# Vitamin D (25(OH)D and 1,25(OH)<sub>2</sub>D<sub>3</sub>) levels and abdominal aortic calcification in patients with chronic kidney disease



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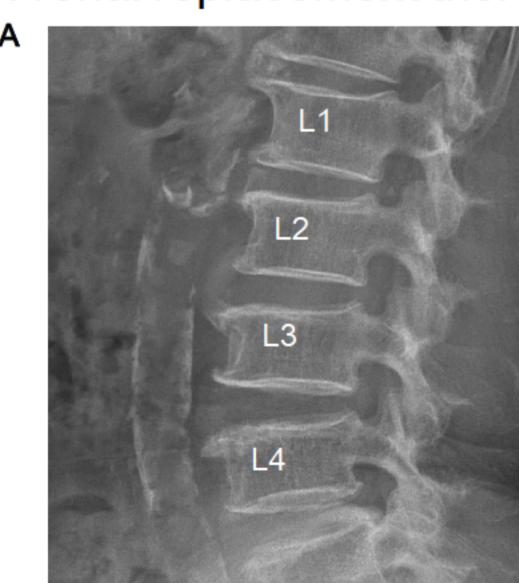


## Background

- Vascular calcification is an independent risk factor for cardiovascular disease, which is the leading cause of death in patients with chronic kidney disease (CKD). Low vitamin D levels might have affected vascular calcification. CKD patients revealed a high prevalence of vitamin D deficiency.
- The aim of this study was to investigate the abdominal aortic calcification according to vitamin D levels in patients with advanced kidney disease.

#### Method

- Study design: retrospective
- Study period: Jan. 2011 ~ Aug. 2012
- Inclusion
  - 1. Patients of CKD stage III-V in single center (N=149)
  - 2. age ≥ 20
- Exclusion
- 1. acute infectious disease
- 2. liver disease
- 3. renal replacement therapy (dialysis, renal transplantation)



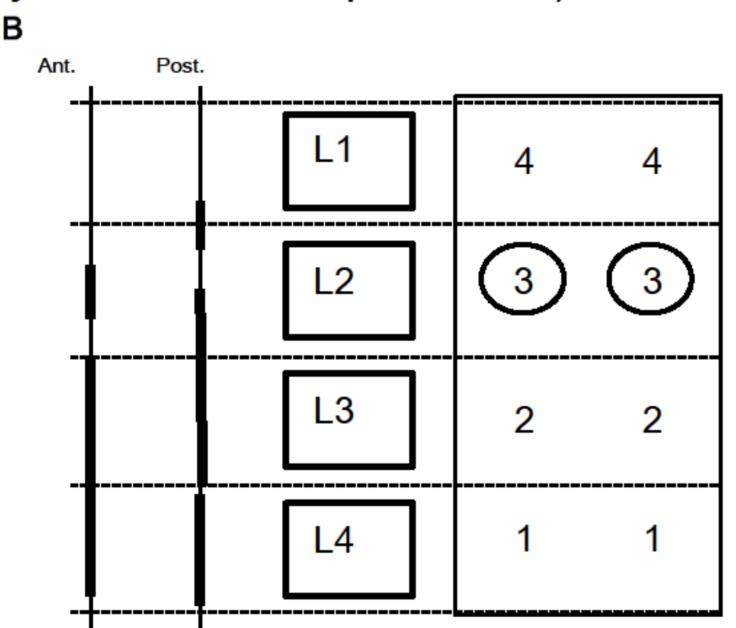


Figure. 1 A) Abdominal aortic calcification (AAC) was shown on plain radiography of lateral lumbar spine view. B) Grading of calcification was assessed at the ant. (anterior) and the post. (posterior) walls of the abdominal aorta adjacent to vertebrae L1-L4 and the composite score determined. AAC score of this patients were 6.

