

Effect of a single dose of a bradykinin potentiating factor isolated from scorpion venom (*Buthus occitanus*) on total protein and albumin in serum of irradiated growing male Guinea pigs.

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ABSTRACT

Whole body gamma irradiation of growing male Guinea pigs at the dose of 150 rads/animal reduced the levels of total protein and albumin. Growing Guinea pigs males (250 ± 30 gm. b. w.) were divided into two main groups; non-irradiated and irradiated groups and every main group were divided into four sub-groups. Irradiated growing Guinea pigs males groups were exposed to 150 Rads/animal by cesium-137 irradiation unit with a dose route of 1.06483 rad per second in a uniform manner and divided into four sub-groups. In irradiated groups; the first sub-groups were interaperitoneally (i.p) injected with 100 μ L saline solution. The second sub-groups were i.p. injection of bone marrow cells from their twins (2.5×10^6 cells/animal). The third sub-groups injected i. p with BPF ($1 \mu\text{gm/gm. b. w.}$) which was dissolved in saline solution.

The fourth sub-group injected with i.p solution of BPF ($1 \mu\text{g/gm. b. w.}$) together with bone marrow cells from their twins (2.5×10^6 cells/animal). The non-irradiated sub-groups were injected with the same manner of irradiated sub-groups. The level of total protein and albumin in serum were measured at periods of 1, 3, 7, 14 and 21 days. Irradiated animals i.p. injection of BPF (single dose); appears slight to improve the changes which occurred in both total protein and albumin compared to irradiated sub-group. However, irradiated Guinea pigs injected with single dose of BPF together with bone marrow cells were more improvement of these parameters than sub-groups injected with only single dose of BPF. The improvement of these parameters may be attributed to stimulate growth factors and/or growth hormones to increase protein synthesis as an effect on the liver that necessitates future investigations.

Key words: Irradiation, Bradykinin potentiting factor, total protein, Albumin and Guinea pigs.

INTRODUCTION

Ionizing radiation interacts with biological system through free radicals generated by water radiolysis. It is generally agreed that this indirect action plays an important role in the induction of oxidative stress leading to cellular damage and organ dysfunction (Berroud *et al.*, 1996 and Othman, 1998). It worthy to mention that apoptosis, as a physiological contrast of mitosis, arises during early post-irradiation period as a response to blockade of mitotic activity and causes it's abortive effect on bone marrow cellularity (Vlasov and Kvacheva, 1998; El-Missiry *et al.*, 1998 and 2007 and El-Missiry, 1999).

Serum proteins are synthesized and secreted by several cell types depending on the nature of the individual serum protein. The major functions of these proteins is maintenance of osmotic pressure intravascularly and hence maintenance of blood pressure and fluids in the circulation. They also carry out transport and storage function for several minerals, growth factors and hormones. Another strategic

function of serum proteins is a defensive function mediated by gamma globulins produced by mature B-lymphocyte (Guyton and Hall, 2000). It worthy to mentioning that, whole body gamma irradiation caused changes in the serum globulins in irradiated Guinea pigs (Salman 2002). Moreover, bradykinin stimulates the mitotic activity of bone marrow and thymus cells of irradiated mice (Sukhanova *et al.*, 1977 and Sukhanova, 1986). Furthermore, Nassar and Soliman (1989) isolated BPF1, BPF2 and BPF3 from the Egyptian scorpions, *Buthus occitanus* and *Leiurus quinquestriatus* as well as the Cobra; *Naja haje haje* respectively. *In vitro* study of physiological activities of these fractions was found to be similar to those of BPFp and BPFc isolated from venom of *Bothrops jararaca* BPF of *Buthus occitanus* venom showed growth factor-like activity. This activity was decreased by addition of bradykinin inhibitors on cellular cultures of Baby Hamster kidney (BHK21) cell line. Also in female mice, this fraction enhanced development of the ovarian follicles and the cellular growth of the uterus as well as the level of female sex hormone estradiol (Nassar *et al.*, 1989, 1990, 1991 and 1992). It worth to mentioning that, bradykinin potentiating factors have been isolated from snakes, vipers, scorpions and jelly fish venoms (Abu-Amra, 1994, 2000, 2001a and b, Abd-El-Rahim, 1990 and 1995, Abu-Amra and Abd-El-Rahim, 1992 a and b; Abd-El-Rahim and Abu-Amra, 1992; Salman, 1995; Abd-El-Rahim *et al.*, 1996 and Salman, 2002 and 2008). Additionally, Salman (1995) showed that scorpion venom fraction (BPF) accelerates burn wound repair. The mechanism of action was attributable to direct growth factor-like effects and/or indirect stimulation of endogenous bradykinin, which in turn stimulated endogenous prostaglandin biosynthesis. In particular, the elevated level of prostaglandin E2 fraction, a cytoprotective regulatory substance, might be the responsible effectors for such an accelerated burn wound healing. Remarkably, the extract induced complete re-epithelialization of the surface with abundant connective tissue in the dermis (Salman, 1995). Moreover, (Salman, 2002) showed that injection of BPF in sub-lethally-irradiated and non-irradiated Guinea pig accelerated the generation of thymus and spleen cellularity and completely recovered; the normal platelets, WBCs, RBCs and blood picture without noticeable toxic effects in non-irradiated control animals. Based on the aforementioned proliferation-inducing and cytoprotectant abilities of BPF of *Buthus occitanus* venom, this work has been designed to assess the ameliorating effect of interaperitoneally (i. p.) injected BPF only and i.p. injection of bone marrow (BM) cells from their twins (2.5×10^6 cells/animal) together with BPF on irradiated male Guinea pigs.

