- 1.Hemodialysis,cardiovascular events (age, sex)
- 2.Coronary artery calcification (CAC), atherosclerotic vascular disease, predict future adverse cardiovascular (dialysis Therapy)
- 3.The prevalence of CAC(CRF) ?
- 4.Electron-beam computed tomography, high speed spiral computed tomography (CT), scoring calcified plaques in coronary arteries.

### **METHODS**

- 1.Consecutive outpatients: glomerular filtration rate (GFR) unchanged for at least 12 months
- 2.Controls: (ie,healthy volunteers and patients with essential hypertension and normal renal function).
- 3.Exclusion criteria :symptoms of heart failure or coronary artery disease, previous history of myocardial infarction, coronary bypass surgery, angioplasty; stroke, diabetes, rapidly progressive renal disease, arrhythmia (that would exclude gating during spiral CT).

## METHODS

- 1. CT : model CT Light Speed Plus multislice spiral scanner
- 2.The left main coronary artery, left anterior descending artery, circumflex artery, right coronary artery, and cardiac valves were scored.
- 3. A technician and radiologist

### **METHODS**

 1. calcium, phosphorus, intact parathyroid hormone (iPTH), homocysteine, and C-reactive protein (CRP).

serum calcium, phosphorus, and iPTH: Kidney Disease Outcomes Quality Initiativem (K/DOQI) guidelines.

- 2.GFR : clearance of creatinine.
- 3.calcium score categories :
  - (1) up to 100,
  - (2) 101 to 400,
  - (3) 401 to 1,000,
  - (4) 1,000 or greater.

Table 1. Clinical and Biochemical Characteristics of Patients With CRF and Controls

|  | Normal Value | Patients With CRF (n = 85) | Controls<br>(n = 55) |
|--|--------------|----------------------------|----------------------|
|  |              |                            |                      |
| Sex (% men)                                  |              | 87                         | 79                   |
| Age (y)                                      |              | 52 ± 13                    | 47 ± 11              |
| Body mass index (kg/m²)                      |              | $27.2 \pm 3.9$             | $27.5 \pm 3.4$       |
| Duration of CRF (y)                          |              | $6.3 \pm 5.6$              | _                    |
| GFR (mL/min)                                 | 90-120       | 33 ± 16                    | 99 ± 10*             |
| Urea (mg/dL)                                 | 18-53        | $97 \pm 40$                | 38 ± 6*              |
| Glucose (mg/dL)                              | 60-110       | 92 ± 7                     | 94 ± 6               |
| iPTH (pg/mL)                                 | 10-75        | 143 ± 121                  | 35 ± 17*             |
| Calcium (mg/dL)                              | 8.9-10.3†    | $9.5 \pm 0.5$              | 9.5 ± 0.2            |
| Phosphorus (mg/dL)                           | 3.0-4.5†     | $4.1 \pm 0.9$              | $3.8 \pm 0.8$        |
| Calcium-phosphorus product (mg²/dL²)         |              | $38.7 \pm 7.7$             | $37 \pm 7.8$         |
| Total proteins (g/dL)                        | 6.5-8.2      | $7.2 \pm 0.7$              | $7.3 \pm 0.5$        |
| Albumin (g/dL)                               | 3.6-5.2      | $4.0 \pm 0.5$              | $4.3 \pm 0.3$        |
| Total cholesterol (mg/dL)                    | 120-200      | 190 ± 43                   | 190 ± 39             |
| Triglycerides (mg/dL)                        | 50-200       | 159 ± 88                   | 125 ± 76             |
| High-density lipoprotein cholesterol (mg/dL) | >35          | 46 ± 11                    | 44 ± 11              |
| Low-density lipoprotein cholesterol (mg/dL)  | <180         | 120 ± 51                   | 106 ± 33             |
| CRP (mg/dL)                                  | 0.0-0.5      | $0.5 \pm 0.7$              | $0.3 \pm 0.1$        |
| Homocysteine (µmol/L)                        | 5-16         | $32.4 \pm 14.6$            | 5.7 ± 2*             |

NOTE. Values expressed as mean  $\pm$  SD or percent. To convert GFR from mL/min to mL/s, multiply by 0.01667; urea from mg/dL to mmol/L, multiply by 0.357; glucose from mg/dL to mmol/L, multiply by 0.05551; iPTH from pg/mL to mg/mL, multiply by 1; calcium from mg/dL to mmol/L, multiply by 0.2495; phosphorus from mg/dL to mmol/L, multiply by 0.3229; total protein from g/dL to g/L, multiply by 10; cholesterol, high-density lipoprotein, and low-density lipoprotein cholesterol from mg/dL to mmol/L, multiply by 0.02586, triglycerides from mg/dL to mmol/L, multiply by 0.01129.

