A study on the relationship between parameters of renal osteodystrophy and the development of peripheral and aortic vascular calcification in chronic renal failure patients

Thesis

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By Fadia Moris Bolis M.B.B.Ch

Cairo University

Supervised by Professor Dr/ Tarek Mahmoud Hussein Elshabony

Professor of Internal Medicine and Nephrology Faculty of medicine - Cairo University

Professor Dr/ Dr.Mohamed NasrAllah

Assistant Prof. of Internal Medicine and Nephrology Faculty of medicine - Cairo University

Professor Dr/ Amal Rashad ELShehaby

Assistant Prof. of Biochemistry Faculty of medicine - Cairo University

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To my great Parents,

(Two true gems in my life)

To my great brother Fady

And

To my dear husband Akram

Abstract

Background and aims:

Hemodialysis (HD) patients belong to the group of patients with a high prevalence of cardiovascular disease. Atherosclerosis is the most frequent cause of cardiovascular morbidity in patients with end-stage renal disease (ESRD). There is a high prevalence of vascular calcification among ESRD patients on hemodialysis and this is correlated to the presence of preexisting cardiovascular disease.

The aim of the present study was to test the correlation between various calcification scores and cardiovascular morbidity in patients with end stage renal disease on dialysis.

Subjects and methods:

91 ESRD patients on regular hemodialysis at Kasr ElAini Hemodialysis Unit and Kasr ElAini Nephro-Urology Unit (King Fahd Unit) were recruited to participate in the study in observational period 18 months. The included patients were 44 males and 47 females . Patients below age 18 years and pregnant women were excluded.

All subjects incorporated in the study were evaluated thoroughly by history taking, clinical examination and routine laboratory investigations for hemodialysis patients. Vascular calcification was assessed by non-contrast CT abdomen ,lateral lumbar x ray and x rays of both hands and pelvis.

Results:

We found a high prevalence of vascular calcification using six different radiological scores among ESRD patients on hemodialysis which increased progressively from proximal to distal along the abdominal aorta. This prevalence was higher in our study when measured by CT compared to plain X rays. Frequency distribution of vascular calcification by CT thorax score was 24.3%, CT suprarenal score was 33.7%, CT infrarenal score was 47.2%, CT bifurcation score was 42.3%, Kauppila score was 24.4%, and by Adraogo score was 34.1%.

We also found a positive correlation between calcification scores and diabetes mellitus, age and other factors correlating to the presence of vascular calcification including phosphate level, triglycerides, parathyroid hormone and dialysis adequacy as measured by kt/v urea.

We also found that cardiovascular disease in our study population was correlated in logistic regression analysis with age (P value = 0.001) and BMI (P value = 0.02) and negatively correlated with Kt/v urea (P value = 0.006).

We also found that vascular calcification scores were also correlated positively with the presence of cardiovascular disease.

4 Conclusions:

There is a high prevalence of vascular calcification among hemodialysis patients and this is more readily predicted by CT based technique. This vascular calcification correlated positively with the presence of pre-existing cardiovascular disease.

Key words: Chronic kidney disease-Cardiovascular disease-Vascular calcification.

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List of abbreviations

AACS Abdominal Aorta Calcification score

AB Apoptotic bodies

ABI Ankle-brachial index

ACC American College of Cardiology

ACEi Angiotensin converting enzyme inhibitor

ACI Aortic calcification index

ADMA Asymmetric dimethylarginine

AF Atrial fibrillation

A.G.E Advanced glycation end products

AHA American Heart Association

ALP Alkaline phosphatase

AMI Acute myocardial infarction

AngII Angiotensin II

ANP Atrial natriuretic peptide

AoAC Aortic arch calcification score

AOPPs Advanced oxidation protein products

AVC Aortic valve calcification

AVG Arteriovenous grafts

BCP Bony calcium phosphate

BNP Brain natriuretic peptide

CAC Coronary artery calcium scores

CAD Coronary artery disease

CAPD continuous ambulatory peritoneal dialysis

Cbfa1 Core-binding factor 1alpha

CETP Cholesteryl ester transfer protein

CHD Coronary heart disease

CKD Chronic kidney disease

CO Cardiac output

CRP C-reactive protein

CVD Cardiovascular disease

CVSSC Cardiovascular Special Studies Center

DCOR Dialysis Clinical Outcomes Revisited

DDAH Dimethylarginine dimethylaminohydrolase.

EBCT Electron beam computed tomography

ECSOD Extracellular- superoxide dismutase

E.D Endothelial dysfunction

eNOS Endothelial NO synthase

ESRD End-stage renal disease

FFA Free fatty acid

FGF23 Fibroblast growth factor 23

Fig. Figure

GFR Glomerular filtration rate

Hb Haemoglobin

HbA1c Glycated hemoglobin

HD Hemodialysis

HDL High density lipoprotein

HF Heart failure

HOMA Homeostasis model assessment method

HSMC Human Smooth muscle cell

HTN Hypertension.

ICAM Intercellular adhesion molecule.

ICD Implantable cardioverter-defibrillator

IDDM Insulin dependent diabetes mellitus

IE Infective endocarditis

IHD Ischemic heart disease

IL Interleukin

IVUS Intravascular ultrasound

KEEP Kidney Early Evaluation Program

LAV Left atrial volume

LDL Low density lipoprotein

Lp(a) Lipoprotein(a)

LVH Left ventricular hypertrophy

LVMI Left ventricular mass index

MAC Mitral annular calcification

MDA Malondialdehyde

MetS Metabolic syndrome.

MGP Matrix Gla protein

MI Myocardial infarction

MPO Myeloperoxidase

MRA Magnetic resonance angiography

MSCT Multislice computed tomography

MV Microvesicles

MWFS Midwall fractional shortening

NCBPs Nitrogen-containing bisphosphonates

NO Nitric oxide

NT-pro-BNP N-terminal pro-brain natriuretic peptide

OPBs Calcium-based oral phosphate binders

OPG Osteoprotegerin

OPN Osteopontin

 $O_{2^{T}}$ Superoxide anion radical

OxLDL Oxidized LDL

PAD Peripheral arterial disease