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# بسم الله الرحمن الرحيم

مركز الشبكات وتكنولوجيا المعلومات

قسم التوثيق الإلكتروني



Safaa Mahmoud



# جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم

## قسم

نقسم بالله العظيم أن المادة التي تم توثيقها وتسجيلها  
على هذه الأقراص المدمجة قد أعدت دون أية تغييرات





**Comparative Study on the Effect of Mesenchymal Stem  
Cells Derived Exosomes Versus Platelets Rich Plasma on  
Healing of Skin Burn in Albino Rat.  
A Histological Study**

*Thesis*

*Submitted for Partial Fulfillment of Master's Degree  
in Histology and Cell Biology*

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*2022*

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ

قَالَ

سُبْحَانَكَ لَا مَعْلَمَ لَنَا  
إِلَّا مَا عَلَّمْتَنَا إِنَّكَ أَنْتَ  
الْعَلِيمُ الْعَظِيمُ

صدق الله العظيم

سورة البقرة الآية: ٣٢





## Acknowledgement

*First and foremost, thanks to **Allah** the kindest and the most merciful to whom I relate any success in achieving any work in my life.*

*I would like to express my sincere thanks and gratitude to **Prof. Dr. Amany Mohamed El Shawarby**, Professor of Histology and cell biology, Faculty of Medicine, Ain Shams University, for her kind supervision, valuable advice, and consistent encouragement and support. It was such a great honor to work under her guidance.*

*I am particularly very grateful and appreciative to **Ass: Prof. Dr. Sara Abdel Gawad El Sebay**, Assistant Professor of Histology and cell biology, Faculty of Medicine, Ain Shams University, for her great help, sincere efforts, and continuous encouragement and support throughout the journey, which made the completion of this work much easier. It was a pleasure working under her supervision.*

*I would further like to thank **Dr. Mohamed Ahmed Abdou Hegazi**, Lecturer of Histology and Cell Biology, Faculty of Medicine, Ain Shams University for her great help and precious instructions throughout this work.*

*Finally, I would like to express my sincere gratitude, love, respect, and appreciation to all my professors and my colleagues for their continuous support till this work was completed.*

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*Mai Mostafa Mohamed Naser*

# Dedication

*Words cannot describe my love, thanks, gratefulness, and respect to my parents, my sister my brothers, and my husband for their warm kindness and genuine support. Without their care, patience, encouragement, and support, I would have never achieved any success. To all of them, I dedicate my work.*

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## *List of Abbreviations*

Abb.	Full-term
ADSCs.....	Adipose tissue derived stem cells.
AMPs.....	Antimicrobial peptides.
ANOVA .....	One-way analysis of variance.
DAB .....	Diaminobenzidine tetrahydrochloride.
DEPA .....	Dose, efficiency, purity, activation process of the PRP.
DMEM .....	Dulbecco's modified Eagles medium.
DNA .....	Deoxyribonucleic acid.
ECM .....	Extracellular matrix.
EGF .....	Epidermal growth factor.
EMT .....	Epithelium mesenchymal transition.
EPC .....	Endothelial progenitor cells.
FBS.....	Fetal bovine serum.
FGF-2 .....	Fibroblast growth factor-2.
FGF-9 .....	Fibroblast growth factor-9.
GFs .....	Growth factors.
H&E .....	Haematoxylin and eosin.
H2O2.....	Hydrogen Peroxide.
HPF .....	High power field.
HRP.....	Horse-radish peroxidase.



## *List of Abbreviations (Cont.)*

Abb.	Full-term
IFN- $\gamma$ .....	Interferon- $\gamma$ .
IGF-1 .....	Insulin-like growth factor-1.
IL-1 $\beta$ .....	Interleukin-1 $\beta$ .
IL-6.....	Interleukin-6.
IP .....	Intraperitoneal.
LP-PRF.....	Leukocyte-poor platelet-rich fibrin.
LP-PRP.....	Leukocyte-poor PRP.
LR-PRF .....	Leukocytes-rich platelet rich fibrin.
LR-PRP .....	Leukocyte-rich PRP.
LSD .....	least significance difference.
MASRI.....	Medical Ain Shams Research Center.
mRNA .....	Messenger Ribonucleic acid.
MSCs.....	Mesenchymal stem cells.
MVEs .....	Multivesicular endosomes.
NF-kB.....	Nuclear factor kappa B.
PAP .....	Platelet average plasma.
PBS.....	Phosphate buffer saline.
PCNA .....	Proliferating cell nuclear antigen.
PDGF.....	platelets derived growth factor.
PPP .....	Platelet poor plasma.

