Evaluation of Serum Level of Salusin Beta in Psoriatic Patients

S.H. Ahmed1, E.M. Akil1, N.A. Abdul Hafez2, E.E. Shehata 1 and S.M. Abd El Aal 1

1 Dermatology, Venereology and Andrology Dept., Faculty of Medicine, Benha Univ., Benha
2 Clinical Pathology and Immunology Dept., Faculty of Medicine, Benha Univ., Benha, Egypt
3 Cardiology Dept., Faculty of Medicine, Ain Shams Univ., Egypt
E-Mail: Shady@gmail.com

Abstract
Psoriasis is a chronic, immune-mediated, inflammatory disease of the skin. It is a multifactorial malady with hereditary foundation activated by environmental factors and immunological middle people. Numerous Epidemiologic examinations proposed that patients with psoriasis are at expanded danger of cardiovascular disease. To assess serum salusin beta level in psoriatic patients. In this relative case-control study, 50 psoriatic patients (case gathering) and 30 age-and sexual orientation coordinated solid people filling in as a benchmark group, were chosen from the outpatients' facilities of the Dermatology, Venereology, and Andrology Department, Faculty of Medicine, Benha University, during the period from October, 2018 to May, 2019. This investigation found that the serum salusin-beta levels were significantly higher in psoriasis patients contrasted with control group. This study found that the serum salusin-beta levels were significantly higher in psoriasis patients contrasted with control gathering.

Keywords: Salusin beta, Psoriatic, Diabetes mellitus.

1. Introduction
Psoriasis is a ceaseless, immune-mediated illness influencing around 100 million individuals around the world. Psoriasis influences people all things considered and can show in a wide range of structures, the most well-known being psoriasis vulgaris (or plaque psoriasis). Plaque psoriasis is portrayed by patches of erythema canvassed in a silvery-white scale, which is the aftereffect of quick hyperproliferation and dysregulated separation of epidermal keratinocytes [12].

The etiology of psoriasis is multifactorial and incorporates an unpredictable interchange of hereditary, natural, irresistible and way of life factors [3]. Genome-wide affiliation considers have distinguished various psoriasis-associated quality loci including the Human leukocyte antigen (HLA)-Cw6 quality explicitly the HLA-class 1 allele, HLA-C, situated inside Psoriasis Susceptibility Locus 1 (PSORS1 on 6p21.3) [25]. Different variations, are identified with natural insusceptible pathways, antigen introduction, and T-cell actuation and separation [15].

Salusin-beta is a strong bioactive peptide that was initially anticipated utilizing in silico investigation of a human cDNA library [20]. Salusin-beta-like immunoreactivity was later shown in human plasma and pee [6]. Salusin-beta applies consolidated fundamental and neighborhood organic activities, including hypotensive and bradycardic impacts, that are intervened by means of foundational parasympathetic incitement and negative cardiotropic inotropic. Focal salusin-beta directs hemodynamic homeostasis by actuating antidiopsgenesis, circulatory strain height and incitement of vasopressin and oxytocin discharge. Fringe exercises of salusin-beta incorporate its intense proatherosclerotic impacts and concealment of cardiovascular renovating. Notwithstanding such overpowering organic exercises, it has likewise been proposed as a biomarker for certain human maladies [19].

Metabolic Syndrome (MS) is characterized as a group of hazard factors including focal stoutness, atherogenic dyslipidemia, hypertension and glucose narrow mindedness, is a solid indicator of cardiovascular illness. Expanded mortality from cardiovascular infection in patients with serious psoriasis has been archived and psoriasis might be a free hazard factor for myocardial dead tissue, particularly in youthful patients. Psoriasis is related with MS, free of its seriousness. A few variables may add to a horrible cardiovascular hazard profile in patients with psoriasis, for example, cigarette smoking, liquor utilization, corpulence, physical idleness, homocysteinemia, mental pressure, and misery, which are all increasingly predominant in patients with psoriasis [14].

