

Adipose Tissue Remodeling : The Link Between FGF23-Klotho-Vitamin D3 Axis and Renin-Angiotensin-Aldosterone Axis In The Development And Progression of Obesity Related Kidney Disease

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METHODS

OBJECTIVES

Obesity is established as an important contributor of increased diabetes mellitus, hypertension and cardiovascular disease, all of which can promote chronic kidney disease (CKD). Recently, there is a growing appreciation that even in the absence of these risks, obesity itself significantly increases CKD and accelerates its progression. The aim of this work is to evaluate the link between Renin-Angiotensin-Aldosterone System (RAAS) and FGF23-Klotho-1,25D3 axis and their impact in obese and non-obese CKD patients.

In a cross sectional randomized multi centers study ,two hundred twenty six CKD patients stage III and IV (eGFR 20-60 ml/min/m²) have enrolled in this study as follows : group I ; 87 non diabetic CKD patients aged 20-40 years with body mass index (BMI) between 20-25 kgm/m²; group II ; 130 non diabetic CKD patients aged 20-40 years with (BMI) > 30 kgm/m² and group III ; 89 CKD patients aged >60 years. All patient have been tested for plasma leptin levels, 1,25 dihydrocholiciferole (1,25D3), plasmaparathormone (PTH) Serum calcium (Ca), serum phosphorus (PO₄), and plasma FGF-23 , plasma renin activity (PRA), plasma angiotensinogen receptor 1 & 2 (AT1 & AT2) and plasma aldosterone (ALD) .

References:

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2. Martin H. de Borst, Marc G. Vervloet, piet M. ter Wee, and Gerjan Navis. Cross Talk Between the Renin-Angiotensin-Aldosterone System and Vitamin D-FGF-23-klotho in Chronic Kidney Disease. *J Am Soc Nephrol*. 2011 September; 22(9): 1603–1609.

CONCLUSIONS

Obesity per se is an independent risk factor for the development and progression of chronic kidney disease specially in young age patients.

	Group I lean 20-40 y N = 87	Group II Obese 20-40 y N = 130	Group III Old Age > 60 y N = 89	P value Group I Vs Group II	P value Group II Vs Group III	P value Group I Vs Group III
eGFR ml/min/m ²	49.3 ± 7.51	37.71 ± 13.6	41.0 ± 13.47	P < 0.01	P < 0.05	P < 0.05
Diabetes %	Nil	Nil	36 %	---	---	---
PRA pg/ml/hr	4.21 ± 2.07	2.08 ± 1.45	3.46 ± 1.36	P < 0.05	P < 0.05	P = 346
ALD pg/ml	47.55 ± 21.8	89.91 ± 22.6	86.18 ± 16.8	P < 0.001	P = 0.34	P < 0.05
ALD/PRA ratio	11.29 ± 4.1	43.23 ± 14.9	24.91 ± 12.1	P < 0.001	P < 0.01	P < 0.01
AT1 pg/ml	290.3 ± 210.6	653.3 ± 352.6	407.1 ± 370.4	P < 0.001	P < 0.01	P < 0.05
AT2 pg/ml	25.46 ± 11.4	48.1 ± 4.8	33.68 ± 7.63	P < 0.001	P < 0.01	P < 0.05
PWV m/sec.	9.48 ± 0.8	9.31 ± 0.9	10.21 ± 0.68	P = 0.334	P < 0.05	P < 0.05
S Ca mg/dl	9.19 ± 0.64	9.27 ± 0.88	9.24 ± 0.49	NS	NS	NS
S PO ₄ mg/dl	3.85 ± 0.92	4.74 ± 1.61	4.09 ± 0.42	P < 0.05	NS	NS
PTH pg/ml	59.18 ± 24.7	77.63 ± 32.4	70.9 ± 15.3	P < 0.01	P = 336	P < 0.05
1,25 D pg/ml	24.4 ± 4.51	19.85 ± 3.6	25.0 ± 5.81	P < 0.05	P < 0.05	NS
S Leptin fg/l	1.92 ± 1.61	24.13 ± 7.81	5.51 ± 3.21	P < 0.001	NS	P < 0.01
FGF23 R u/ml	132.8 ± 126.1	259.6 ± 138.6	179.3 ± 237.4	P < 0.001	P < 0.01	P < 0.05
P insulin uU/ml	5.59 ± 2.31	13.73 ± 2.38	10.7 ± 1.68	P < 0.01	P < 0.05	P < 0.01
S albumin gm/l	37.53 ± 4.5	39.1 ± 3.81	37.48 ± 4.2	NS	NS	NS

Table (2) : Mean ± SD; NS, non-significant; eGFR, estimated glomerular filtration rate; PRA, plasma renin activity; ALD, aldosterone; AT1 & AT2, angiotensin receptor 1 & 2; PWV, pulse wave velocity; S. Ca, serum calcium; S. PO₄, serum phosphate; PTH, parathormone; 1,25D3, 1,25 dihydrocholiciferol; S. Leptin, serum leptin; FGF23, fibroblast growth factor 23; plasma insulin; serum albumin.

