



SOME ENZYMATIC ACTIVITY IN SERUM WITH CHRONIC RENAL FAILURE

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ABSTRACT

The research was concerned with measuring the activity of the following enzymes: Paraoxinase (PON1), proilase (PRO), aryl esterase (Ary), Lactonase, Glutathion – Stransferase (GsT), polymine oxidase (PAO), Glucoolatonase (GIN), Nitric oxide (NO) and Superoxide dismutase (SOD), in serum with chronic renal failure (CRF) undergoing hemodialysis, seventy nine patients with CRF before heomdialysis and eighty healthy controls were in clouded in this study. Two groups of CRF patients and healthy subjects were divided according to age (20 – 40) and (40 – 65) respectively. The result showed a significant decrease in serum PONI, Ary, Lactonase (Lac),

GIN, NO, SOD, ADA activities in CRF patients while PAO, GST increase significantly in serum patients compared to normal individuals.

KEYWORD: PONI, PR, Ary, Lactonase, GST, PAO, ADA, NO, SOD, CRF.

- This work is taken from her ph. D thesis

1- INTRODUCTION

Chronic kidney disease (CKD) is a worldwide public health problem, with dverse outcomes of kidney failure and premature death. CKD is defined as kidney damage of glomerular filtration rate (GFR) $<60\text{ml /min/ } 1.73\text{m}^2$ for 3 months or more, irrespective of cause. Kidney damage in many kidney diseases can be a scertained by the presence of albuminuria, defined as albumin – to creatinine ratio $>30\text{mg/g}$ in two of three spot urine specimens. GFP can be estimated from calibrated serum creatinine and estimating equation, (Levey *et al.*, 2005).

PONI Enzyme

The enzyme paraoxonase (PON1; EC. 3.1.8.1) detoxifies activated (Oxon) organ phosphorus pesticides (OPS) which are neurotoxic. PONI also is an antioxidants and several studies indicate that it inhibits oxidation of Low – density Lipoprotein, a marker of oxidative stress. (Karen, *et al.*, 2009).

The enzyme arylesterase (Ary) (EC3.1.1.2) showing paraoxonase activity were less understood than other esterase, especially carboxylesterase and lipase until recently most of the arylesterase and/ or paraoxonase reported were from mammalian sources. Which plays important roles in the detoxification of the organophosphoric compounds as well as in the prevention of atherosclerosis (Aharon, *et al.*, 2004; Gur, *et al.*, 2007).

The enzyme ADA is essential and widely distributed enzyme of purine catabolism that catalyzes the hydrolytic deamination of adenosine and deoxyadenosine to inosine and deoxyinosine. (Black, *et al.*, 1996).

The PAO (PAO EC1.5.3.11) are FAD – dependent enzymes involved in poly amine catabolism (Con *et al.*, 2006) they catalyze the oxidation of amines to produce the aldehyde and H₂O₂. (Pohicelli, *et al.*, 2012).

The enzyme GIN is a 34 protein whose tissue levels in the liver, kidney, and lung (Hasegawa, 2010). And GIN which functions enzymatically as an alactonase hydrolyzes various carbohydrate lactonase. (Fujil, *et al.*, 1999).

The GST (GST EC. 2.5.1.18) enzyme catalyzes the covalent conjugation of glutathione (GSH) to electrophilic substrates. GSH conjugation inactivates electrophilic molecules. (Guenenger *et al.*, 1987).

By replacing the electronegative functional group with sulfur atom of glutathione. GST catalyzes covalent conjugation of GSH via the cysteine thiolate of GSH. (Hayes *et al.*, 2005) copper/zinc superoxide dismutase (Cu/Zn – SOD). Which catalyzes the conversion of superoxide anion into hydrogen peroxide and works concomitantly with hydrogen peroxide. Removing enzymes such as catalase and glutathione peroxidase (Aymelek *et al.*, 2002).

