ANTI-HIV DRUG

a guide to the interactions found between anti-hiv drugs and common hiv-related therapies

With the number of drugs available to treat HIV and with those that prevent or treat opportunistic infections (OIs), the potential for drug interactions increases. Developing a health management plan and deciding which therapies to include in that plan may seem a daunting task. Not only does each therapy have possible side effects, but each might increase or decrease the benefit of the other drugs you take.



Drug interactions can take various forms, occurring immediately or over several weeks. Some drugs simply should not be used together, while others can be combined only if done with careful monitoring to detect emergency problems. Interactions can also occur when one therapy alters the effect of another. This includes how the other is absorbed in the body, broken down (*metabolized*), distributed or excreted.

A common interaction can occur when two drugs have similar side effect profiles. For example, both ddI and ddC can cause a tingling or pain in the legs, hands or feet (*peripheral neuropathy*). It's not recommended that they be used together because the similar side effects may increase the potential for neuropathy. Similarly, AZT and ganciclovir (a treatment for CMV) may both cause bone marrow suppression, resulting in anemia. However, adding a third drug, G-CSF (Neupogen), can help manage this interaction. In addition, higher blood levels of a drug increase the chance of more side effects.

A PUBLICATION FROM



AUGUST 2004

talking to your doctor about drug interactions

Many people take a variety of therapies at the same time. These can range from experimental and approved anti-HIV drugs to complimentary and over-the-counter medicines. Drug interactions may play a major role in the success of any treatment plan. Unfortunately, they're not always considered when developing a treatment strategy. The following are some suggestions to help prevent drug interactions:

- BROWN BAG MEDICINE CHECKUP. Each time you see your healthcare provider, put all your meds—including over-the-counte and complimentary products—in a bag. Take the bag with you and have your doctor review those medicines for safety, appropriatene compatibility and instructions for use.
- NEW PRESCRIPTIONS. Each time your doctor prescribes you a r medication, ask him or her if it can be combined safely with your other therapies.
- REGULAR DISCUSSION. Talk to your doctor about making the 'medicine checkup' part of your regular visits. Discuss how best to check for possible drug interactions. (Bring Project Inform's publication, *Drug Interactions*, with you to your appointment.)



As it has become standard medical practice to prevent multiple OIs with different drugs, drug interactions become more of a concern. In some prevention regimens, drug interactions may even cause more harm than good. For example, one drug might reduce the blood levels of another drug, leading to drug resistance. This could result in a particular disease (like HIV or hepatitis) becoming unresponsive to treatment. The added toxicity of many therapies taken together may also outweigh their hopeful benefit for preventing disease. Therefore, healthcare providers and people with HIV should make informed decisions about combining therapies and OI prevention regimens and should carefully check for drug interactions and other side effects.

Unfortunately, most drug interaction studies have compared only two drugs, although most people with HIV often take many more than two. As a result, very little is known about how all the commonly used drugs interact with each other.

In the meantime, it's important to discuss possible drug interactions with your doctor and pharmacist. Before starting a new therapy (experimental, approved or complimentary), consider the possible drug interactions and side effects. Not everyone experiences side effects; and many drug interactions can be managed by monitoring carefully, adjusting the doses, or stopping the therapy as needed.

The following chart should only be used as a guide for possible drug interactions. Remember that these interactions might occur in some people, but not in others. This chart was put together with information from prescription package inserts, anecdotal reports, discussions with pharmacologists and doctors who treat HIV disease, and discussions with drug companies.

glossary and instructions

Throughout the text of drug interactions on pages 3–27, you may find terms and instructions you're not familiar with. A glossary of these terms and dosing instructions can be found on page 27.

Abacavir (Ziagen) plus ...

) 3TC

Decreases 3TC level by 15% in blood. No dose adjustment recommended.

> Alcohol

Should be used together with caution. Increases abacavir level by 41% in blood. No dose adjustment recommended.

- > Amprenavir Increases amprenavir level by 29% in blood. No dose adjustment recommended.
-) AZT

Increases AZT level by 10% in blood. No dose adjustment recommended.

- Chloral hydrate Should be used together with caution. May increase chloral hydrate level in blood.
- > Methadone May decrease methadone level in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Tenofovir Should be used together with caution. Although blood levels not affected, tenofovir and abacavir used in combination may speed the development of resistance to both.