2. Aim of the work
The aim of this study is to evaluate serum salusin beta level in psoriatic patients.

3. Subjects and methods
Study Type: This examination is a near case-control study.

Ethical Consideration: This examination was done after endorsement of the Dermatology, Venereology and Andrology Department and the Ethics advisory group of Faculty of Medicine, Benha University.

Subjects and strategies: In this relative case-control study, 50 psoriatic patients (case gathering) and 30 age-and sex coordinated sound people filling in as a benchmark group, were chosen from the outpatients' facilities of the Dermatology, Venereology, and Andrology Department, Faculty of Medicine, Benha University, during the period from October, 2018 to May, 2019.

Consideration rules: Patients experiencing psoriasis vulgaris with the accompanying measures:

Patients with summed up plaque psoriasis. Psoriasis zone and seriousness file (PASI) score > 8. Age: over
18 years of age. Length of the malady > a half year. Sex: the two guys and females.

Prohibition rules: Patients who got any fundamental or skin treatment for psoriasis inside the a month or fourteen days individually preceding the examination. Pregnant or lactating female patients. Patients experiencing any fundamental (hepatic, renal sickness) or other dermatological maladies.

Subjects

Definite ace sheet is introduced in reference section A. The two patients and controls were exposed to the accompanying.

3.1 Methods

A. Clinical assessment:

All patients were exposed to the accompanying:

I. Full history taking including

Personal history. Present history.

History of present sickness including: Onset, course and term of psoriasis. Nearness of activating variables, for example, chafirvoyant pressure and occasional variety. Medication history of past medicines, type, term, reaction and the date of halting the last treatment and history of some other prescriptions. Family ancestry of psoriasis. History of Cardiovascular infection, renal sickness, diabetes mellitus, hypertension, hyperlipidemia, smoking or admission of liquor.

II. General clinical assessment

Circulatory strain was assessed by means of physical assessment and was recorded as the normal of two estimations after the members had been sitting for 5 minutes. Body weight was estimated in light attire, without shoes. Body Height was estimated. Weight list (BMI) was determined as the proportion of weight (kg) to tallness (m2) squared (kg/m2) accordingly in condition W/t/H2.

III. Dermatological assessment

Patients were analyzed to decide: site, size, circulation, number of sores. The psoriasis territory and seriousness file (PASI) score was estimated as respects; erythema, thickness, scaling and surface region and both power and degree of the psoriatic plaques were determined (Schmitt and Wozel, 2005).

The different body areas were weighted to mirror teir separate extent of body surface region (BSA).

Patients were grouped by PASI score into: Mild psoriasis: PASI < 7. Moderate psoriasis: PASI (7 – 12). Extreme psoriasis: PASI > 12.

Examinations

Research center examinations:

1- Venous examples were taken as follow

80m Peripheral blood tests were gathered from patients and controls. Followed by Centrifugation of blood tests to isolate plasma from the blood. Serum tests were put away at - 20 °C.

2- Serum levels of Salusin β was estimated by ELISA method (Bioassay Technology Laboratory Korain Biotech Co., Ltd. Feline # E 1272 Hu. Part # E 1812004).

Measure method: 0.1ml of the example cradle was included into the control well (Zero well) at that point, 0.1ml of each appropriately weakened example was added to each void well and the plate was hatched at 37°C for 90 min. 0.1ml of biotinylated hostile to human salusin B counter acting agent working arrangement into each well were included and the plate was the hatched at 37 °C for an hour. Plate was washed multiple times. 0.1ml of arranged ABC working arrangement were included into each well and the plate was hatched at 37°C for 30 min. Plate washed multiple times. 90 µl of arranged TMB shading forming was included into each well and the plate was hatched at 37°C in dull for 15-20 min. 0.1 ml of arranged TMB stop arrangement was included into each well. The O.D. absorbance was perused at 450 nm and determined by standard bend.

