***Original Research Article***

**Squamous Cell Carcinoma of the Skin in Ocular-Cutaneous, Cutaneous, and Ocular Albinism: A Case Study in Southeastern Nigeria**

**Abstract**

I**ntroduction**: Squamous Cell Carcinoma (SCC) is a prevalent skin cancer, with a higher incidence among individuals with albinism due to their lack of melanin protection [1]. **Aim:** This study investigates the association between SCC and different types of albinism ocular-cutaneous, cutaneous, and ocular by evaluating serum markers such as Squamous Cell Carcinoma Antigen (SCCag), Human Melanoma-Associated Antigen (HMAA), and Alpha-Tumor Necrosis Factor (Alpha-TNF) and was conducted in Southeastern Nigeria, where albinism prevalence is relatively high [2]. **Method: A total 300 individuals with albinism consisting of 100** oculo-cutaneous (OCA), 100 cutaneous (CA), and 100 ocular albinism (OA) participated in this study. They were aged-matched with 100 controls. Eight ml of venous blood sample was collected from each participant. Serum SCCag, HMAA and alpha-TNF levels were determined using ELISA technique. Data generated were subjected to statistical analysis using IBM SPSS version 23. **Results:** The results revealed significantly higher serum levels of SCCag, HMAA, and Alpha-TNF in individuals with ocular-cutaneous, cutaneous and ocular albinism compared to controls (p=0.000 in each case). Suggesting a heightened risk of SCC development. **Conclusion:** These findings underscore the need for targeted interventions, including regular skin screenings and preventive measures [3].

**Keywords:** Squamous Cell Carcinoma, Albinism, SCCag, HMAA, Alpha-TNF, Nigeria, Skin Cancer.

**Introduction**

Squamous Cell Carcinoma (SCC) is a common non-melanoma skin cancer arising from keratinocytes, often linked to excessive ultraviolet (UV) exposure [4]. Individuals with albinism, particularly those with ocular-cutaneous albinism, face a higher risk of developing SCC due to their lack of melanin, which provides natural UV protection [5]. In Nigeria, where sun exposure is high, SCC cases among albinos are frequently reported [1]. However, limited studies have assessed the biochemical markers associated with SCC risk in albinos [5]. This study evaluates the association of SCC with serum markers in different types of albinism, providing insight into their vulnerability and potential preventive strategies.

Squamous Cell Carcinoma (SCC) is a prevalent form of non-melanoma skin cancer that arises from keratinocytes and is frequently associated with excessive ultraviolet (UV) radiation exposure. Chronic sun exposure is a well-established risk factor for SCC, particularly in individuals with reduced skin pigmentation, as melanin plays a crucial role in protecting the skin from UV-induced DNA damage [1,2]. Albinism, a genetic condition characterized by the partial or complete absence of melanin, increases the risk of developing SCC, particularly among individuals with oculo-cutaneous albinism (OCA), cutaneous albinism (CA), and ocular albinism (OA). OCA affects the skin, hair, and eyes, rendering individuals highly susceptible to UV damage [3]. CA primarily affects the skin and hair, while OA primarily impacts the eyes, with minimal or no skin involvement. Among these subtypes, individuals with OCA and CA face a significantly higher risk of SCC due to their extensive sun exposure and lack of protective melanin [4,5]. In tropical regions such as Nigeria, where sun exposure is intense, cases of SCC among albinos are frequently reported [6]. Despite this, research focusing on the biochemical markers associated with SCC in albinos remains limited. Understanding the molecular and biochemical changes that predispose individuals with albinism to SCC is essential for developing early diagnostic tools and effective preventive strategies [7].

**Justification for the Study**

Studies have shown that individuals with albinism, particularly in sun-intense regions, experience disproportionately high rates of SCC [8]. Due to the lack of melanin, these individuals suffer from frequent sunburns, actinic keratosis, and subsequent malignant transformations leading to SCC. The limited availability of effective sunscreen and protective measures exacerbates their vulnerability [9].  
While the link between albinism and SCC is well recognized, few studies have investigated the biochemical markers that could serve as indicators of SCC development in different types of albinos [10]. Most existing research focuses on clinical observations rather than molecular or biochemical assessments [11]. The importance of serum biomarkers in early detection is identifying serum biomarkers associated with SCC in albinos could provide valuable insights into the disease's pathogenesis and aid in early diagnosis [12]. Biomarkers such as inflammatory cytokines, oxidative stress indicators, and genetic mutations linked to SCC could be explored to develop non-invasive diagnostic tools [13].