Acyclovir (Zovirax) plus ...

> Probenecid Increases acyclovir level in blood.

Adefovir (Hepsera) plus ...

> Amikacin Should not be used together. Increases risk of kidney toxicity.

- > Amphotericin B Should not be used together*. Increases risk of kidney toxicity.
- Cidofovir Should not be used together*. Increases risk of kidney toxicity.
- > Foscarnet Should not be used together*. Increases risk of kidney toxicity.
- > Gentamicin Should not be used together*. Increases risk of kidney toxicity.
- > Ibuprofen Should be used together with caution. Increases adefovir level by 23% in blood and may increase risk of kidney toxicity.
- > Kanamycin Should not be used together*. Increases risk of kidney toxicity.
- > NSAIDS Should not be used together*. May increase risk of kidney toxicity.
- > Pentamidine (IV) Should not be used together*. Increases risk of kidney toxicity.
- > Tenofovir Should not be used together*. Increases risk of kidney toxicity.

Adefovir plus ... (continued)

- > Tobramycin
 - Should not be used together*. Increases risk of kidney toxicity.

Amphotericin B (Fungizone) plus ...

> Adefovir

Should not be used together.* Increases risk of kidney toxicity.

- Amikacin Should be used together with great caution. May increase risk of kidney toxicity.
- Anticancer drugs Should be used together with great caution. May increase risk of kidney toxicity and low blood pressure.
-) azt
- May increase risk of bone marrow toxicity. > Cidofovir
- Should not be used together*. Increases risk of kidney toxicity.
- Clotrimazole May interfere with the activity of amphotericin B.
- Corticosteroids Should be used together with caution. May decrease potassium level and increases risk of heart problems.
- Cyclosporine Should be used together with great caution. May increase risk of kidney toxicity.
- > ddC May increase ddC level in blood and increase risk of peripheral neuropathy.
- Digitalis
 Use together with great caution. May increase risk of heart problems.
- > Fluconazole May interfere with the activity of amphotericin B.
- Flucytosine Increases antifungal activity in test tubes but also increases risk of bone marrow and kidney toxicity.
- Foscarnet Should be used together with great caution. May decrease number of red blood cells and increases risk of kidney toxicity.
- Ganciclovir Should be used together with great caution. May increase risk of bone marrow toxicity.
- Gentamicin Should be used together with great caution. May increase risk of kidney toxicity.
- > Itraconazole May interfere with the activity of amphotericin B.
- Kanamycin Should be used together with great caution. May increase risk of kidney toxicity.
- > Ketoconazole May interfere with the activity of amphotericin B.

Amphotericin B plus ... (continued)

> Miconazole

May interfere with the activity of amphotericin B.

- > Pentamidine (IV) Should be used together with great caution. May increase risk of kidney toxicity.
- > Tenofovir Should not be used together*. May increase risk of side effects.
- > Tobramycin Should be used together with great caution. May increase risk of kidney toxicity.

Amprenavir (Agenerase) plus ...

- > Abacavir
 - Increases amprenavir level by 29% in blood. No dose adjustment recommended.
- Alcohol Should not be used together. Alcohol and the oral formulation of amprenavir increases risk of side effects.
- Alprazolam May increase alprazolam level in blood.
- Amiodarone Should be used together with caution. May increase risk of side effects. Requires monitoring concentration of amiodarone.
- > Antacids Amprenavir should be taken 1 hour before or 1 hour after taking antacids.
- > Astemizole Should not be used together*. Increases risk of side effects.
- > Atazanavir Atazanavir may increase amprenavir levels in blood.
- > Atorvastatin Should be used together with great caution. May increase atorvastatin level in blood.
- AZT Increases AZT level by 31% and increases amprenavir level by 13% in blood. No dose adjustment recommended.
- > Bepridil Should not be used together*. Increases risk of side effects.
- Carbamazepine Should be used together with caution. May decrease amprenavir level in blood.
- Cimetidine May increase amprenavir level in blood.
- Cisapride Should not be used together*. Increases risk of side effects.
- Clarithromycin Increases amprenavir level by 18% in blood. No dose adjustment recommended.
- Clorazepate May increase clorazepate level in blood.
- Clozapine May increase clozapine level in blood.