3.2 Statistical analysis

Inforation were gathered, reconsidered, coded and entered to the Statistical Package for Social Science (IBM SPSS) rendition 23. The quantitative information were introduced as mean, standard deviations and extents when their circulation discovered parametric. Additionally, subjective factors were introduced as number and rates. The examination between bunches with respect to subjective information were finished by utilizing Chi-square test. The examination between two free gatherings with quantitative information and parametric dispersion were finished by utilizing Independent t-test. Spearman relationship coefficients were utilized to serve the connection between's two quantitative boundaries in a similar gathering. Recipient working trademark bend (ROC) was utilized to survey the best cut off point for salusin β among patients and controls and furthermore between patients with positive and negative pressure ECG with its affectability, particularity, positive prescient worth (PPV), negative prescient worth (NPV) and region under bend (AUC).

The certainty stretch was set to 95% and the wiggle room acknowledged was set to 5%. Thus, the p-esteem was viewed as noteworthy as the accompanying: P > 0.05: Non critical. P ≤ 0.05: Significant.

4. Results

This study included 50 clinically diagnosed psoriatic patients and 30 age and sex-matched healthy controls. The patients were 32 females (64%) and 18 males (36%) with mean age 41.62±8.02 years old (range 26-56). The controls included 30 age and gender-matched healthy volunteers. They were 17 females (56.7%) and 13 males (43.3%) with mean age 42.73±7.80 years old (range 28-56) Tables (1).
Table (1) Comparison between patients’ group and control group regarding demographic data weight, height and BMI.

<table>
<thead>
<tr>
<th>Patients group</th>
<th>Control group</th>
<th>Test value</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean±SD</td>
<td>41.62 ± 8.02</td>
<td>42.73 ± 7.80</td>
<td>-0.607•</td>
</tr>
<tr>
<td>Range</td>
<td>26 – 56</td>
<td>28 – 56</td>
<td></td>
</tr>
<tr>
<td>Sex</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Females</td>
<td>32 (64.0%)</td>
<td>17 (56.7%)</td>
<td>0.425*</td>
</tr>
<tr>
<td>Males</td>
<td>18 (36.0%)</td>
<td>13 (43.3%)</td>
<td></td>
</tr>
<tr>
<td>Weight (kg)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean±SD</td>
<td>84.28 ± 13.87</td>
<td>80.90 ± 12.85</td>
<td>1.084•</td>
</tr>
<tr>
<td>Range</td>
<td>58 – 110</td>
<td>58 – 105</td>
<td></td>
</tr>
<tr>
<td>Height (m)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean±SD</td>
<td>1.68 ± 0.07</td>
<td>1.72 ± 0.09</td>
<td>-1.956*</td>
</tr>
<tr>
<td>Range</td>
<td>1.56 – 1.85</td>
<td>1.57 – 1.89</td>
<td>S</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean±SD</td>
<td>29.79 ± 4.29</td>
<td>27.47 ± 3.51</td>
<td>2.492*</td>
</tr>
<tr>
<td>Range</td>
<td>22.04 – 40.4</td>
<td>20.76 – 35.79</td>
<td></td>
</tr>
</tbody>
</table>

Independent t-test; *: Chi-square test

P ≥ 0.05: Non significant; P ≤ 0.05: Significant

There was non significant difference between both studied groups regarding age, sex and weight. Patients group had higher height and BMI than control group with significant difference between both studied groups (P<0.05) Table (1).

Table (2) Clinical history of patients group.

<table>
<thead>
<tr>
<th>Patients group</th>
<th>No. = 50</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age of onset (years)</td>
<td>Mean±SD</td>
</tr>
<tr>
<td>Range</td>
<td>13 – 45</td>
</tr>
<tr>
<td>Duration of disease (years)</td>
<td>Mean±SD</td>
</tr>
<tr>
<td>Range</td>
<td>1 – 30</td>
</tr>
<tr>
<td>Family (Hx)</td>
<td>Negative</td>
</tr>
<tr>
<td>Positive</td>
<td>5 (10.0%)</td>
</tr>
<tr>
<td>Psoriatic Arthritis</td>
<td>No</td>
</tr>
<tr>
<td>Yes</td>
<td>12 (24.0%)</td>
</tr>
<tr>
<td>PASI score</td>
<td>Mean±SD</td>
</tr>
<tr>
<td>Range</td>
<td>8.1 – 34.1</td>
</tr>
</tbody>
</table>