Understanding the molecular pathways involved in SCC progression among albinos could inform the development of targeted preventive and therapeutic strategies. These could include enhanced sun protection protocols, antioxidant-based interventions, and personalized treatments tailored to the specific needs of albino subpopulations[14].This research has significant public health relevance, particularly in regions with a high albino population and intense UV exposure [15]. The findings could contribute to policy recommendations for improved healthcare access, education on sun protection, and early cancer screening programs for albinos [16].

**Inclusion criteria**: Subjects were male and female between the ages 18-60 years were characterized by those with skin, hair and eyes hypo pigmented, characteristic skin lesions distribution, reduced visual acuity, nystagmus and strabismus. The cutaneous were primarily characterized by hypo pigmented skin without any direct involvement of the eyes while Ocular were characterized only by those with ocular involvement with characteristics like nystagmus, strabismus and poor visual acuity [ 5 ]

**Exclusion criteria**: Pigmented healthy subjects without any of the characteristics listed for albinism [5].

**Materials and Methods**

**Study Population and Design**

Sample size was calculated based on 95% confident interval, desired accuracy of 0.05. This study was conducted in some South-eastern Nigerian Universities, namely; Anambra State University teaching hospital Awka, Imo State University teaching hospital Orlu and University of Nigeria teaching hospital Enugu. Focusing on individuals diagnosed with ocular-cutaneous, cutaneous, and ocular albinism. An approval was obtained from the albino foundation of Nigeria and informed consent was obtained from all participants.

Participants were clinically examined and categorized based on standard diagnostic criteria for albinism. The study population included 100 OCA, 100 CA 100 OA and 100 control group of non-albino individuals for comparison.

**Sample Collection and Storage**

8 millimeter of venous Blood samples were collected from participants under aseptic conditions. They were stored in plain tubes, centrifuged and serum separated into aliquots, stored at -20oC for makers for determination of squamous carcinoma antigen, Human melanoma antigen and alpha-TNF.

**Laboratory Analysis Methods**

Enzyme-linked Immunosorbent Assay (ELISA) method were used for the estimation of serum SCCag, Alpha-TNF and Human melanoma Associated antigen (HMAA) concentration, using reagent kits all supplied by Melsin Medical Company Limited.Catalog numbers EKHU-0259 for SCCag, EKHU-2228 for HMAA, and EKHU-2141 for alpha-TNF.

Principle of ELISA technique; The Enzyme-linked Immuosorbent Assay (ELISA) is an analytical technique designed to detect and quantify soluble substances such as peptides, proteins, antibodies and hormones. The fundamental principle involves on the specific interaction between an antigen and its corresponding antibody, utilizing an enzyme linked detection system to produce a measurable signal indicative of the presence and concentration of target analyte [6].

**Statistical Analysis**

Data were analyzed using SPSS version 25.0. The mean ± standard deviation (SD) of SCCag, HMAA, and Alpha-TNF   levels were compared between albino individuals groups and controls using independent t-tests. A p-value < 0.05 was considered statistically significant

**Results**

**TABLE 1**

**Mean±SD values of serum markers of squamous cell carcinoma of skin; SCCag, HMAA, and alpha- TNF in Ocular-Cutaneous albino and Control**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| VARIABLES  MEAN± SD | OCULAR-CUTANEOUS, N=100 | CONTROL  N= 100 | t-Value | p-Value  0.05 |
| SCCag mg/dl  Lower 95% C.I  Upper 95% C.I | **288.72±98.97**  **265.07**  **312.00** | **217.51±30.45**  **211.50**  **223.52** | **4.421** | **0.000** |
| HMAA(mg/dl)  Lower 95% C.I  Upper 95% C.I | **2.31±1.90**  **1.94**  **2.70** | **1.07±0.57**  **0.90**  **1.23** | **4.433** | **0.000** |
| TNF(mg/dl)  Lower 95% C.I  Upper 95% C.I | **99.73±83.10**  **85.78**  **125.71** | **74.59±7.68**  **72.40**  **76.77** | **2.137** | **0.038** |