Materials and Methods:

Two hundreds male growing guinea pigs of approximate age and weight (4 month and about 250 ± 30 gm b. w.) were selected from the Animal House Facility of the Egyptian Organization for Biological Products and Vaccines (VACSERA), Helwan, Cairo, Egypt. The animals were housed in the Animal House of the Faculty of Science, South Valley University, Qena, Egypt, for two weeks under the natural day and night periods and supplied with a balanced diet and water *ad libitum*. Bradykinin potentiating factor was previously isolated from the venom of the scorpion *Buthus occitanus* (Salman, 1995 and 2002) according to the chemical method of Ferreira (1965). The LD50 of this factor was determined according to the method of Meier and Theakston (1986). The animals were allocated into two main groups; non-irradiated and irradiated groups (100 individual each). The first main group (non-irradiated group) was subdivided into four sub-groups (25 animals each). The first sub-group was injected with saline solution (100 μ L) and considered as control. The second sub-group was i. p. injection 1 μ gm/gm b. w. of bradykinin potentiating factor

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isolated from the venom of scorpion, *Buthus occitanus* and was solvent in saline solution. The third sub-groups were i.p. injection of bone marrow cells from their twins (2.5×10^6 cells/animal). The fourth sub-group injected i. p of BPF ($1\mu\text{g}/\text{gm}$. b. w.) was dissolved in saline solution together with i. p. injection of bone marrow cells from their twins (2.5×10^6 cells/animal). Irradiated sub-groups were also divided into four sub-groups. Irradiated growing Guinea pigs males sub-groups were exposed to 150 Rads/animal by cesium-137 irradiation unit with a dose rate of 1.06483 rads per second in a uniform manner and divided into four sub-groups. The irradiated sub-groups were injected with the same manner of non-irradiated sub-groups

Biochemical analysis

Sample collection: Five animals were sacrificed from each sub-group of non-irradiated and irradiated at 1, 3, 7, 14 and 21 days post-irradiated and treatment with bradykinin potentiating factor. Peripheral blood was left in clean tubes at room temperature to clot. After an hour, serum was separated by centrifugation for 30 minutes at 3000 rpm. (Dacie and Lewis, 1975). The sera were collected in aliquots in labeled- Epindorff's tubes and stored at -20°C until used for determination of total protein and albumin.

Determination of total serum protein level: Determination of total serum protein level was estimated according to Peters (1968) method.

Reagents: Potassium-sodium-tartrate (15 mmol), sodium iodide (100 mmol), potassium iodide (15 mmol), copper sulphate (5 mmol) produced by LABKIT-Josep Tarradellas, cat. no. 80 08029 Barcelona, Spain code no. 30350.

