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Effect of combined administration of vitamin D<sub>3</sub> and vitamin K<sub>2</sub> on bone mineral density of the lumbar spine in postmenopausal women with osteoporosis

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## **Abstract**

The effect of the combined administration of vitamin D<sub>2</sub> and vitamin K<sub>2</sub> on bone mineral density (BMD) of the lumbar spine was examined in postmenopausal women with osteoporosis. Ninety-two osteoporotic women who were more than 5 years after the state of the smenopause, aged 55–81 years, were randomly divided into four administration groups: vitamin  $D_3$  (1 $\alpha$  hydroxyvitamin  $D_3$ ) o.75µg/day) (D group; n 5 29), vitamin K2 (menatetrenone, 45mg/day) (K group; n 5 22), vitamin D3 plus vitamin K2 (DK group, n 5 21), and calcium (calcium lactate, 2g/day) (C group; n 5 20). BMD of the lumbar spine (L2–L4) was measured by dual energy X-ray  $absorptiometry\ at\ o,\ a,\ and\ a\ years\ after\ the\ treatment\ started.\ There\ were\ no\ significant\ differences\ in\ age,\ body\ mass\ index,\ years\ after\ the\ treatment\ started.$ since menopause, and initial BMD among the four groups. One-way analysis of variance (ANOVA) with repeated measurements showed a significant decrease in BMD in the C group (P, 0.001). Two-way ANOVA with repeated measurements showed a significant increase in BMD in the D and K groups compared with that in the C group (P. 0.05 and P. 0.001, respectively), and a significant increase in BMD in the DK group compared with that in the C, D, and K groups (P, 0.0001, P, 0.05 and P, 0.01, respectively). These  $findings\ indicate\ that\ combined\ administration\ of\ vitamin\ \textbf{$K_2$}, compared\ with\ calcium\ administration\ , appears\ to\ administration\ , appe$ be useful in increasing the BMD of the lumbar spine in postmenopausal women with osteoporosis.

This paper is only available as a PDF.

## Key words

Vitamin D<sub>3</sub> · Vitamin K<sub>2</sub> · Postmenopausal women · Osteoporosis · Bone mineral density (BMD)

ORTHOPAEDIC SCIENCE

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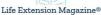
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#### Vitamin K2 Reduces Atherosclerosis

Vitamin K2 blocks the progression of arterial thickening and st Studies show higher vitamin K2 intake reduces the risk of dying from heart disease by 57%

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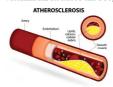
In a new study, scientists have presented powerful evidence that **vitamin K2** can reduce the progression of **atherosclerosis**, the "blockage" of the arteries that can lead to heart attacks and strokes.

This is tremendous news for the millions of Americans who are at risk for cardiovascular disease, which remains the number one killer in the US.1 Scientists long ago learned that vitamin K2 plays a crucial role in activating proteins that help keep calcium where it

belongs, in the bones, and out of blood vessels where it can cause proble Published data show that people with higher intake of vitamin K2 have a 57% reduction in the risk of dying from

cardiovascular disease, and as much as an 81% reduction in non-vertebral fractures. Here, we'll review the important new study on atherosclerosis, and then examine previous findings that explain how

#### vitamin K2 contributes so importantly to both cardiovascular and bone health in aging adults. Vitamin K2 Reduces the Progression of Arterial Thickening



Numerous studies have demonstrated that people with higher intakes of vitamin K2 have a reduced risk for cardiovascular disease. \*\* Intrigued by this connection, Polish researchers from the Medical University at Lodz teamed up with researchers from Maastricht University in the Netherlands and Poland's International Science and Health Foundation to determine if vitamin K2 supplementation could  $\it reduce the progression$  of existing rosclerosis\*

The scientists evaluated the progression of atherosclerosis in a group of 42 patients with chronic kidney disease.6 These patients were ideal for this type

of study because they are known to experience a rapid reduction in both of memory and the study because they are known to experience a rapid reduction in both of memory and the memory and the study of the study are also subject to equally excessive deposits of calcium in tissues where it doesn't belong—particularly in the walls of major arteries.\*

udy, the subjects were divided into two groups. One group re eived vitamin K2 (90 mcg per day) plus vitamin D3 (400 IU per day). The second group received only vitamin D3 (400 IU per day).6

After nine months, it was already evident that the subjects taking the combination of vitamins K2 and D3 experienced a slower progression of the Common Carotid Intima Media Thickness, which is a good indicator of atherosclerosis well as a predictor of cardiovascular episodes and death. Specifically, the thickness of the carotid (major neck) as increased by 13.73% in the group taking vitamin D3, but in the group taking both vitamins, it only increased by 6.32%. Remember that the group of subjects in this study have a tendency for an increased carotid intima media thickness as a result of calcium losses from bone.

