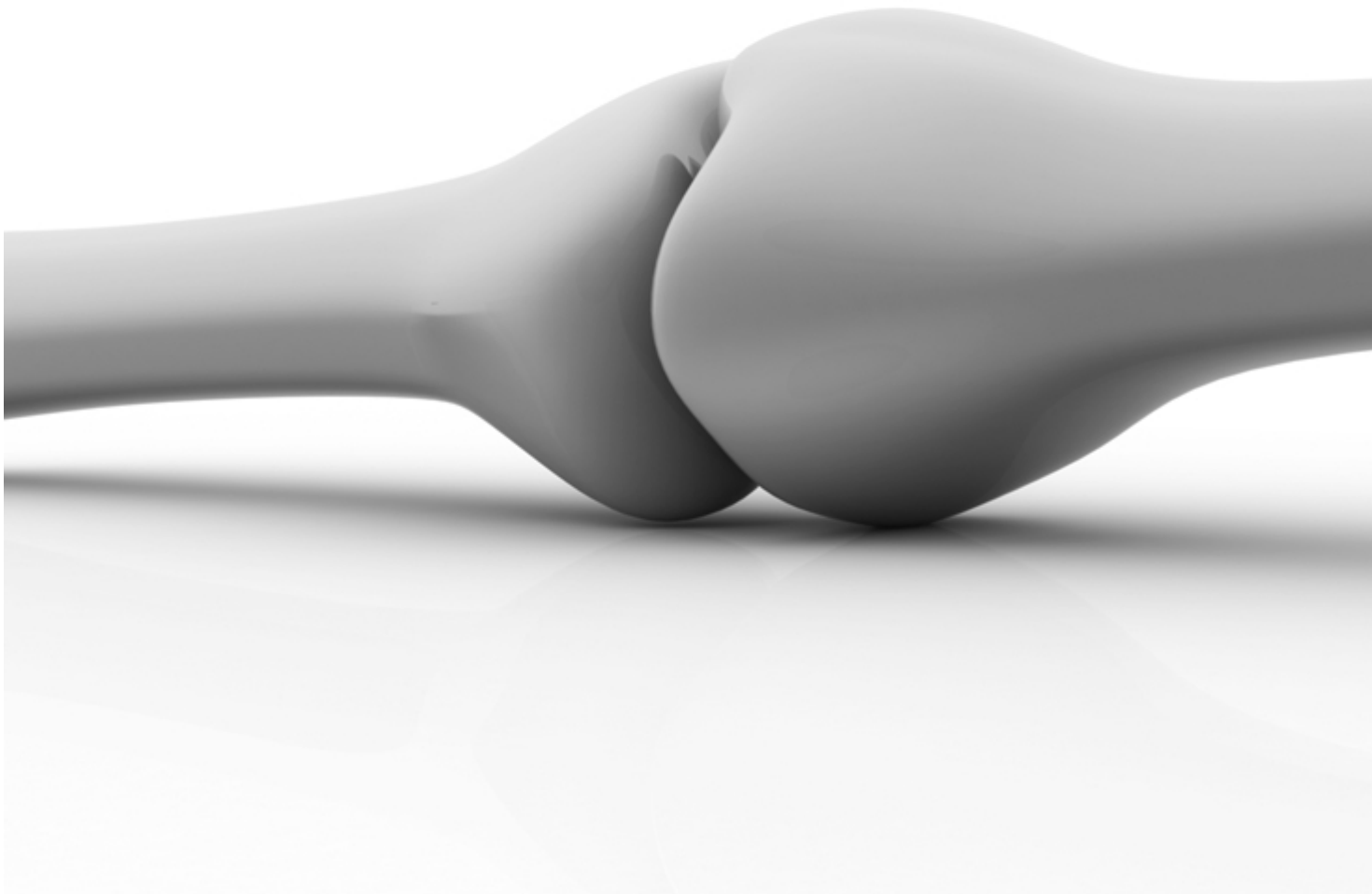


## Bone Health <sup>[1]</sup>

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## The Importance of Healthy Bones

Bones play many important roles in your body. They support you, help you move, and protect some of your organs from injury. Bones also store minerals such as calcium and phosphorus and make many of the cells that circulate in your blood (e.g., red blood cells, white blood cells, platelets).

Bones are constantly changing throughout your life. Every day, your body removes old bone and adds new bone in its place. In young people, more bone is added than removed. After age 30, more bone is removed than added. As a result, the bones become lighter and weaker, which puts them at greater risk for injury.