**OCA: Ocular-Cutaneous Albino**

**SCC: Squamous Cell Carcinoma**

**HMAA: Human Melanoma Associated Antigen**

**TNF: Cellular Damage Marker**

**TABLE 2**

**Mean±SD values of Serum markers of Squamous Cell Carcinoma of skin; SCCag, HMAA, and alpha- TNF in Cutaneous albino and Control**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| VARIABLES  MEAN± SD | CUTANEOUS N=100 | CONTROL  N= 100 | t-Value | p-Value  0.05 |
| SCCAg mg/dl  Lower 95% C.I  Upper 95% C.I | **298.62±120.65**  **274.91**  **322.53** | **217.51±30.45**  **211.50**  **223.52** | **4.513** | **0.000** |
| HMAA(mg/dl)  Lower 95% C.I  Upper 95% C.I | **1.81±0.14**  **1.79**  **1.85** | **1.07±0.57**  **0.95**  **1.18** | **8.497** | **0.000** |
| TNF(mg/dl)  Lower 95% C.I  Upper 95% C.I | **120.9475±111.46**  **99.30**  **143.29** | **74.59±7.68**  **73.07**  **76.11** | **2.928** | **0.005** |

**OCA: Ocular-Cutaneous Albino**

**SCC: Squamous Cell Carcinoma**

**HMAA: Human Melanoma Associated Antigen**

**TNF: Cellular Damage Marker**

**TABLE 3**

**Mean ±SD values of Serum markers of Squamous Cell Carcinoma of skin; SCCag, HMAA, and alpha- TNF in Ocular Albino and Control**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| VARIABLES  MEAN± SD | OCULAR  N=100 | CONTROL  N= 100 | t-Value | p-Value  0.05 |
| SCCag mg/dl  Lower 95% C.I  Upper 95% C.I | **260.49±11.72**  **258.60**  **263.24** | **217.51±30.45**  **211.50**  **223.52** | **9.277** | **0.000** |
| HMAA(mg/dl)  Lower 95% C.I  Upper 95% C.I | **1.98±0.08**  **1.96**  **1.99** | **1.07±0.57**  **0.95**  **1.18** | **10.823** | **0.000** |
| TNF(mg/dl)  Lower 95% C.I  Upper 95% C.I | **86.05±1.59**  **85.71**  **86.33** | **74.59±7.68**  **73.07**  **76.11** | **10.383** | **0.000** |

OCA: Ocular-Cutaneous Albino

SCC: Squamous Cell Carcinoma

HMAA: Human Melanoma Associated Antigen

TNF: Cellular Damage Marker

**Presents Mean±SD values of serum markers of squamous cell carcinoma of skin; SCCag, HMAA, and alpha- TNF in Ocular-Cutaneous albino and Control groups.**

There were statistically significant higher serum levels of SCCag (288.7296 ±89.9769 vs 217.5150 ± 30.4535, P=0.000), HMAA (2.3168 ± 1.9005 vs 1.0726± 0.5756, P=0.000) and alpha-TNF (99.7359 ± 83.1010 vs (74.5924 ± 7.6895, p=0.000) in Cutaneous albino compared to the control (Table 1).

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**Presents Mean±SD values of serum markers of squamous cell carcinoma of skin; SCCag, HMAA, and alpha- TNF in Cutaneous albino and Control groups.**

There were significantly higher serum levels of SCCag (298.6292±120.6539 vs. 217.5150±30.4535, p=0-000), HMAA (1.8154±0.14897 vs. 1.0726±0.5756, p=0.000) and Alpha-TNF (120.9475±111.4628 vs. 74.5924±7.6895, P=0.005) in Cutaneous Albino compared to the control (Table 2).