### Results

Table 1. Demographic of total subjects (N=149)

| Variables                       | Mean±SD or n (n of patients = 149) |
|---------------------------------|------------------------------------|
| Age, years                      | 64.5±13.0 (20-88)                  |
| Male : Female (%)               | 90 (60.4) : 59 (39.6)              |
| eGFR, mL/min/1.73m <sup>2</sup> | 12.5±9.0                           |
| CKD stage (%)                   |                                    |
| III, IV                         | 40 (26.8)                          |
| V                               | 109 (73.2)                         |
| Cause of CKD (%)                |                                    |
| DM                              | 95 (63.8)                          |
| HTN                             | 18 (12.1)                          |
| CGN                             | 24 (16.1)                          |
| others                          | 12 (8.1)                           |
| AACS, 0-8 (%)                   |                                    |
| AACS, 0-3                       | 117 (78.5)                         |
| AACS, 4-8                       | 32 (21.5)                          |

Table 2. Laboratory parameters of patients with advanced kidney disease

|   | <u> </u>                      |
|---|-------------------------------|
| Variables                                     | Mean±SD (n of patients = 149) |
| Hemoglobin, g/dL                              | 8.8±1.8 (4.1-13.7)            |
| Albumin, mg/dL                                | 3.3±0.5 (1.8-4.7)             |
| BUN, mg/dL                                    | 73.4±35.0 (19.5-194.5)        |
| Creatinine, mg/dL                             | 6.6±4.2 (1.5-25.6)            |
| Alkaline phosphatase                          | 248.6±124.9 (83-904)          |
| Calcium, mg/dL                                | 7.8±0.9 (3.7-10.0)            |
| Phosphate, mg/dL                              | 5.3±2.1 (1.8-15.1)            |
| Calcium X Phosphate, mg²/dL²                  | 41.1±14.1 (12.3-95.1)         |
| Intact PTH, pg/mL                             | 229.1±182.9 (5.3-973.9)       |
| 25(OH)D, ng/mL                                | 8.7±5.5 (4.0-52.0)            |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> , pg/mL  | 9.6±7.8 (1.6-56.2)            |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> /25(OH)D | 1.28±1.09 (0.06-7.40)         |
| eGFR, mL/min/1.73m <sup>2</sup>               | 12.58±9.0 (2.0-48.9)          |

Table 3. Comparison of clinical and laboratory parameters of patients stratified according to advanced kidney disease

|   | CKD stage III, IV | CKD stage V   | P value |
|---|-------------------|---------------|---------|
| Patients (%)                                  | 40 (26.8)         | 109 (73.2)    |         |
| Age, years                                    | 68.0±11.4         | 63.2±13.4     | 0.047*  |
| Male (%)                                      | 22 (55.0)         | 68 (62.4)     | 0.414   |
| Diabetes (%)                                  | 24 (60.0)         | 71 (65.1)     | 0.563   |
| WBC, 10 <sup>3</sup> /µL                      | 8890.0±4853.9     | 8247.7±3950.9 | 0.456   |
| Hemoglobin, g/dL                              | 9.6±1.8           | 8.6±1.7       | 0.001*  |
| Albumin, mg/dL                                | 3.2±0.5           | 3.3±0.4       | 0.105   |
| BUN, mg/dL                                    | 47.7±21.3         | 82.9±34.4     | <0.001* |
| Creatinine, mg/dL                             | 2.6±0.6           | 8.0±4.1       | <0.001* |
| Alkaline Phosphatase                          | 256.1±135.8       | 245.8±121.3   | 0.658   |
| Calcium, mg/dL                                | 8.1±0.7           | 7.7±1.0       | 0.012*  |
| Phosphate, mg/dL                              | 4.0±0.9           | 5.8±2.2       | <0.001* |
| Calcium X Phosphate, mg²/dL²                  | 33.2±8.6          | 44.1±14.6     | <0.001* |
| Cystatin-C, mg/L                              | 3.1±1.7           | 4.4±1.3       | <0.001* |
| CRP, mg/L                                     | 29.5±49.4         | 29.0±47.4     | 0.960   |
| Intact PTH, pg/mL                             | 118.6±92.3        | 270.4±191.3   | <0.001* |
| 25(OH)D, ng/mL                                | 8.5±3.6           | 8.8±6.0       | 0.717   |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> , pg/mL  | 11.3±8.4          | 8.9±7.5       | 0.097   |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> /25(OH)D | 1.46±1.12         | 1.22±1.07     | 0.233   |
| AACS, 0-8                                     | 2.1±2.1           | 1.9±2.0       | 0.616   |
| eGFR, mL/min/1.73m <sup>2</sup>               | 24.8±8.3          | 8.0±3.3       | <0.001* |