Determination of serum albumin level: Serum albumin was determined according to the method described by Doumas *et al.* (1971 and 1972)

Reagent: Bromocresol green pH 4.2, 50 mmol produced by LABKIT-plató, 6 cat. no. E-08021 Barcelona, Spain, cod. No.30120. Using spectrophotometer (Spectronic 601 Item 335104 MB, Serl. 3614203016, Milton Ro. Company, 820 Linden Avenue, Rochester, NY 14625m U.S.A)

Statistical analysis

Data are presented as standard error (S.E.) of the means and were statistically analyzed using SPSS version 8.0 of Software. Differences between moments were analyzed by Kruskal-Wallis Test and parameters of each test group were compared with control group by Mann-Whitney Test. Results were considered significant when p value was lower than 0.05.

RESULTS

1) Serum total protein content:

Effect of BM cell, BPF or both in non-irradiated animals. The results presented in table (1) show that administration of B. M. cells (2.5×10^6 cells/animal) to the normal non irradiated guinea pigs did not induce significant difference change of their serum total protein content from their corresponding control values during the 21 days test period. On the other hand, i.p. injection of a single dose BPF isolated from the scorpion, *B. occitanus* venom to the normal animals induced a significant increase of their mean serum total protein content. This increase was a pronouncing and reached 93.13% ($P < 0.01$) after one day 34.7% ($P < 0.5$) after three days and 8.92% ($P < 0.05$) after 7 days of treatment respectively. However, after 14 and 21 days of treatment there was no significant change of the mean serum total protein level from that of the corresponding control value (Table 1). Furthermore, the combination treatment with both of BM cells and a single venom fraction BPF dose enhanced the

increase of mean serum total protein level determined after each of 24 h, 3 days and 7 days test period (Table 1). The increases recorded after 14 and 21 days were insignificant from the corresponding control value (Table 1).

2) Effect of whole body irradiation on serum total protein:

On the other hand, whole body irradiation of guinea pigs (1.5 Gy) induced a remarkable decrease in serum total protein level (29.06%; $P < 0.05$) after 24 h of irradiation. Such decrease was evident after 3 days (35.81%; $P < 0.05$) and after 7 days (65.05%; $P < 0.001$) of radiation exposure (Table 1). After 14 and 21 days of irradiation the mean serum total protein level was still significantly lower (63.62%; $P < 0.001$ and 60.16%; $P < 0.01$ respectively) than the corresponding control level (Table 1).

Table (1): Effect of intraperitoneal injection of a single dose ($1\mu\text{g}/\text{gm}$ body weight) of BPF from scorpion; *Buthus occitanus* (BPF), 2.5×10^6 BMCs or BMCs together with BPF on total serum protein level of non-irradiated and groups of growing male Guinea pigs at 1, 3, 7, 14 and 21 days post-treatment.

Days	Groups	Non-irradiated groups				Irradiated groups			
		Treatments	Control	BMCs	BPF	BMCs+BPF	I	I+BMCs	I+BPF
After One days	Mean \pm S.E. gm/dL	5.3104 \pm 0.163	5.4158 \pm 0.386	10.2558 \pm 0.390	11.5682 \pm 0.373	3.7672 \pm 0.264	4.1530 \pm 0.293	5.5022 \pm 0.288	7.01582 \pm 0.290
	%of change		+1.98	+93.13	+117.84	-29.06	-21.79	+3.61	+32.114
	Significance		NS	$P < 0.01$	$P < 0.001$	$P < 0.05$	$P < 0.05$	NS	$P < 0.05$
After 3 days	Mean \pm S.E. gm/dL	5.2042 \pm 0.188	5.4314 \pm 0.168	7.0100 \pm 0.165	8.6342 \pm 0.197	3.3406 \pm 0.199	4.3808 \pm 0.187	5.5048 \pm 0.199	5.886 \pm 0.180
	% of change		+4.37	+34.70	+65.91	-35.81	-15.82	+5.776	+13.10
	significance		NS	$P < 0.05$	$P < 0.01$	$P < 0.05$	$P < 0.05$	NS	$P < 0.05$
After 7 days	Mean \pm S.E. gm/dL	5.155 \pm 0.1856	5.2768 \pm 0.19169	5.615 \pm 0.108	6.0062 \pm 0.178	1.8016 \pm 0.193	2.5802 \pm 0.187	3.8032 \pm 0.180	4.3084 \pm 0.180
	%of change		+ 2.36	+ 8.92	+16.51	- 65.05	- 49.95	- 26.22	- 16.42
	significance		NS	$P < 0.5$	$P < 0.05$	$P < 0.001$	$P < 0.01$	$P < 0.05$	$P < 0.05$
After 14 days	Mean \pm S.E. gm/dL	5.230 \pm 0.144	5.044 \pm 0.181	5.250 \pm 0.121	5.3866 \pm 0.126	1.9027 \pm 0.185	2.630 \pm 0.150	3.1230 \pm 0.140	3.500 \pm 0.187
	%of change		-3.56	+0.38	+2.99	- 63.62	- 49.71	- 40.29	- 33.08
	significance		NS	NS	NS	$P < 0.001$	$P < 0.01$	$P < 0.01$	$P < 0.05$
After 21 days	Mean \pm S.E. gm/dL	5.2446 \pm 0.1730	5.280 \pm 0.110	5.448 \pm 0.194	5.368 \pm 0.193	2.089 \pm 0.138	2.957 \pm 0.171	3.408 \pm 0.150	4.982 \pm 0.180
	%of change		+0.67	+3.88	+2.353	- 60.17	- 43.62	- 35.02	- 5.01
	significance		NS	NS	NS	$P < 0.01$	$P < 0.01$	$P < 0.05$	NS