Many people have weak bones and do not know it. This is because bone loss often happens over a long period of time and does not hurt. For many people, a broken bone or a fracture is the first sign of weakened bones. This weakness is often referred to as low bone mass or low bone density. Bone mass or bone density refers to how much internal bone structure is in a given section of bone. For example, more dense bone has more calcium and phosphorus ? two key minerals that make up bone ? per square inch of bone.

## Bone Loss and HIV

Bone loss and weakened bones occur more often in people living with HIV (HIV+). Experts are not exactly sure why. It could be due to HIV itself, HIV drugs, or people living with HIV getting older. People living with HIV are more likely to experience three bone conditions: osteopenia, osteoporosis, and osteonecrosis.

### Osteopenia

Osteopenia is caused by a loss of bone minerals that leads to lower-than-normal bone density. Most often, osteopenia has no symptoms. The only way to know if you have this condition is to have a bone density test. If you do learn you have osteopenia, there are things you can do to stop it and possibly even reverse it (see the Physical Activity and Diet sections below).

Osteopenia can lead to a more serious condition called osteoporosis (see below), but having osteopenia does not mean you will definitely develop osteoporosis; however, it does mean that you are more likely to develop it. Although bone loss with osteopenia is generally less severe than with osteoporosis, it does indicate that bones are weaker and may be more likely to break.

## **Osteoporosis**

Osteoporosis is a more serious condition than osteopenia and refers to a loss of bone density and bone mass. Over time, breaking down more bone than is replaced causes lower bone mineral density, also called bone mineral density, which means there are lower-than-normal levels of minerals in the bones. Bones become weak and are more likely to break. People with osteoporosis most often break bones in the wrist, spine, and hip. Sometimes bones break without an injury.

## **Osteonecrosis (avascular necrosis)**

Osteonecrosis means bone death. It is caused by a loss of blood supply to the bone. Osteonecrosis may cause pain at the affected joint. It usually affects the head of the femur, the ball-shaped part of the thigh bone that connects it to the hip.

# **How to Know if You Have Bone Problems**

## **Osteoporosis and Osteopenia**

Bone mineral density (BMD) tests are the only way to find out if you have osteoporosis or osteopenia. The most widely used BMD test is a DEXA (Dual Energy X-ray Absorptiometry) scan. The DEXA scan is a kind of x-ray, and is an easy and painless test that takes about 15 minutes.

## **Osteonecrosis**

Osteonecrosis can cause pain in the joints, usually in the hip area. At first the pain may only occur when you put weight on the joint. In more severe cases the pain can be constant. An MRI (magnetic resonance imaging) scan can spot early stages. X-rays and other scans can detect advanced osteonecrosis.

## **Guidelines for Bone Care**

The World Health Organization (WHO) recently released new guidelines for the management of bone health among people living with HIV. The guidelines recommend that all premenopausal women living with HIV who are 40 years old or older have their risk of fracture measured using the Fracture Risk Assessment Tool (FRAX) without a DEXA scan. This assessment looks at risk factors for fracture (e.g., age, smoking, family history of fracture, etc.) and calculates a probability of future fracture.

The guidelines also suggest that DEXA scans be conducted for all postmenopausal women living with HIV, people living with HIV who have a history of fragility fracture, people living with HIV who are receiving chronic glucocorticoid treatment, and people living with HIV who have a

high risk of falls.

Lastly, the guidelines recommend that people living with HIV who are at risk for bone loss and/or fracture have their HIV treatment regimens adjusted so that they do not contain tenofovir disoproxil fumarate (TDF; see Tenofovir section below).

## Factors Affecting Your Bone Health

There are several things that can affect your bone health and put you at risk for bone loss. Some of these are things you cannot control:

- Age: Your chances of getting osteoporosis increase with age
- Sex: Women have a greater chance of getting osteoporosis than men
- Menopause [2]: Women lose bone due to hormone changes that happen after menopause. While hormone replacement therapy is no longer routinely recommended, it is an effective therapy for bone loss that does not respond to other treatments.
- Family history: Having a parent with osteoporosis, especially a mother or father who has had a hip fracture, increases your risk of bone loss

The good news is that there are also several things that you can control ? things that you can do to maximize your bone health, regardless of your age, sex, or family history.

### Diet and Body Weight

Even though you cannot control all of the things that lead to bone disease, you can control your diet [3]. The mineral [4] calcium makes up a large part of your bones. This means that if you do not get enough calcium in your diet, your bones may get weaker. Having too little body weight ? being too thin ? can also increase your chances of having weaker bones.