SD: Standard deviations

PASI: Psoriasis area and severity index

The mean age of psoriasis onset and duration was 28.96±6.33, 12.64±7.20 respectively. The majority had negative family history (90%) and only 24% had psoriatic arthritis. The mean PASI score was 14.08±5.70 Table (2).

Table (3) Comparison between patients’ group and control group regarding Salusin β.

<table>
<thead>
<tr>
<th>Patients group</th>
<th>Control group</th>
<th>Test value</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Salusin β (Pg)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean±SD</td>
<td>301.52 ± 105.10</td>
<td>236.67 ± 82.93</td>
<td>2.882•</td>
</tr>
<tr>
<td>Range</td>
<td>180 – 580</td>
<td>110 – 410</td>
<td></td>
</tr>
</tbody>
</table>

Independent t-test; *: Chi-square test

p > 0.05: Non significant; p < 0.05: Significant

This table shows that Salusin β had higher significant level in patients’ group when compared to control group (p = 0.005).
Evaluation of Serum Level of Salusin Beta in Psoriatic Patients

Table 1

<table>
<thead>
<tr>
<th>Variables</th>
<th>Cut off point</th>
<th>AUC</th>
<th>Sensitivity</th>
<th>Specificity</th>
<th>PPV</th>
<th>NPV</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Salusin β</td>
<td>&gt;420</td>
<td>0.958</td>
<td>100.00</td>
<td>91.67</td>
<td>33.3</td>
<td>100.0</td>
<td>0.0001</td>
</tr>
</tbody>
</table>

AUC= Area under the curve; PPV= positive predictive value; NPV= Negative predictive value.

The Receiver operating characteristic curve (ROC) shows that the best cut off point for Salusin β to differentiate between cases and controls was found > 420 with sensitivity of 100%, specificity of 91.67% and area under curve (AUC) of 0.958 with a highly statistically significant p-value (<0.001).

Table 4

<table>
<thead>
<tr>
<th></th>
<th>Salusin β</th>
<th>r*</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td>0.147</td>
<td>0.309</td>
<td></td>
</tr>
<tr>
<td>Age of onset (years)</td>
<td>-0.097</td>
<td>0.503</td>
<td></td>
</tr>
<tr>
<td>Duration of disease</td>
<td>0.295</td>
<td>0.037</td>
<td></td>
</tr>
<tr>
<td>Weight (kg)</td>
<td>0.002</td>
<td>0.986</td>
<td></td>
</tr>
<tr>
<td>Height (m)</td>
<td>0.038</td>
<td>0.795</td>
<td></td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>-0.113</td>
<td>0.435</td>
<td></td>
</tr>
<tr>
<td>PASI score</td>
<td>0.720</td>
<td>0.000</td>
<td></td>
</tr>
</tbody>
</table>

*r=Spearman correlation coefficients; Bold indicates statistically significant difference p<0.05

Table (4) demonstrates significant positive correlation of salusin β to duration of disease (r= 0.295, p= 0.37) and PASI score (r= 0.720, p= 0.000) Figs (2, 3).

Fig (1) Receiver operating characteristic curve (ROC) for Salusin β between patients and controls.

![Receiver operating characteristic curve (ROC)](image1)

![Receiver operating characteristic curve (ROC)](image2)

Fig (2) Showing significant positive correlation between Salusin β and duration of disease.

Fig (3) Showing significant positive correlation between Salusin β and PASI score.
In this investigation our point was to assess serum salusin beta level in psoriatic patients.

This examination was led on 50 psoriatic patients (bunch An) and 30 age and sex coordinated solid control subjects (bunch B).