- N = 5 animals were used in each group.
- P = Significantly different from the control.
- NS = Insignificantly different from the control.
- BPF = Bradykinin potentiating factor (isolated scorpion venom fraction).
- BM = Bone marrow.
- I = Irradiated.

Effect of treatment with BM cells, BPF or both on irradiated animals: Treating the irradiated animals with BM cells 3 h after irradiation slightly ameliorated the total protein content to be 21.79% ($P < 0.05$) lower than that of the control level after 24 h of treatment. However, while this amelioration was complete after 3 days of radiation exposure it was also incomplete after 7, 14 and 21 days of treatment and total protein level was significantly lower than that of corresponding control value by 15.82% ($P < 0.05$), 49.95% ($P < 0.01$) and 49.71% ($P < 0.01$) respectively as shown in (Table 1). Treating the irradiated animals with a single dose of BPF showed also a recovery of the mean serum total protein content of guinea pigs and the level was 3.61% higher than that of the corresponding control value after one day of treatment. This effect was enhanced after three days and the increase of serum total protein level was 5.78%

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higher than that of the control value. After 7 days the total protein level was significantly lower from that of the mean control value, (26.22%; $P < 0.05$) as shown in table (1). This decrease of the serum total protein level of irradiated animals that were treated with a single dose of BPF was a significantly lower than that of the corresponding control after 14 days (40.29%; $P < 0.01$) and 21 days (35.02%; $P < 0.05$) respectively (Table 1). On the other hand, the combined BM cells and a single dose of BPF treatments of the irradiated guinea pigs resulted in remarkable increase of serum total protein level after one (32.11%; $P < 0.05$) and 3 days (13.11%; $P < 0.05$) of treatment in comparison with that the corresponding control animals as shown table (1). However, this amelioration was incomplete after 7 days of treatment and the mean serum total protein level was significantly lower than that of corresponding control value by 16.50% ($P < 0.05$) as shown in (Table 1). These results indicate a significant decrease also of serum total protein level due to this combined treatment by 33.08% ($P < 0.05$) and 5.0% after 14 and 21 days of irradiation respectively (Table 1).

Serum albumin content:

1) Effect of BM cell, BPF or both in non-irradiated animal in table (2) show that administration of BM cells (2.5×10^6 cells/animal) to the normal non irradiated Guinea pigs did not induce significant difference change of their serum albumin content from their corresponding control values during the 21 days test period. On the other hand, i.p. injection of a single dose BPF isolated from the scorpion, *B. occitanus* venom to the normal animals induced a significant increase of their mean serum albumin content. This increase was a pronouncing and reached 36.20% ($P < 0.01$) after one day 9.20% ($P < 0.5$) after three days and 8.92% ($P < 0.05$) after 7 days of treatment respectively. However, after 14 and 21 days of treatment there was no significant change of the mean serum albumin level from that of the corresponding control value (Table 2).