## *List of Abbreviations (Cont.)*

Abb.	Full-term
PRF.....	Platelet-rich fibrin.
PRP.....	Platelet-rich plasma.
RCMB .....	Regional Center for Mycology and Biotechnology.
ROS .....	Reactive oxygen species.
rpm .....	Round per minute.
SD.....	Standard deviation.
SKP .....	Skin derived precursor.
SPSS.....	Statistical Package for the Social Sciences.
TBI .....	Traumatic brain injury.
TEM .....	Transmission electron microscopy.
TGF $\alpha$ .....	Transforming growth factor- $\alpha$ .
TGF- $\beta$ .....	Transforming growth factor-beta.
TNF- $\alpha$ .....	Tumor necrosis factor- $\alpha$ .
UV .....	Ultraviolet.
VEGF .....	Vascular endothelial growth factor.
$\gamma$ - $\delta$ .....	Gamma delta.

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# ABSTRACT

## **Background:**

Burn injury is considered one of the major causes of trauma to the human body that may lead to death and disability. Secondary infections, long healing periods, and healing with scars still appear as the main difficulties in burn wound management. Therefore, new strategies are needed to promote and help in wound healing and repair.

Recent studies have demonstrated that exosomes help tissue repair because of high stability, non-immune rejection and easy control of dosage and concentration.

Platelet-rich plasma (PRP) is an autologous concentration of platelets in concentrated plasma. It has been used to promote soft and hard tissue healing.

**Aim of the work:** to compare the effect of Mesenchymal stem cells (MSCs) derived exosomes versus platelets rich plasma (PRP) on the promotion of healing of experimentally induced second degree partial thickness burn injury in adult male albino rats.

## **Materials and methods:**

The thirty-two adult male albino rats were divided randomly into 4 groups. **Group I** (control). **Group II** (burn injury group) that will be left for spontaneous healing. **Group III** (burn injury treated with PRP intradermal injection). **Group IV** (burn injury treated with intradermal MSCs derived exosome). All the animals were sacrificed at 7th day (subgroup a) and 21th day (subgroup b). The skin biopsies were obtained and processed for histological and immunohistochemical studies. Morphometrical & statistical analysis was also done.

## **Results:**

H&E stained section of untreated burn (**subgroup IIa**) showed coagulative necrosis of the epidermis and papillary layer of the dermis. Sign of epithelial migration from the edge of the wound appeared. Significant decrease in both area percentage of collagen fibers and CD34 positive cells was detected in the dermis compared to all other subgroups. In **Subgroup IIb** the surface of the wound was covered by an eschar that converted partial thickness burn into complete thickness burn. Absence of epidermis occurred with significant decrease in area percentage of collagen fibers in the dermis. The dermis also showed apparent increase in the inflammatory cells together with significant increase in CD34 positive cells. In **PRP treated group**, the epidermis revealed incomplete differentiation with significant decrease in the epidermal thickness

compared to exosomes treated group. After **1 week** PCNA positive cells showed significant decrease While, area percentage of collagen fibers and CD34 positive cells showed significant increase compared to exosomes treated subgroup. After **3 weeks, PRP treated subgroup** showed significant increase in PCNA positive cells compared to exosomes treated subgroup. However, the area percentage of collagen fibers and CD34 positive cells were similar to control group. **One-week exosomes treated subgroup** revealed well differentiated epidermis with significant increase in the epidermal thickness and PCNA positive cells compared to PRP treated subgroup. However, there was significant decrease in the area percentage of collagen fibers and CD34 positive cells compared to PRP treated subgroup. **Three weeks exosomes treated subgroup** showed statistical results similar to control group.

**Conclusion:**

The current study revealed that burn treated exosomes group was better than PRP treated group. In addition, PRP treated group revealed hyperplastic changes in epidermis with abnormal remodeling phase of burn healing in 20% of specimens.

**Keywords:** partial thickness skin burn, MSCs derived exosomes, PRP, male albino rat.