In the interim Salusin-β levels were seen as essentially higher in psoriatic patients contrasted with control gathering. As far as we could possibly know, Salusin-β was estimated in psoriatic patients in a single report as it were. Xu et al. [24]; Cakir et al. [1]; Zhou et al. [25] additionally detailed that Salusins are related with aggravation and oxidative pressure.

In the interim Salusin-β levels were seen as essentially higher in other immune system sicknesses which are portrayed by aggravation like psoriasis, As Ozgen et al. [16]; Erden [4] examined whether Salusins were related with aggravation and fiery illnesses and found that serum Salusin-α levels and Salusin-β levels were higher in patients with Behcet sickness.

Moreover, Cakir [2] found that serum Salusin-β levels were essentially higher in Relapsing-Remitting Multiple Sclerosis (RRMS) patients than in control gathering, realizing that RRMS is an immune system illness of focal sensory system and its pathology is described by irritation.

In this examination there was a significant positive connection between's the serum levels of Salusin-β with the span and seriousness of psoriasis. This examination additionally found that patients with psoriatic joint inflammation had noteworthy higher mean degree of Salusin-β than those without psoriatic arthritis.

Furthermore, table 5 shows that although male patients and those with negative family history of psoriasis had higher level of Salusin-β than their counter patients the difference was insignificant (p > 0.05).

5. Discussion

Psoriasis is described by hyperproliferation and unusual separation of epidermal keratinocytes, lymphocyte invasion for the most part of T lymphocytes and different endothelial vascular changes in the dermal layer, for example, angiogenesis, dilatation and high endothelial venule (HEV) arrangement Guenther and Ortonne [7].

A few investigations exhibited that patients with psoriasis have higher dangers of myocardial localized necrosis, angina, atherosclerosis, fringe vascular ailments, and stroke Wakkee [14]. The immediate advancement of cardiovascular maladies, e.g., by endothelial brokenness brought about by for all time raised degrees of go betweens, for example, VEGF, significantly raises the recurrence of atherosclerosis even among patients with psoriasis who have none of the old style chance components [10].

Mehlis estimated a potential clarification for the relationship among psoriasis and atherosclerosis which is the nearness of constant aggravation that happens in view of persevering discharge of TNF-alpha and other proinflammatory cytokines, for example, interleukin-1 and interleukin-6, which hasten psoriasis. This proinflammatory cytokines invigorate interminable foundational irritation and actuate endothelial brokenness, adjusted glucose digestion and insulin opposition that assume a noteworthy job in the advancement of cardiovascular maladies, e.g., by development of cardiovascular ailments Guenther and Ortonne [7].

There are two types of Salusin: Salusin-α and Salusin-β. Salusin-α is probably going to forestall atherosclerosis Watanabe et al., [23], while Salusin-β may go about as a potential proatherogenic factor [27].

Shichiri et al. [20] found Salusin-α in human incipient organism. PreproSalusin is communicated in various pieces of the focal sensory system Shichiri et al., [20], Suzuki et al., [21] . Salusins are generally found in various districts of the cerebrum, in the safe framework and in numerous tissues in the body, Salusin-α and Salusin-β likewise were resolved in human serum and pee (Sato et al., 2006).

Studies on Salusins were basically centered around their relationship with the cardiovascular framework. It has been demonstrated that Salusin-β diminishes pulse and circulatory strain when administrated intravenously in rodents [8].

Table (5) illustrates that patients with psoriatic arthritis had significant higher mean level of Salusin-β than those without psoriatic arthritis.