Table (2): Effect of intraperitoneal injection of a single dose ($1 \mu\text{g}/\text{gm}$ body weight) of BPF from scorpion; *Buthus occitanus* (BPF), 2.5×10^6 BMCs or BMCs together with BPF on serum albumin level of non-irradiated and irradiated groups of growing male Guinea pigs at 1, 3 and 7, 14 and 21 days post-treatment.

Days	Groups	Non-irradiated groups				Irradiated groups			
		Treatments	Control	BMCs	BPF	BMCs+BPF	I	I+BMCs	I+BPF
After One day	Mean±S.E. gm/dL	3.4088 ± 0.185	3.3132 ± 0.159	4.6428 ± 0.121	4.9944 ± 0.263	2.4198 ± 0.196	2.7146 ± 0.139	3.162 ± 0.176	3.669 ± 0.195
	%of change		-2.80	+36.20	+46.51	-29.01	-20.36	-7.240	+7.63
	Significance		NS	$P < 0.01$	$P < 0.01$	$P < 0.05$	$P < 0.05$	NS	NS
After 3 days	Mean±S.E. gm/dL	3.3626 ± 0.104	3.4382 ± 0.165	3.6720 ± 0.115	3.7442 ± 0.196	2.0076 ± 0.173	2.6076 ± 0.151	3.000 ± 0.197	3.0864 ± 0.160
	%of change		+2.25	+9.20	+11.35	-40.30	-22.45	-10.78	-8.21
	significance		NS	$P < 0.05$	$P < 0.05$	$P < 0.01$	$P < 0.05$	NS	NS
After 7 days	Mean±S.E. gm/dl	3.3626 ± 0.179	3.392 ± 0.121	3.572 ± 0.120	3.858 ± 0.171	1.2754 ± 0.119	1.754 ± 0.176	2.7164 ± 0.1293	2.9618 ± 0.112
	%of change		+0.87	+6.23	+14.73	-62.07	-47.84	-19.22	-11.92
	significance		NS	NS	$P < 0.05$	$P < 0.001$	$P < 0.01$	$P < 0.05$	NS
After 14 days	Mean ± S.E. gm/dL	3.4026 ± 0.176	3.32 ± 0.101	3.454 ± 0.179	3.5366 ± 0.147	1.000 ± 0.162	1.500 ± 0.113	1.700 ± 0.159	2.000 ± 0.169
	%of change		-2.43	+1.51	+3.94	-70.61	-55.92	-50.04	-41.22
	significance		NS	NS	NS	$P < 0.001$	$P < 0.01$	$P < 0.01$	$P < 0.01$
After 21 days	Mean ± S.E. gm/dL	3.3882 ± 0.111	3.578 ± 0.209	3.558 ± 0.295	3.446 ± 0.194	1.423 ± 0.171	2.100 ± 0.195	2.220 ± 0.186	3.000 ± 0.114
	%of change		+5.60	+5.01	+1.71	-58.00	-38.02	-34.48	-11.46
	Significance		NS	NS	NS	$P < 0.001$	$P < 0.01$	$P < 0.01$	$P < 0.05$

N = 5 animals were used in each group.

P = Significantly different from the control.

NS = Insignificantly different from the control.

BPF = Bradykinin potentiating factor (isolated scorpion venom fraction).

BM= Bone marrow.

I = Irradiated.

Furthermore, the combination treatment with both of BM cells and a single venom fraction BPF dose enhanced the increase of mean serum albumin level determined after each of 24 h, 3 days and 7 days test period (Table 2). The increases recorded after 14 and 21 days were insignificant from the corresponding control value (Table 2).

2) Effect of whole body irradiation on serum albumin:

On the other hand, whole body irradiation of guinea pigs (1.5 Gy) induced a remarkable decrease in serum total protein level (29.01%; $P < 0.05$) after 24 h of irradiation. Such decrease was evident after 3 days (35.81%; $P < 0.05$) and after 7 days (40.31%; $P < 0.01$) of radiation exposure (Table 1). After 14 and 21 days of irradiation the mean serum albumin level was still significantly lower (70.61; $P < 0.001$ and 58.00; $P < 0.01$) respectively than the corresponding control level (Table 2).