Calcium is found naturally in some foods, and it is added to others. Some foods that contain calcium are:

- Dairy products, including milk, yogurt, and cheese
- Soy products: tofu, tempeh, soy milk
- Seeds: sesame seeds, chia seeds, and flax seeds
- Dark leafy greens: spinach, kale, chard, collard greens, turnip greens
- Broccoli
- Beans: black beans, navy beans
- Dried fruits and nuts: figs, almonds, brazil nuts
- Oranges and orange juice (especially calcium-fortified orange juice)
- Salmon or sardines with the bones

Some people living with HIV still need to take calcium pills every day even if their diet includes foods with calcium in them. A registered dietitian or other trained health care provider can help you decide if you should take calcium supplements. If you do take calcium supplements, it may be a good idea to take calcium pills with vitamin D in them, since your body cannot use calcium without vitamin D. According to the Office of Dietary Supplements at the National Institutes of Health in the US, the recommended dietary allowances are:

- Girls and young women nine to 18 years old need 1,300 mg of calcium and 600 IU of vitamin D daily

- Women 19 to 50 need 1,000 mg of calcium and 600 IU of vitamin D daily
- Women 50 to 70 need 1,200 mg of calcium and 600 IU of vitamin D daily
- Women 71 and over need 1,200 mg of calcium and 800 IU of vitamin D daily
- Pregnant and breastfeeding teens need 1,300 mg of calcium and 600 IU of vitamin D daily
- Pregnant and breastfeeding adults need 1,000 mg of calcium and 600 IU of vitamin D daily

It is important that you talk to your health care provider before taking any supplements [4] and not take more than these amounts unless instructed to by your health care provider.

## Physical Activity

If you do not have joint pain, it is important to be physically active [5] on a regular basis. When you exercise your muscles pull against your bones, which helps keep your bones healthy and strong. The best kind of physical activities to keep your bones strong are activities that use weight or resistance such as:

- Walking (you can use ankle weights)
- Working out with weights, weight machines, or bands (also called resistance training)
- Stair climbing
- Hiking
- Aerobics
- Jogging

If you cannot do high-impact weight-bearing activities, try lower-impact ones. For example, try walking or stair climbing instead of jogging. If you have not exercised regularly for a while, check with your health care provider before beginning a new exercise program.

Once you have your health care provider's approval, start your exercise routine slowly. Every week or two, make your routine five minutes longer. In the end, it is recommended that you do 150 minutes of moderate-level physical activity each week (e.g., five 30-minute workouts).

## Tobacco, Alcohol, and Caffeine

Using tobacco products, whether it's smoking, using snuff, or chewing tobacco, can increase your chances of bone loss. If you use tobacco, it is a good idea to try and quit. Stopping smoking is one of the single best things you can do for your health. Talk with your health care provider about stopping smoking [6] ? there are many tools to help you quit. You can also find lots of information and support online (<http://www.smokefree.gov> [7]).

Drinking a lot of alcohol [8] and/or caffeine can also put you at greater risk of bone loss.

## Using Certain Medicines

Long-term use of some medicines can increase the chances of bone loss and lead to weakened bones:

- Depo-Provera (an injectable form of birth control [9])
- Glucocorticosteroids (drugs such as prednisone and cortisone)
- Thyroid hormones
- Anticonvulsants (anti-seizure medications)

- Heparin (blood thinner)
- Pentamidine (antibiotic)
- Ketoconazole (anti-fungal)
- Tenofovir disoproxil fumarate (TDF or Viread; see below)

## Tenofovir: TDF and TAF

Tenofovir is an HIV drug that comes in two forms: TDF (tenofovir disoproxil fumarate) and TAF (tenofovir alafenamide). TDF is the tenofovir also known by the brand name Viread and found in other combination drugs [10] such as Truvada. Previous studies have shown that treatment with TDF can lead to a reduction in kidney function and bone loss [1]. Treatment with TAF, which can be given at lower doses than TDF while achieving the same level of HIV suppression, appears to produce less bone loss and less reduction in kidney function.

A study of over 12,000 Europeans living with HIV found that treatment with TDF increases not just bone loss, but the risk of actual bone fracture in adults. This is an especially important consideration for women, who often experience bone loss naturally after menopause [2]. The risk of fracture did not increase with increasing time on TDF, which matches our understanding that most of the loss of bone loss associated with taking tenofovir disoproxil occurs early in treatment.