†See text for other normal values.

<sup>\*</sup>P < 0.05 versus patients with CRF.

#### 1. Causes of renal failure

chronic glomerulonephritis (46%), chronic interstitial nephritis (10%), postischemic renal failure, adult polycystic kidney disease, malformations of urinary apparatus (10%), unknown(34%).

#### 2.K/DOQI guidelines,

8 patients :stage 2 (9.4%) (GFR, 60 to 89 mL/min);

40 patients :stage 3 (47%) (GFR, 30 to 59 mL/min);

24 patients: stage 4 (28.2%) (GFR, 15 to 29 mL/min;

13 patients:stage 5(15.3%) (GFR 15 mL/min).

■ 1. Spiral CT showed in 34 CAC / 85 patients with CRF (40%).

2.Calcium scores

<100 :12 patients,

101 to 400 :12 patients,

401 to 1,000 : 6 patients,

> 1,000 : 4 patients.

Table 2. Clinical and Biochemical Characteristics of Patients With and Without CAC

|   | Patients With CAC<br>(n = 34; 40%) | Patients Without CAC<br>(n = 51) |
|---|------------------------------------|----------------------------------|
| Sex (% men)   | 91                                 | 87                               |
| Age (y)   | $59.2 \pm 8.2$                     | 46.4 ± 12.2*                     |
| Body mass index (kg/m²)   | 26.8 ± 3.6                         | 26.5 ± 4.0                       |
| Smokers (%)   | 22                                 | 24                               |
| Duration of CRF (y)   | $6.5 \pm 5.2$                      | $6.0 \pm 5.9$                    |
| Systolic blood pressure (mm Hg)   | 134 ± 11                           | 132 ± 13                         |
| Diastolic blood pressure (mm Hg)  | 81 ± 4                             | $79 \pm 13$                      |
| GFR (mL/min)  | 33 ± 16                            | $33 \pm 17$                      |
| iPTH (pg/mL)  | 122 ± 110                          | 153 ± 128                        |
| Calcium (mg/dL)   | $9.3 \pm 0.5$                      | $9.4 \pm 0.5$                    |
| Phosphorus (mg/dL)  | $3.9 \pm 0.9$                      | 4.2 ± 0.8                        |
| Calcium-phosphorus product (mg²/dL²)                                      | $36.9 \pm 8.3$                     | $39.8 \pm 7.4$                   |
| Total protein (g/dL)  | $7.2 \pm 0.7$                      | $7.1 \pm 0.7$                    |
| Serum albumin (g/dL)  | $4.0 \pm 0.5$                      | $4.1 \pm 0.5$                    |
| Total cholesterol (mg/dL)   | 189 ± 43                           | 194 ± 45                         |
| Triglycerides (mg/dL)   | 180 ± 115                          | 144 ± 60                         |
| High-density lipoprotein cholesterol (mg/dL)                              | 46 ± 8                             | 46 ± 13                          |
| Low-density lipoprotein cholesterol (mg/dL)                               | 111 ± 39                           | 125 ± 44                         |
| CRP (mg/dL)   | $0.7 \pm 1.2$                      | $0.4 \pm 0.2$                    |
| Homocysteine (μmol/L)   | 36.7 ± 16                          | 32.2 ± 18                        |
| Phosphate P binders (%)   | 27                                 | 27                               |
| Angiotensin-converting inhibitors/angiotensin II receptor antagonists (%) | 51                                 | 50                               |
| Calcium channel blockers (%)  | 40                                 | 38                               |
| Statins   | 22                                 | 19                               |
| Vitamin D analogues   | 14                                 | 15                               |

NOTE. Values are expressed as mean  $\pm$  SD or percent. To convert GFR in mL/min to mL/s, multiply by 0.01667; urea in mg/dL to mmol/L, multiply by 0.357; glucose in mg/dL to mmol/L, multiply by 0.05551; iPTH in pg/mL to ng/L, multiply by 1; calcium in mg/dL to mmol/L, multiply by 0.2495; phosphorus in mg/dL to mmol/L, multiply by 0.3229; total protein in g/dL to g/L, multiply by 10; cholesterol in mg/dL to mmol/L, multiply by 0.02586; triglycerides in mg/dL to mmol/L, multiply by 0.01129.