3) Effect of treatment with BM cells, BPF or both on irradiated animals:

Treating the irradiated animals with BM cells 3 h after irradiation slightly ameliorated the albumin content to be 20.36; ($P < 0.05$) lower than that of the control level after 24 h of treatment. However, while this improvement was complete after 3 days of radiation exposure it was also incomplete after 7, 14 and 21 days of treatment and serum albumin level was significantly lower than that of corresponding control value by 47.83% ($P < 0.05$), 55.91% ($P < 0.01$) and 38.02% ($P < 0.01$), respectively as shown in (Table 2). Treating the irradiated animals with a single dose of BPF showed also a recovery of the mean serum albumin content of Guinea pigs and the level was 7.24% lower than that of the corresponding control value after one day of treatment. However, while this improvement was complete after 3 days of radiation exposure it was also incomplete after 7, 14 and 21 days of treatment and serum albumin level was significantly lower than that of corresponding control value by 19.21% ($P < 0.05$), 50.04% ($P < 0.01$) and 34.48% ($P < 0.01$) respectively as shown in (Table 2). On the other hand, the combined BM cells and a single dose of BPF treatments of the irradiated guinea pigs resulted in increase of serum albumin level after one day (7.63%). However, this amelioration was incomplete after 3 and 7 days of treatment and the mean serum total protein level was significantly lower than that of corresponding control value by 8.21% and 11.92% respectively as shown in (Table 1). These results indicate a significant decrease also of serum albumin level due to this combined treatment by 41.22% ($P < 0.05$) and 11.45% ($P < 0.05$) after 14 and 21 days of irradiation respectively (Table 2).

DISCUSSION

Recovery from irradiation effects has strongly attracted the attention of radiobiologist. They tried different means of treatment against irradiation effects as chemical and biological substances (Khamis, 1986 and Dong *et al.*, 2009). This paper deals with the trial of using one of the most important biological cells in the body (bone marrow cells, bradykinin potentiating factor and combination bone marrow cells and bradykinin potentiating factor) for treatment post-irradiation. The liver is a major producer for most of serum proteins and its total level in the blood is a main liver function test. Irradiation has a destructive effect on all body tissues including the liver. This is very important in the context of the present studies considering the fact that the liver is the major source for endogenous kininogens (El-Asmar *et al.*, 1975; Kehrer *et al.*, 1990 and Abu-Amra and Abd-El-Rahim, 2000). It was reported that BPF injection causes activation of glycogen synthetase enzyme that increases

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liver content of glycogen. This increase has a liver protective effect and potentiates the detoxification efficiency of the liver (Basu *et al.*, 1992; Abu-Amra, 1994; Muriel, 1998 and Schmaier *et al.*, 1998). Furthermore, the hepatic synthesis of a number of plasma protein increased by IL-1 that is bradykinin-dependent in its secretion (Oppenheim *et al.*, 1982; Tiffany and Burch, 1989; Zündorf *et al.*, 1996 and Öztürk, 2001). It was noted that the irradiation-induced reduction in serum total protein and albumin were improved by the different treatments applied in the present study. BPF effect was dose-frequency dependent improvement the concentration total protein and albumin but did not reach complete recovery. Combined BPF and BM cells treatment was more efficient in normalizing serum total protein and albumin. It is worthy to mention that, radiation-induced renal injury (radiation nephropathy) is characterized by proteinuria, hypertension, and progressive decline in renal function (Cohen, 2001; Cohen *et al.*, 2002 and Cohen and Robbins, 2003). In the present study the significant decrease in the serum level of total protein and albumin after irradiation was in agreement with previous results (Marinova *et al.*, 1991). It is established that liver is the main source of plasma albumin. The decreases in serum total protein and albumin is mainly due to the diminishing of its synthesis in hepatic cells, accompanied by losses of large amounts of albumin into the urine and gastrointestinal tract due to damage kidney and intestinal mucosa (West, 1985). Karalenka *et al.*, (1993) found that gamma irradiation leads to cessation of the transport of albumin into rat plasma. They concluded that the speed and the characters of that radiation effect depend both on dose and duration after irradiation. Albumin catabolism (Simsa and Mráz, 1974 and Waldmann, 1977) may differ fundamentally from the catabolism of the other plasma protein, since albumin is duration withdrawal the only plasma protein that is not a glycoprotein (Sharon and Lis, 1981). On the other hand, there are interactions between vasoactive polypeptides such as bradykinin and classical hormonal transmitters. Example of such interactions is that active polypeptides stimulate the synthesis or release of prolactin and growth hormone (Vijayan *et al.*, 1979; Enjalbert *et al.*, 1980; Frawley and Neil, 1981 and Chihara *et al.*, 1982). More recently, Dong *et al.* (2009) reported that, the effect of scorpion venom polypeptide accelerating the hematopoietic recovery was potentially through a mechanism of stimulating the release of cytokines. Consequently, it is could be suggested that the endogenous bradykinin potentiated by venom fractions enhanced the release of growth hormone, cytokine and other growth factors, resulting in the stimulation of increase total protein and albumin. In conclusion the improvement of these parameters may be attributed to the induction of hematopoietic growth factors and/or amelioration of the injury effect of radiation on the liver that necessitates future investigation.