**Presents Mean±SD values of serum markers of squamous cell carcinoma of skin; SCCag, HMAA, and alpha- TNF in Ocular albino and Control groups.**

There were significantly higher serum levels of SCCag (260.4962±11.772553 vs. 217.5150±30.4535, p=0-000), HMMA (1.9825±0.08336 vs. 1.0726±0.5756, p=0.000) and Alpha-TNF (86.0535±1.5975 vs. 74.5924±7.6895) in Ocular Albino compared to the control (Table 3).

**Discussion**

In table 1, 2 and 3 shows significant higher serum levels of SCCag, in ocular-cutaneous (282.75±98.97 mg/dl), cutaneous (298.62±120.65 mg/dl) and ocular albino (260.49±11.72 mg/dl) compared to controls (217.51±30.45 mg/dl), with p-values of 0.000. This suggests a higher risk or presence of SCC in albino individuals, likely due to increased UV radiation susceptibility in individuals with reduced melanin protection levels [6].These findings reaffirm that albinos are at a higher risk of SCC due to their inherent lack of melanin, leading to increased UV-induced DNA damage [6]. The cutaneous albino group had the highest SCCag levels, which may be attributed to greater cumulative UV exposure compared to ocular and ocular-cutaneous albinos [7].

HMAA levels were also significantly increased in ocular-cutaneous albino (2.31±1.90 mg/dl) and cutaneous albino (1.81±0.14 mg/dl) and ocular (1.98±0.08 mg/dl) compared to controls (1.07±0.57 mg/dl), with p-values of 0.000. HMAA is associated with melanoma and other skin-related malignancies, and its elevation in albino populations suggests increased cellular stress or premalignant transformation [1]. This suggests a possible predisposition to melanoma-like changes or increased skin cell turnover in albino populations [1].

The higher HMAA levels in ocular-cutaneous albinos could indicate a more widespread systemic response to UV damage compared to purely cutaneous or ocular albinos [3].

Alpha-TNF, a pro-inflammatory cytokine involved in tumor progression, was significantly higher in cutaneous albino (120.947±111.46 mg/dl, p=0.005), Ocular-cutaneous albino (99.73±83.10 mg/dl, p=0.038) and Ocular albino 86.05±1.59 mg/dl, p=0.000) compared to controls (74.59±7.68 mg/dl). This suggests a heightened inflammatory response in albinos, likely due to chronic UV exposure and oxidative stress leading to SCC development [8]. The higher Alpha-TNF in cutaneous albino compared to ocular-cutaneous albino supports the idea that inflammation-driven carcinogenesis is more pronounced in individuals with extensive skin exposure [1].

The findings of this study support existing evidence that albino individuals, particularly those with ocular-cutaneous and cutaneous forms, have a significantly higher risk of developing SCC [9]. The elevated SCCag and Alpha-TNF levels indicate increased inflammation and tumor activity, aligning with SCC pathogenesis [10].

Comparisonwithprevious studies have demonstrated that SCC is more prevalent among albinos in sun-intensive regions [11]. The high SCCag levels found in this study reinforce the role of SCCag as a reliable biomarker for early SCC detection in albinos [5]. The increase in Alpha-TNF further suggests a strong inflammatory response, which may contribute to tumor progression [12].

**Conclusion**

The significant differences in serum markers among albinos highlight the urgent need for early screening programs in Nigeria. Sunscreen use, protective clothing, and community awareness campaigns should be prioritized to mitigate SCC risk. This study demonstrates that albino individuals, especially those with ocular-cutaneous and cutaneous albinism, have significantly higher SCCag, HMAA, and Alpha-TNF levels, indicating a higher risk of SCC. The findings emphasize the need for proactive skin cancer surveillance and protective measures among albinos in Southeastern Nigeria.

**Recommendations**

Early Screening Programs and regular dermatological screenings for all albinos’ individuals is necessary. These findings reinforce that albino populations, particularly cutaneous albinos, are at greater risk of SCC due to increased biomarker expression linked to skin malignancies. Regular dermatological screening and protective measures such as sunscreen application and reduced UV exposure are strongly recommended.

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