Nitricoxide (NO) is one of the main factors involved in the anti atherosclerotic effects of endothelium several investigators have focused their attention on a possible role played by NO in the development of. Uremic symptoms. (Sarkar, *et al.*, 2004).

The PR enzyme. An enzyme found in human and animal tissues, plays an important role in collagen catabolism and synthesis. (Karna, *et al.*, 2000; Myara, *et al.*, 1984).

2- MATERIAL AND METHODS

This research was conducted on a random sample of patients with CRF who attended the artificial kidney unit in BN – SINA hospital in Nineveh.

And the account patients (87) male and females ranging in their age between 20 – 65 years these samples were diagnosed clinically and radiologically as having CRF undergoing hemodialysis.

Parameters	References
PON (U/L)	Pock, <i>et al.</i> , 2008
Ary (U/L)	Rock, <i>et al.</i> , 2008
Lactonase (U/L)	Rock, <i>et al.</i> , 2008
PR (U/L)	(Myara., <i>et al.</i> , 1984)
ADA (U/L)	(Glustili, 1981)
PAO (U/L)	(Flayeh, 1988; Alkatib, 2000)
GST (U/L)	Habig <i>et al.</i> ,(1974)
SOD (U/L)	(Brown and Goldsrin, 1983)
NO (U/L)	(Green, <i>et al.</i> , 1982)
G/N (U/L)	Hucho and Wallenfels (1972)

RESULTS AND DISCUSSIONS

The results in table (1) showed that is a significant decrease ($p < 0.05$) in PON1 activity in patients in comparison with control the results of the present study were agree with (Dantonine, *et al.*, 1998 Para G. 1998).

Also results show that PON1/Ary activities are reduced in patients with CRF (Mockness ML., *et al.*, 1993). And agree with (La Du, *et al.*, 1993; Mackness, *et al.*, 1996; Durrington, *et al.*, 2001; Aviram, *et al.*, 1999; Gasselwander, *et al.*, 1999). While few studies has previously pointed out findings about the level ADA activity in uremic patients especially those suffering from CRF in this study, table (1) and this results with agree (Sufrin, *et al.*, 2006).

Also the results show there was a significant decrease in PR activity in CRF compared with normal subjects. This agrees with (Gehyof *et al.*, 1983) also found higher activity in PAO activity in CRF patients. This agrees with (Lgarashi, *et al.*, 2006). While also we found increase this agrees with (Founier, *et al.*, 1992) also the results show the enzyme lactonase is decreased in patients with CRF compared controls (table). This result agrees with (Teber, *et al.*, 2003).

Also in this study was found that GIN is lower in patients with CRF and NO any studies about it. Also this study shows that NO is 10% lower in CRF patient compared with control this agrees with (Saran, *et al.*, 2003; Baylis, *et al.*, 2006). While also there is a significant decrease $p(<0.05)$ in SOD activity. This is in agreement with (Vaziri, *et al.*, 1984, Nagane, *et al.*, 2009).

We found a significant positive correlation between paroxonase and all enzymes except enzymes GST a negative correlation at the level 0.01 table (2) while found a negative correlation between enzyme lactonase and all enzymes above except enzymes GST and prolidase positive correlation table (2). Also found a negative correlation between enzyme prolidase and all enzymes above except enzyme Lac positive correlation. Also found a negative correlation between PAO and all enzymes except Lac and PR table (2). Also found NO correlation between all enzymes and NO, SOD. Also we found positive correlation between ADA and all enzymes except PON1. A negative correlation table (2).

Table (1) theenzymes activity and concentration of the parameters in blood serum of hemodialysis patients and control subjects.