Table 4. Comparison of demographics and clinical laboratory parameters of advanced kdiney disease patients stratified according to abdominal aortic calcification score (0-8)

|  | AACS≤3        | AACS>3        | P value  |
|--|---------------|---------------|----------|
| Patients (%)                                   | 117 (78.5)    | 32 (21.5)     |          |
| Age, years                                     | 62.9±13.1     | 70.1±11.0     | 0.005*   |
| Male (%)                                       | 71 (60.7)     | 19 (65.6)     | 0.893    |
| Diabetes (%)                                   | 74 (63.2)     | 21 (65.6)     | 0.804    |
| CKD stage V (%)                                | 86 (73.5)     | 23 (71.9)     | 0.854    |
| eGFR, mL/min/1.73m <sup>2</sup>                | 12.6±9.2      | 12.3±8.5      | 0.870    |
| WBC, 10 <sup>3</sup> /µL                       | 8099.1±4091.7 | 9593.7±4467.0 | 0.075    |
| Hemoglobin, g/dL                               | 8.8±1.7       | 8.9±1.8       | 0.748    |
| Albumin, mg/dL                                 | 3.3±0.5       | 3.2±0.5       | 0.609    |
| BUN, mg/dL                                     | 72.7±34.9     | 76.2±35.8     | 0.619    |
| Creatinine, mg/dL                              | 6.6±4.3       | 6.4±4.0       | 0.811    |
| Alkaline Phosphatase                           | 245.2±120.8   | 260.8±140.5   | 0.534    |
| Calcium, mg/dL                                 | 7.8±1.0       | 7.8±0.7       | 0.791    |
| Phosphate, mg/dL                               | 5.3±2.0       | 5.5±2.4       | 0.620    |
| Calcium X Phosphate                            | 40.8±13.7     | 42.5±15.5     | 0.551    |
| Cystatin-C, mg/L                               | 4.0±1.6       | 4.1±1.1       | 0.638    |
| CRP, mg/L                                      | 27.2±44.8     | 36.3±57.8     | 0.348    |
| Intact PTH, pg/mL                              | 234.2±178.3   | 2107±200.4    | 0.522    |
| 25(OH)D, ng/mL                                 | 8.8±6.0       | 8.5±2.9       | 0.757    |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> , pg/mL   | 10.3±8.4      | 6.9±4.1       | 0.034*   |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> ,/25(OH)D | 1.39±1.15     | 0.87±0.48     | < 0.001* |
| AACS   | 1.06±1.00     | 5.47±1.27     | < 0.001* |

Table 5. Pearson's Correlation Coefficient

|             | Cr       | BUN      | Ca       | Р       | iPTH   | 25(OH)D  | 1,25(OH)2D3 |
|-------------|----------|----------|----------|---------|--------|----------|-------------|
| BUN         | 0.701**  |          |          |         |        |          |             |
| Ca          | -0.418** | -0.323** |          |         |        |          |             |
| Р           | 0.742**  | 0.633**  | -0.382** |         |        |          |             |
| iPTH        | 0.541**  | 0.465**  | -0.360** | 0.519** |        |          |             |
| 25(OH)D     | 0.036    | 0.056    | -0.005   | -0.010  | -0.062 |          |             |
| 1,25(OH)2D3 | -0.235** | -0.171*  | 0.085    | -0.194* | -0.052 | 0.045    |             |
| VitD ratio  | -0.195*  | -0.166*  | 0.006    | -0.096  | -0.010 | -0.283** | 0.849       |

Table 5. Multivariate analysis of risk factors for abdominal aortic calcification score > 3

|                                      | Odd Ratio | 95% CI      | P value |
|--------------------------------------|-----------|-------------|---------|
| Male                                 | 0.782     | 0.283-2.161 | 0.636   |
| Age                                  | 1.075     | 1.026-1.126 | 0.002*  |
| DM                                   | 0.925     | 0.352-2.432 | 0.874   |
| CKD stage V                          | 1.338     | 0.405-4.417 | 0.633   |
| Calcium                              | 0.820     | 0.469-1.432 | 0.485   |
| Phosphate                            | 1.205     | 0.869-1.671 | 0.263   |
| Cystatin-C                           | 1.149     | 0.827-1.596 | 0.407   |
| Intact PTH                           | 0.999     | 0.996-1.002 | 0.606   |
| 25(OH)D                              | 0.988     | 0.915-1.067 | 0.758   |
| 1,25(OH) <sub>2</sub> D <sub>3</sub> | 0.888     | 0.796-0.991 | 0.033*  |

# Conclusions

These results suggest that vitamin D deficiency was common, and age and serum 1,25(OH)<sub>2</sub>D<sub>3</sub> level were independent risk factors for vascular calcification in patients with advanced kidney disease.





