

New biomarkers in kidney donors, in patients with chronic kidney disease, and in a healthy control population

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OBJECTIVES

Background: 27% of kidney transplantations in Norway are from living donors(1). Concern is raised about increased mortality and increased risk for developing ESRD in kidney donors (2).

Aim: To evaluate the serum levels of the new biomarkers neutrophil gelatinase-associated lipocalin (NGAL), soluble Klotho (sKlotho) and fibroblast growth factor 23 (FGF23) in kidney donors with normal kidney function, in healthy controls and in patients with CKD stages 3-5.

METHODS

Cross-sectional, observational single-center study including 40 kidney donors with an estimated glomerular filtration rate (eGFR) ≥ 60 ml/min/1.73m², 22 CKD stage 3 patients (eGFR 30-59 ml/min/1.73m²), 18 patients with CKD stage 4 (eGFR 15-29 ml/min/1.73m²), 20 patients with CKD stage 5 (eGFR < 15 ml/min/1.73m²) and 35 healthy controls, Table 1.

Serum NGAL, sKlotho and FGF23 were measured by commercially available ELISA kits from Bio-Porto Diagnostics, Gentofte, DK, Immuno-Biological laboratories (IBL), Fujioka, Japan and Kainos Laboratories Inc., Tokyo, Japan respectively.

Group:	Control	Donor	CKD stage 3	CKD stage 4	CKD stage 5	P-value
	n=35	n=40	n=22	n=18	n=20	
NGAL (ng/ml)	97.6±18.3	110.6±31.7	209.2±67.5	459.5±111.3	1012.6±374.1	.000
FGF23 (pg/ml)	51.8 (25.9-90)	62.1 (6.6-112)	97.5 (44-308)	337.0 (139-11000)	806.0 (121-16100)	.000
sKlotho (pg/ml)	736.7± 170	660.8±191	628.7±153	478.4±123	415.1±149.3	.000
25 (OH) VitD (nM)	58.67±25.2	77.73±17.5	44.01±19.4	53.55±22.5	59.65±29.5	.000
eGFR (ml/min)	99.0±13.1	73.6±13.1	43.7±9.8	19.1±5.8	7.3±2.6	.000
Creatinine (umol/L)	73.1±12.6	91.8±16.5	141.0±28.1	280.1±84.2	621.2±203.6	.000

RESULTS

NGAL was higher in donors compared to healthy controls (110.6 ng/ml \pm 31.7 vs 97.6 ng/ml \pm 18.3 p<0,05), and increased significantly with declining kidney function, Table 1.

FGF23 levels were non-significantly higher in donors compared to controls, and increased significantly with declining kidney function.

sKlotho levels were significantly lower in CKD stages 4 and 5 compared to controls but no difference was revealed between controls and kidney donors.

There was no significant difference in levels of calcium, phosphate, PTH or FePO₄ between donors and controls.

CONCLUSIONS

Kidney donors have significantly higher levels of NGAL than healthy controls. This may reflect a partial renal loss-of-function in kidney donors compared with a healthy control group. Renal hyperfiltration as a consequence of the nephrectomy, may induce cellular stress.

NGAL could be a valuable marker for predicting donors at risk of developing CKD and premature mortality.

References

1 Leivestad, Annual report 2013 The Norwegian Renal Registry
<http://www.nephro.no/nmr/AARSM2013.pdf>
 2 Mjoen et.al, Kidney Int 2014; vol 86: 162-7