Amprenavir plus ... (continued)

> Dapsone

May increase dapsone level in blood.

- > ddl or ddl EC Should be taken 1 hour before or after taking ddI tablets. ddI EC may be taken at the same time as amprenavir, but only on an empty stomach.
- > Delavirdine May increase amprenavir level in blood. No dose adjustment recommended.
- > Diazepam May increase diazepam level in blood.
 > Dibudra association
- Dihydroergotamine Should not be used together*. Increases risk of side effects.

 > Diltiazem May increase diltiazem level in blood.

- Disulfiram Should not be used together with amprenavir oral solution*. Increases risk of side effects.
- Efavirenz Decreases amprenavir level by 36% and increases efavirenz level by 15% in blood. Amprenavir dose may need to be increased.
- > Ergotamine Should not be used together*. Increases risk of side effects.
- > Erythromycin May increase erythromycin and amprenavir levels in blood.
- > Ethinyl estradiol Should not be used together. Amprenavir may decrease ethinyl estradiol and norethindrone levels in blood. Use another form of contraception.
- > Felodipine
- May increase felodipine level in blood.
- > Flurazepam
- May increase flurazepam level in blood.
- Food (high fat) Decreases amprenavir level in blood and should be avoided.
- Indinavir Increases amprenavir level by 33% and decreases indinavir level by 38% in blood. No dose adjustment recommended.
- Itraconazole May increase itraconazole and amprenavir level in blood. Check for toxicity.
- > Kaletra

Decreases lopinavir level by 15% and increases amprenavir trough level about 2fold in blood. Amprenavir dose should be reduced to 750mg twice a day or less. However, optimal dose is unclear.

- Ketoconazole Increases amprenavir level by 31% and increases ketoconazole level by 44% in blood. Impact of interaction is uncertain.
- > Lidocaine (systemic) Should be used together with caution. Requires monitoring concentration of lidocaine.
- > Loratidine
- May increase loratidine level in blood.

Amprenavir plus ... (continued)

> Lovastatin

Should be used together with great caution. May increase lovastatin level in blood.

- > Metronidazole Should not be used together with amprenavir oral solution*. Increases risk of side effects.
- > Midazolam Should not be used together*. Increases risk of side effects.
- > Nelfinavir Increases nelfinavir level by 15% in blood. No dose adjustment recommended.
- > Nevirapine May decrease amprenavir level in blood. Dose of amprenavir may need to be increased.
- > Norethindrone Should not be used together. Amprenavir may decrease ethinyl estradiol and norethindrone levels in blood. Use another form of contraception.
- Nicardipine May increase nicardipine level in blood.
- > Nifedipine May increase nifedipine level in blood.
- > Nimodipine May increase nimodipine level in blood.
 > Phenobarbital Should be used to gether with coution. May
- Should be used together with caution. May decrease amprenavir level in blood.
- > Phenytoin Should be used together with caution. May decrease amprenavir level in blood.
- > Pimozide
- May increase pimozide level in blood.
- Pravastatin Should be used together with great caution. May increase pravastatin level in blood.
- Quinidine Should be used together with caution. Requires monitoring concentration of quinidine.
- Rifabutin Decreases amprenavir level by 15% and increases rifabutin level by 193% in blood. A lower dose of rifabutin required.
- > Rifampin Should not be used together*. Significantly decreases amprenavir level in blood.
- > Ritonavir Increases amprenavir level by 70% in blood. Alternate dosing is amprenavir 600mg + ritonavir 100mg twice a day, or amprenavir 1,200mg + ritonavir 200mg once daily.
- > Saquinavir (Fortovase) Decreases amprenavir level by 32% and saquinavir level by 19% in blood. No dose adjustment likely required.
- Sildenafil Should be used together with caution. May increase sildenafil level in blood.
- Simvastatin
 Should be used together with great caution.
 May increase simvastatin level in blood.

Amprenavir plus ... (continued)

- > St. John's Wort
- Should be used together with great caution. May significantly decrease amprenavir level in blood.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > Triazolam Should not be used together*. Increases risk of side effects.
- Tricyclic antidepressants
 Should be used together with caution.
 Requires monitoring concentration of tricyclic antidepressants.
- Vardenafil Should be used together with caution. May increase vardenafil level in blood.
- Vitamin E Amprenavir capsules contain more than the recommended daily allowance for vitamin E. Additional supplementation, especially when used together with drugs like warfarin could result in hemorrhage.
- > Warfarin Should be used together with caution. Requires monitoring concentration of warfarin.