<sup>\*</sup>P < 0.05 versus patients with CRF.

■ 1. > 50 y/o (85% of cases, CAC)  $_{\prime}$  > 1 coronary artery

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4-vessel: 21%;
3-vessel: 48%;
2-vessel: 15%;
1-vessel disease:15%
The left anterior descending artery (76%);
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2. the calcification process affected

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the right coronary artery (70%) left main coronary artery (67%) circumflex artery (64%).
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3.Mitral valve calcification : 2 patients.

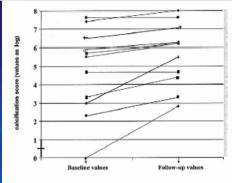


Fig 2. Calcification score in patients with CRF (n = 18) who underwent a repeated scan. Patients (n = 7) who remained free of calcification are not included.

1. Spiral CT was repeated in 18 patients (15 men,3 women; 8 patients without

CAC, 10 patients with CAC at

baseline)

mean follow-up: 7.9 months (range, 7 to 9 months).

2. free of CAC: 7 patients
 ex novo appearance of CAC: 1 patient
 unchanged calcium score :2 patients
 significant increase in calcification score: 8 patients

- 1. Coronary artery disease.
  - the effects of some antihypertensive drugs (ie, calcium channel blockers versus diuretics) on plaque progression.
- 2.CAC is present in 40% of patients (13% of controls).
- 3.Calcium scores are higher than 100 in 65% of patients; 1 control subject had a calcium score higher than 100.

Table 3. Multivariable Regression With CAC as Dependent Variable

|                  | P     | Odds Ratio (95%<br>Confidence Interval) |
|------------------|-------|---|
| Age              | 0.000 | 1.137 (1.137-1.218)                     |
| Duration of CRF  | 0.721 | 1.001 (0.994-1.009)                     |
| GFR              | 0.698 | 1.010 (0.960-1.063)                     |
| iPTH             | 0.268 | .997 (0.991-1.003)                      |
| Serum calcium    | 0.476 | 1.573 (0.451-5.471)                     |
| Serum phosphorus | 0.800 | .896 (0.383-2.093)                      |

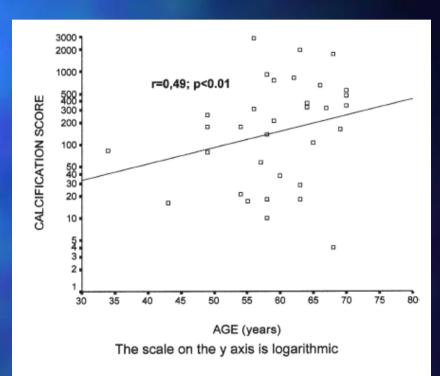


Fig 1. Correlation between calcification score and age in patients with CRF with CAC (n = 34).

1. Amplifying effect of dialysis on calcifications?

\*very low prevalence of cardiac valve calcification in our patient, whereas calcification of the mitral and aortic valves is present largely in patients on dialysis therapy

#### 2.variables:

greater derangement of mineral metabolism, greater iPTH production, excessive calcium load by dialysate, vitamin D intoxication, enhanced inflammation.

1.Chronic inflammation is: CRP and homocysteine
 CRP, homocysteine level NOT associated with calcification score.

2.in many patients: >1 coronary vessel , electron-beam CT , 2.2x cardiovascular

progression is fast:14%: 26 weeks

25%:52 weeks

(on dialysis therapy)

3. In asymptomatic patients with no history of coronary artery stenosis and renal failure?

■ 1. CRF -> CAC?

2. Evaluate asymptomatic patients not yet on dialysis therapy?