

GENERAL INSTRUCTIONS

DAIDS Core Risk List (side effects/risks of study drug) Template for DAIDS Informed Consent Form

GENERAL INSTRUCTIONS for DAIDS Medical Officers (MOs): Address the Virologic Failure and Development of Resistance in the risk section of the DAIDS Sample Informed Consent Form, if appropriate. This topic is not addressed in this core DAIDS Risk List (side effects/risks of drugs used in the study).

The risks of side effects from the study drug listed below apply to the risk section of the DAIDS Sample Informed Consent Form. These side effects/risks should be addressed in the risk section of the DAIDS Sample Informed Consent Form.

Include the appropriate risk language for ritonavir in the risk section of the DAIDS Sample Informed Consent Form. Refer to the DAIDS ritonavir Core Risk List if applicable.

DAIDS Core Risk List (side effects/risks of study drug) Template for DAIDS Informed Consent Form

General Disclaimer

The drugs used in this study may have side effects, some of which are listed below. Please note that these lists do not include all the side effects seen with these drugs. These lists include the more serious or common side effects with a known or possible relationship. It is very important that you tell your study doctor of any changes in your medical condition while taking part in the study. At any time during the study, if you believe you are experiencing any of these side effects, you have the right to ask questions on possible and /or known risks.

There is a risk of serious and/or life threatening side effects when non-study medications are taken with the study drugs. For your safety, you must tell the study doctor or nurse about all medications you are taking before you start the study and you must ask approval for taking any new medication while you are on the study.

Use of Combination Antiretroviral Drugs

Immune Reconstitution Syndrome:

In some people with advanced HIV infection, symptoms from other infections or certain diseases may occur soon after starting combination anti-HIV treatment but can also occur later. Some of these symptoms may be life threatening. If you start having new symptoms, or notice that existing symptoms are getting worse after starting your antiretroviral therapy, tell your healthcare provider right away.

The use of potent antiretroviral drug combinations may be associated with an abnormal placement of body fat and wasting. Some of the body changes include:

- Increase in fat around the waist and stomach area
- Increase in fat on the back of the neck
- Thinning of the face, legs, and arms
- Breast enlargement

Protease Inhibitors

The use of protease inhibitors may be associated with the following:

- *Increases in the amount of triglycerides and/or cholesterol in the blood*
- *Development of diabetes or the worsening of high blood sugar*

There have been reports of increased bleeding in HIV-infected persons with hemophilia who were treated with protease inhibitors. It is not known if protease inhibitors were the cause of these bleeding episodes.

Darunavir (DRV/rtv, PREZISTA/rtv, PREZISTA™*)

The following serious side effects have been associated with the use of darunavir given together with ritonavir (RTV). Contact your healthcare provider if you have any of the symptoms of liver problems that are listed and any skin changes with the symptoms listed below.

- People taking darunavir together with ritonavir may develop severe liver problems, which may be life-threatening. People who have increased liver function tests before starting darunavir and people with liver diseases such as Hepatitis B or C have an increased risk of worsening liver disease. If you are developing liver problems, you may have one or more of the following symptoms: yellowing of the skin or whites of your eyes, dark urine, and pain on the right side of your stomach, loss of appetite, upset stomach or vomiting, pale colored stools, itchy skin.
- Rash, which may be severe or life-threatening? Contact your Healthcare Provider if you develop a rash.
- If you develop any skin changes with the following symptoms stop using DRV/rtv combination and contact your Health Care Provider right away:
 - Fever
 - Tiredness
 - Muscle or joint pain
 - Blisters
 - Mouth sores
 - Red or inflamed eyes.

Additional side effects include:

- Diarrhea
- Nausea
- Stomach pain
- Vomiting
- Headache

- Other changes in blood test results that may show problems with the liver , kidneys and cholesterol levels
- Inflammation of the pancreas. When the pancreas becomes inflamed, it can cause pain in the stomach, nausea, vomiting
- Joint stiffness and bone pains may occur, rarely death of bone tissue and collapse of the bone may occur.

Note: Before starting darunavir, you should inform your healthcare provider if you are allergic to sulfa medicines.

Your healthcare provider can provide more complete information about the side effects of ritonavir (the medicine you need to take together with DRV).