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ARABIC SUMMARY

تأثير جرعة واحدة من مفضل من سم العقرب بوشس اوكسيتانس كعامل منشط للبراديكينين علي البروتين الكلي والالبومين في مصل خنازير غينيا النامية المشعة

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يهدف هذا البحث دراسة تأثير مفضل من سم العقرب كعامل منشط للبراديكينين علي مصل حيوانات خنازير غينيا النامية المشعة ذلك لمعرفة مدي قدرته في التأثير علي البروتين الكلي و الالبومين في مصل الحيوانات كدلائل للشفاء من آثار الإشعاع المدمرة .
ومن ثم فقد تم تقسيم الحيوانات الي مجموعتين رئيسيتين إحداهما غير مشعة، والأخرى مشعة. ثم تم تقسيم المجموعة الرئيسية الغير مشعة إلي أربع مجموعات فرعية:-
(١) مجموعة فرعية غير مشعة حقنت بمحلول فسيولوجي، و تركت بدون إشعاع أو علاج (مجموعة ضابطة).
(٢) مجموعة فرعية غير مشعة حقنت بخلايا نخاع العظم من توأم الحيوانات.
(٣) مجموعة فرعية غير مشعة حقنت بواحد ميكرو جرام/جرام من المستخلص السمي.
(٤) مجموعة فرعية غير مشعة حقنت بواحد ميكرو جرام من المستخلص السمي، وخلايا نخاع العظم.
إما المجموعة الرئيسية الثانية المشعة فقد قسمت أيضاً إلي أربع مجموعات:-
(١) مجموعة فرعية مشعة حقنت بالمحلول الفسيولوجي.
(٢) مجموعة فرعية مشعة حقنت بخلايا نخاع العظم من توأم الحيوانات.
(٣) مجموعة فرعية مشعة حقنت بواحد ميكرو جرام/ جرام من المستخلص السمي.
(٤) مجموعة فرعية مشعة حقنت بواحد ميكرو جرام من المستخلص السمي، وخلايا نخاع العظم . ثم ذبح من كل مجموعة فرعية خمس حيوانات بعد مرور ١، 3، ٧، ٢١ يوم من التعرض للإشعاع، والعلاج بالمستخلص السمي، ونخاع العظم، و أعد المصل لتقدير الألبومين، والبروتين الكلي في جميع المجموعات.
ولقد أظهرت نتائج التحليل البيوكيماوية أن تركيز البروتين الكلي، والألبومين في المجموعات التي شععت، وعولجت بالمستخلص السمي تحسناً، وظل هذا التحسن لمدة ١٤ يوم بعد العلاج. أما المجموعة التي عولجت بخلايا نخاع العظم، والمستخلص السمي فكانت هي الأكثر تحسناً. بيد أن الإشعاع أدي إلي انخفاض حاد في مستوي تركيز البروتين الكلي والألبومين في المجموعة التي حقنت، وشععت بأشعة جاما، وظل هذا الانخفاض إلي نهاية التجربة (٢١ يوماً). وهذا يدل علي أن لهذا المستخلص دور في تحسين البروتين الكلي، والألبومين من تأثير الإشعاع المدمر، وربما يكون ذلك من خلال إطلاق عوامل النمو والسيتوكينات. مما يرغب في المزيد من الدراسة والتوضيح في هذ المضمار.