In addition, subjects taking the combination of vitamins K2 and D3 showed a reduction in carotid artery calcification score in all patients except those with the highest scores at baseline. Finis indicates that calcium was staying in the bones, where it belongs, and out of the arteries.

These results clearly indicated that vitamin K2 does indeed reduce the progression of atherosclerosis.

The impact of vitamin K2 on atherosclerosis is due to its role in activating a group of proteins whose job it is to keep calcium in the bones and out of the arteries.32

In bones, vitamin K2 activates a specific protein called **osteocalcin**. When osteocalcin is activated by vitamin K2, it binds calcium tightly to bone minerals to create strong bones. In arteries, vitamin K2 activates a protein called **matrix Gla protein**. When matrix Gla protein is activated by vitamin K2, it prevents calcium from being deposited in arteries. 9,10

When vitamin K2 is not present in sufficient enough quantities to activate these two proteins, the result is an increased rosis and atheroscler sis because of calcium loss from the bones

#### High Praise

The findings from this recent Polish study were significant enough to merit published comments by other experts in the

The experts observed that patients in the combined vitamins K2 plus D3 arm of the study already had more severe kidney disease, despite being randomly assigned to the treatment groups. This means that the group taking vitamin K2 along with D3 succeeded despite a less favorable starting point compared with the group taking only vitamin D3.14

Of course, while chronic kidney disease is common and deadly, patients with the disease are far from the only ones a risk for cardiovascular disease and bone loss arising from insufficient vitamin K2 intake. But given their extreme risk those conditions, this patient population made an ideal test group to demonstrate the value of supplementation with both vitamin K2 and D3.

### Vitamin K2's Dual Benefits



The recent Polish study further validated previous studies demonstrating vitamin K2's heart- and bone-health benefits.

Numerous studies have shown that populations with higher daily vitamin K2 intake (more than 32 mcg) have a 50% reduction in the risk of death from cardiov 32 mog) have a 50% reduction in the risk of death from cardiovascular usease compared those with low intake. <sup>32</sup> Calculated another way, for every 10-mog per day-increase in vitamin K6 intake, the risk of coronary heart disease falls by 9%. <sup>53</sup> In another study, women with the highest

intake of vitamin K2 were found to be at a 20% lower risk for coronary artery calcification compared with women who

These benefits have been found to extend to bone health as well. Postmenopausal women taking 1,500 mg of calcium along with 45 mg of vitamin K2 daily experienced an increase in bone mineral density and a 55,9% reduction in *inactive* osteocalcin levels,<sup>15</sup> a marker reported to be elevated in osteoporotic patients and related to an increased hip fracture risk.<sup>18,17</sup> Another study showed that when **180 mcg** of vitamin K2 daily was given for 3 years, it increased the amount of activated osteocalcin, produced significant improvements in bone mineral content and density, and increased bone strength.1

#### Western Diet is Deficient in Vitamin K2

The average American diet lacks enough vitamin K2—which is found primarily in organ meats, egg yolks, cheese, and in a Japanese staple dish of fermented soybean called natto-to properly activate those vitamin K-depe Compounding the problem, vitamin K2 deficiency worsens with advancing age.10

Fortunately, studies have shown that sustained vitamin K2 supplementation produces significant improvements in bone ion and strength and also in standard meas ents of cardiovascular health.92

The take-home message is that supplementing with vitamin K2 is practically mandatory for the average American desiring both healthy bones and a healthy cardiovascular system.

Vitamin~K2~has~the~unique~ability~to~activate~proteins~that~send~calcium~to~its~proper~destination, into~bone, while simultaneously preventing~calcium~from~being~deposited~in~blood~vessel~walls.

A new study has shown that supplemental vitamin K2 can reduce the progression of arterial thickening and stiffening in a population of kidney disease patients, a group that notoriously has cardiovascular complications arising from excessive calcium deposition.

Thus, supplementation with vitamin K2 should be part of a daily longevity program for any aging adult

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