

# Atrial Fibrillation and Stroke Prevention: Anticoagulants

## Stroke prevention is the cornerstone of AFib treatment.

More than three million Americans have Atrial Fibrillation (AFib), which is the most common heart rhythm disorder. AFib is caused by chaotic electrical signals, which make the upper chambers of the heart (the atria) quiver, instead of properly contracting. During AFib, blood pools in the atria, which can allow a clot to form. If a blood clot breaks free, it can enter the bloodstream and cause a stroke.

People with AFib have a stroke risk five times higher than those who do not have AFib. AFib causes approximately 120,000 ischemic strokes each year. Ischemic stroke happens when a clot breaks free, lodges in a blood vessel, and blocks the flow of blood and oxygen to the brain. One out of every 4 strokes is due to AFib. Because of the adverse effects a stroke can have on quality and duration of life, stroke prevention is a primary treatment goal in AFib. There are a variety of treatments to prevent ischemic stroke, but a medication called an anticoagulant is the most common “first-line”

treatment. Anticoagulants are highly effective at lowering the likelihood of ischemic stroke.

### *What is an Anticoagulant?*

Anticoagulants, which are sometimes called blood thinners, interrupt the blood’s normal clotting (coagulation) process. This complex system, which is called the coagulation cascade, involves many cell proteins that work together to stop bleeding.

### **Anticoagulant Types**

There are several anticoagulant medications available for patients with AFib. Anticoagulants target different parts of the coagulation cascade so blood clots cannot form.

#### ■ **Vitamin K antagonists**

Many of the cell proteins involved in the coagulation cascade rely on vitamin K for synthesis. Vitamin K antagonists (VKAs) interrupt the production of these clotting proteins. VKAs have the longest track record of use in AFib-related stroke prevention. The first VKA was approved in

1954. Warfarin (Coumadin) is a vitamin K antagonist.

#### ■ **Direct thrombin inhibitors**

Thrombin is one of the enzymes involved in clot formation. By inhibiting thrombin, the coagulation cascade is interrupted, so blood clots form less readily. Dabigatran (Pradaxa) is a direct thrombin inhibitor.

#### ■ **Factor Xa inhibitors**

Factor Xa is another enzyme involved in the coagulation cascade. By inhibiting Factor Xa, the coagulation cascade is interrupted. Rivaroxaban (Xarelto) and Apixaban (Eliquis) are Factor Xa inhibitors.

### **Benefits and Risks**

When taken as prescribed, all anticoagulants significantly reduce the risk of ischemic stroke. Patients taking anticoagulants are at risk for excess bleeding because the medications interrupt the coagulation cascade and prevent clotting. Each type of anticoagulant has its own benefits and risks.

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AFib-related strokes have double the mortality rate of non-AFib-related strokes.

### Vitamin K antagonists (VKAs)

**Benefits.** Most doctors are experienced with managing patients taking a VKA. If there is an emergency (such as a car accident) or a planned medical procedure, healthcare professionals can reverse a VKA so that the body's normal clotting abilities return. In addition, VKAs are the least expensive type of anticoagulant.

**Risks.** Certain foods—particularly green, leafy vegetables—have a lot of vitamin K. Eating too many foods that are rich in vitamin K can make VKAs ineffective at stroke prevention. In addition, some medications also interfere with VKAs. Because of this, patients taking a VKA must have their blood tested routinely to make sure it is adequately thinned, but not too thinned. The test measures the International Normalized Ratio (INR), which in general should be maintained between 2.0 and 3.0. If the INR is less than 2, the patient is at risk for an ischemic stroke. If the INR exceeds 3, there is an increased risk of bleeding.

### Direct thrombin inhibitors

**Benefits.** Direct thrombin inhibitors may be easier for some patients to use than a VKA. Direct thrombin inhibitors have fewer dietary restrictions and fewer drug-to-drug interactions than VKAs. Direct thrombin inhibitors do not require frequent blood tests. In addition, direct thrombin inhibitors have a lower risk of bleeding in the brain than VKAs.

**Risks.** Direct thrombin inhibitors offer stroke protection for a certain period of time, so patients cannot skip a dose. Direct thrombin inhibitors may also have a higher risk of major bleeding in the gastro-intestinal organs, including the stomach and intestines.

### Factor Xa inhibitors

**Benefits.** A Factor Xa inhibitor may be easier for some patients to use than a VKA. Similar to direct thrombin inhibitors, Factor Xa inhibitors have fewer dietary restrictions and fewer drug-to-drug interactions than VKAs. Frequent blood tests are not needed. Factor Xa inhibitors also have a lower risk of bleeding in the brain compared to VKAs.

**Risks.** Healthcare professionals are less experienced managing patients taking Factor Xa inhibitors in emergency situations, and there is no approved drug to reverse the anticoagulation effects of Factor Xa inhibitors. Similar to direct thrombin inhibitors, Factor Xa inhibitors offer stroke protection for a certain period of time, so it's important to take the medication as prescribed by the doctor and not skip doses.

## Determining the Appropriate Treatment

Stroke prevention is a primary goal of AFib treatment. Patients should discuss their risk of stroke with their electrophysiologist (a doctor who specializes in heart rhythm disorders), cardiologist or primary care physician. If an anticoagulant is needed, patients should discuss the risks and benefits of the different anticoagulants with their doctor and together, determine which treatment is the best choice.

	Vitamin K Antagonists (Warfarin/Coumadin)	Direct thrombin Inhibitors (Dabigatran/Pradaxa)	Factor Xa Inhibitors (Rivaroxaban/Xarelto and Apixaban/Eliquis)
Dosing frequency	Once/day	Twice/day	Once or twice/day*
Effective at preventing strokes	Yes	Yes	Yes
Risk of excess bleeding	Yes	Yes	Yes
Time to peak effectiveness	Days	0.5-2 hours	2-4 hours
Interaction with food	Yes	No	No
Interaction with other medications	Yes	Fewer	Fewer
Need for frequent blood tests	Yes	No	No
Reversal drug for emergencies	Yes	No**	No**
Need to stop taking drug before medical procedures	Depends	Depends	Depends

\*Some Factor Xa inhibitors only need to be taken once a day with the evening meal. Other Factor Xa inhibitors need to be taken twice a day.

\*\*There is currently no approved reversal drug in the US, however, reversal drugs are under development.

The type of medical or dental procedure will determine whether anticoagulation medication must be stopped. Patients should discuss the planned procedure with their doctor to determine appropriate anticoagulation dosing before and after.