Rang of age	N	PON1 U/L	PR U/L	Lca U/L	PAO U/L	G/N U/L	ADA U/L	Ary U/L	G/N U/L	NO U/L	SOD U/L
40 – 65 patients	43	0.064 \bar{f} 0.001h	34.197 \bar{f} 0.427c	38.566 \bar{f} 1. 315a	15.12 \bar{f} 0.500e	0.159 \bar{f} 0.0055a	0.144 \bar{f} 0.0047a	15.07 \bar{f} 0.23a	6.85 \bar{f} 0.15e	443 \bar{f} 8.3a	3.233 \bar{f} 0.33b
40 – 65 normal	42	0.129 \bar{f} 0.002a	45.28 \bar{f} 0.49d	73.544 \bar{f} 0.924b	6.286 \bar{f} 0.233c	1.559 \bar{f} 0.041c	2.521 \bar{f} 0.0404b	24.77 \bar{f} 0.483b	1.5412 \bar{f} 0.036a	46.707 \bar{f} 3.2b	3.96 \bar{f} 0.36c
20 – 40 patients	40	0.174 \bar{f} 0.003d	26.025 \bar{f} 0.35	140.174 \bar{f} 1.924d	2.4505 \bar{f} 0.054q	0.13G \bar{f} 0.0054b	2.48 \bar{f} 0.0549a	35.193 \bar{f} 1.28c	13.365 \bar{f} 0.235e	56.6 \bar{f} 0.78c	4.15 \bar{f} 0.38d
20 – 40 normal	44	0.246 \bar{f} 0.004e	65.40 \bar{f} 1 .18f	179.263 \bar{f} 71.653e	3.332 \bar{f} 0.049ab	2.423 \bar{f} 0.039e	3.630 \bar{f} 0.045d	76.293 \bar{f} 2.4e	2.399 \bar{f} 0.0858b	58. \bar{f} 0.65D	4.9 \bar{f} 0.15C
Total patients	79	0.153 \bar{f} 0.002c	30.05 \bar{f} 0.56b	91.298 \bar{f} 5. 867e	8.622 \bar{f} 0.762d	0.264 \bar{f} 0.012ab	0.1477 \bar{f} 0.007a	25.80 \bar{f} 1.38b	10.1905 \bar{f} 0.3967d	43.2 \bar{f} 8.1ab	4.5 \bar{f} 0.26cd
Total normal	86	0.159 \bar{f} 0.10dc	50.54 \bar{f} 0.819e	127.633 \bar{f} 75.810d	4.739 \bar{f} 0.194b	1.996 \bar{f} 0.055d	3.0089 \bar{f} 0.067c	49.24 \bar{f} 3.19d	1.9805 \bar{f} 0.066ab	55.5 \bar{f} 0.64ad	3.5 \bar{f} 0.245d

The similar litters means no significant at the level $p < 0.05$.

The different litters means significant level $P < 0.05$.

Table (2) correlation between enzymes in patients with chronic renal failure correlation is significant at the 0.01.

	SOD	NO	ARY	GST	GLN	ADA	PAO	PRO	LAC	PON1
0.6445	0.866	0.832	-0.955	0.514	0.538	0.671	0.975	0.780	0.000	PON1
-0.703	0.604	-0.659	0.771	-0.778	-0.777	-0.745	0.639	0.000	-0.780	LAC
0.608	-0.733	-0.218	-0.972	-0.974	-0.974	-0.876	0.000	0.639	-0.952	PRO
-0.658	-0.89	0.663	-0.607	-0.658	-0.664	0.000	0.876	0.745	-0.671	PAO
-0.633	-0.692	0.831	0.952	0.404	0.000	0.664	0.974	0.777	-0.538	ADA
-0.848	-0.77	0.831	-0.901	0.000	-0.404	0.658	0.974	0.778	-0.514	GLN
-0.584	-0.81	0.821	0.000	0.901	-0.952	0.607	0.972	-0.771	0.953	GST
0.499	-0.692	0.000	-0.821	-0.828	-0.831	-0.663	0.218	0.659	-0.832	ARY
0.599	0.000	0.692	0.81	0.77	0.692	0.89	0.733	-0.604	-0.866	NO
0.000	-0.599	0.499	0.584	0.848	0.633	0.658	-0.608	0.703	-0.6445	SOD

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