Researchers at CROI2016 [11] presented promising results from a comparison of HIV combination treatment with TAF and combination treatment with TDF. In study participants with undetectable viral loads who switched from TDF-containing regimens to TAF-containing regimens, 93 percent continued to have undetectable viral loads [12] while their kidney and bone problems improved.

The US Food and Drug Administration (FDA) recently approved several combination pills containing TAF for HIV treatment. These include Genvoya (elvitegravir, cobicistat, emtricitabine, and tenofovir alafenamide), Odefsey (Emtricitabine, Rilpivirine, Tenofovir Alafenamide), and Descovy (Emtricitabine, Tenofovir Alafenamide). For more information on these HIV drugs, see our HIV Drug Chart [13].

## Drugs to Treat Osteoporosis

Diet and exercise are best for keeping bone loss from occurring. They can also be helpful if you already have osteopenia or osteoporosis, but in some cases, your health care provider may also recommend treatment with medication.

Make sure to ask your health care provider about how to take the medication, possible side effects [14], and whether there are any interactions with HIV drugs you take. Some of the osteoporosis medications that are commonly used include:

### Bisphosphonates

These drugs are widely used to treat and prevent osteoporosis. They include:

- Fosamax (alendronate)
- Boniva (ibandronate)
- Actonel or Atelvia (risedronate)
- Zometa, Zomera, Aclasta and Reclast (zoledronic acid)

It is important to get enough calcium and vitamin D when you are taking a bisphosphonate.

## Hormones

- Estrogen: In women, replacing the hormone estrogen has been shown to decrease the number of fractures. Sometimes estrogen is combined with another hormone called progesterone. However, estrogen replacement therapy can increase the risk of developing other diseases, including certain cancers. Because of this, the US Food and Drug Administration (FDA) recommends using other osteoporosis medications. If estrogen/progesterone are used, the lowest possible doses should be considered.
- Testosterone: Testosterone therapy may be useful to slow or reverse decreased bone density and strength in men
- Miacalcin (calcitonin): This naturally occurring hormone slows bone loss and increases bone density in the spine
- Forteo (teriparatide): Forteo is a parathyroid hormone that has been shown to rebuild bone and increases bone mineral density, especially in the spine

## Selective Estrogen Receptor Modulators (SERMs)

Evista (raloxifene) is in a class of osteoporosis drugs called SERMs. Evista was developed to work like estrogen therapy, but with fewer side effects [14].

## Bone-Modifying Agent

Denosumab (Prolia or Xgeva) is given as a shot (injection), and is used to treat postmenopausal women at high risk for fracture. It works by stopping the development of cells that remove bone.

## Protect Your Bones

It is important that women living with HIV, especially those who have experienced menopause [2], pay careful attention to their bone health. Speak to your health care provider and follow these steps to help protect your bones:

- Ask your health care provider if you need a DEXA scan
- Follow a diet with plenty of calcium and vitamin D
- Seek the advice of a registered dietitian if you need help choosing the right foods
- Take calcium supplements if needed (talk to your health care provider first)
- Ask your health care provider what physical activity is safe for you, and start doing it
- Stop smoking [6] and reduce your intake of caffeine and alcohol
- Tell your health care provider if you are experiencing joint pain, especially in the hip area

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## Additional Resources

Select the links below for additional material related to bone health.

[Menopause and Osteoporosis \(Cleveland Clinic\)](#) [31]

[Bone Health ? Prevent Osteoporosis by Easing Inflammation \(Women to Women\)](#) [32]

[Reduced Bone Density and HIV \(CATIE\)](#) [33]

[Tenofovir Alafenamide Equally Effective but Safer for Kidneys and Bones Than Current Formulation \(AIDSmap\)](#) [34]

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[New Guidelines Recommend Routine Screening for Bone Fracture Risk in Over 40s with HIV \(IOF\)](#) [36]

[What Is Osteoporosis? \(National Osteoporosis Foundation\)](#) [37]

[Bone Problems & HIV/AIDS \(The Body\)](#) [38]

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- [32] <https://www.womentowomen.com/bone-health/bone-health-prevent-osteoporosis-by-easing-inflammation/>
- [33] <http://www.catie.ca/en/treatmentupdate/treatmentupdate-189/bone-health/reduced-bone-density-and-hiv>
- [34] <http://www.aidsmap.com/Tenofovir-alafenamide-equally-effective-but-safer-for-kidneys-and-bones-than-current-formulation/page/2949354/>
- [35] [http://www.poz.com/articles/bone\\_guidelines\\_761\\_26764.shtml](http://www.poz.com/articles/bone_guidelines_761_26764.shtml)
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