<table>
<thead>
<tr>
<th></th>
<th>Students t-test</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sex</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Females</td>
<td>280.94 ± 93.06</td>
<td>180 – 350</td>
</tr>
<tr>
<td>Males</td>
<td>338.11 ± 117.59</td>
<td>188 – 380</td>
</tr>
<tr>
<td>Family (Hx)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Negative</td>
<td>306.40 ± 100.62</td>
<td>180 – 350</td>
</tr>
<tr>
<td>Positive</td>
<td>257.60 ± 145.80</td>
<td>180 – 350</td>
</tr>
<tr>
<td>Psoriatic Arthritis</td>
<td></td>
<td></td>
</tr>
<tr>
<td>No</td>
<td>284.18 ± 94.40</td>
<td>180 – 350</td>
</tr>
<tr>
<td>Yes</td>
<td>356.42 ± 122.08</td>
<td>180 – 350</td>
</tr>
</tbody>
</table>

Table (5) Relation between Salusin-β and the other studied parameters in patients’ group.

Independent t-test: p > 0.05: Non significant; p ≤ 0.05: Significant.
that Salusin-β builds irritation through initiation of NF-
κB.

Additionally, Sato [17] in an in vitro examination
expressed that monocyte/macrophage cells discharge
Salusin-β in the wake of being animated by tumor
putrefaction factor alpha (TNF-α) and lipopolysaccharide.

Unexpectedly, Cakir [2] found that there was no
huge connection between’s the plasma Salusin-β levels
and the illness span of Multiple sclerosis which is
likewise an immune system issue described by
aggravation.

Erden et al. [4] additionally detailed that psoriasis
was a hazard factor for subclinical atherosclerosis.
Erden et al. [5] likewise found that there was a critical
increment in hazard factor for atherosclerosis in
psoriasis with low degree of Salusin-α and significant
levels of Salusin-β and that his outcomes were
predictable with writing.

Niepolski et al [13] revealed that by breaking
down numerous investigations on Salusins and their
relationship with atherosclerosis and lipid unsettling
influences, They accepted that the metabolic reliance
may exist between them. Also detailed that the
statement of Salusin-β is related with more regrettable
that the specific relationship and instruments by which
Salusins influence lipid digestion and atherosclerosis
stay hazy and that further examinations are expected to
fathom these issues.

Taking everything into account high serum levels
of Salusin-β in psoriatic patients could be an early
aware of research the nearness of subclinical
atherosclerosis in coronary veins, and in such cases it is
recommended that activity echocardiography or
radionuclide myocardial perfusion imaging could be
better and increasingly noteworthy in finding of any
conceivable coronary supply route illness or
atherosclerosis.

6. Conclusion
This investigation found that the serum salusin-β
levels were significantly higher in psoriasis patients
contrasted with control gathering. In this examination
we can infer that high serum level of salusin-β in
psoriatic patients could be an early aware of research
the nearness of subclinical atherosclerosis in coronary
veins.

References
effects of Salusin-alpha and Salusin-beta on renal
ischemia/reperfusion damage and their levels in
ischemic acute renal failure. Biotech. Histochem.,
level of plasma Salusin-α and Salusin-β in patients
with multiple sclerosis. Mult Scler Relat Disord,
015354,2014.
and Salusin beta levels in patients with Behcet’s
582, 2014.
and Salusin-beta levels in patients with
53, 2015.
levels of human salusin-β, a potent hemodynamic
and atherogenesis regulator. PLoS One, Vol.8, PP.
e76714, 2013.
psoriasis: science behind therapy. J Cutan Med
as cardiac depressors in rat. Hypertension,
accelerates inflammatory responses in vascular
endothelial cells via NF-kappa B signaling in LDL
receptor-deficient mice in vivo and HUVECs in
possible risk factor for development of coronary
artery calcification. Br J Dermatol, Vol.156,
psoriasis and biologic immunotherapy. J Am Acad
care for the management of psoriasis and psoriatic
arthritis: section 6. Guidelines of care for the
management of psoriasis and psoriatic arthritis:
case-based presentations and evidence-based
conclusions. J Am Acad Dermatol, Vol.65,
adropin: New peptides potentially involved in lipid
metabolism and atherosclerosis. Adv Med Sci,
association between psoriasis and comorbidities.
Genetic Variants in the IL-23 and NF-kappaB
Pathways Discriminates between Mild and Severe
Salusin-beta from human monocytes/macrophages.


