. So it is considered to be a promising target in treating inflammatory diseases.

References

[14] A. Xu, A. W. Tso, B. M. Cheung. Circulating adipocyte-fatty acid binding protein levels predict the development of the metabolic syndrome: a 5-