Atazanavir (Reyataz) plus...

- > Amiodarone Increases risk of side effects. Check amiodarone level and use with caution.
- > Amprenavir Atazanavir may increase amprenavir levels in blood.
- Antacids May decrease atazanavir level in blood. Take atazanavir 2 hours before or 1 hour after taking antacids.
- Atorvastatin Should be used together with great caution. May increase risk of side effects.
- > Bepridil Should not be used together*. Increases risk of side effects.
- Calcium channel blockers Should be used together with great caution. Decrease dose of calcium channel blockers and check ECG.
- Cisapride Should not be used together*. Increases risk of side effects.
- Clarithromycin Increases risk of side effects and may decrease effectiveness of clarithromycin. Consider alternatives or reduce clarithromycin dose by at least 50%.
- > Cyclosporin Should be used together with caution. Check cyclosporine level.

Atazanavair plus ... (continued)

> ddI or ddI EC

ddI buffered tablets can significantly decrease atazanavir level in blood. Take atazanavir (with food) either 2 hours before or 1 hour after buffered ddI tablets. Although ddI EC should not affect atazanavir level, they should be taken at least 1 hour apart. Atazanavir should be taken with food, and ddI EC should be taken on an empty stomach.

) Diltiazem

Should be used together with great caution. Decrease diltiazem dose by at least 50% and check ECG.

- > Ergot derivatives Should not be used together*. Increases risk of side effects.
- > Efavirenz

Decreases atazanavir level in blood. Atazanavir should be boosted with ritonavir if used with efavirenz.

- Ethinyl estradiol Should be used together with caution. Atazanavir increases ethinyl estradiol level in blood. Use lowest effective ethinyl estradiol dose.
- > Food

High fat meal increases atazanavir by 35% and low fat meal increases atazanavir level by 70% in blood. Atazanavir should be taken with food.

Indinavir Should not be used together*. Increases risk of side effects.

- > Irinotecan Atazanavir may interfere with irinotecan metabolism. May increase irinotecan side effects.
- > Lidocaine (systemic) Increases risk of side effects. Check lidocaine level and use with caution.
- > Lovastatin Should not be used together*. Increases risk of side effects.
- > Midazolam Should not be used together*. Increases risk of side effects.
- > Norethindrone Should be used together with caution and decrease norethindrone dose to lowest effective dose. May increase norethindrone side effects, particularly in diabetic women.
- > Pimozide Should not be used together*. Increases risk of side effects.
- > Proton-pump inhibitors Should not be used together. May significantly decrease atazanavir level in blood.
- > Quinidine Increases risk of side effects. Check quinidine level and use with caution.
- Rifabutin Increases rifabutin level. Decrease dose of rifabutin to 150mg every other day or 3 times a week.
- > Rifampin Should not be used together. Decreases level of most protease inhibitors in blood.

Atazanavair plus ... (continued)

- > Ritonavir
- Substantially increases atazanavir blood level. If combined, use atazanavir 300mg (with food) + ritonavir 100mg once daily.
- Saquinavir (Fortovase) Increases saquinavir level by 5–6 fold in blood. Dose adjustments currently under study.
- Sildenafil Increases sildenafil blood level. Decrease sildenafil dose to 25mg every 48 hrs and check for side effects.
- Simvastatin Should not be used together*. Increases risk of side effects.
- Sirolimus Should be used together with caution. Check sirolimus level.
- St. John's Wort Should not be used together. May reduce atazanavir level in blood.
- Tacrolimus Should be used together with caution. Check tacrolimus level.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- Tenofovir Increases tenofovir level by 24% and decreases atazanavir level by up to 40% in blood. Atazanavir should be boosted with ritonavir if used with tenofovir.
- Triazolam
 Should not be used together*. Increases risk of side effects.
- > Tricyclic antidepressants Increases risk of side effects. Check antidepressant level and use with caution.
- > Vardenafil Should be used together with caution. May increase vardenafil level in blood.
- > Warfarin Should be used together with great caution. Has not been studied but may lead to serious side effects.

Atovaquone (Mepron) plus ...

) AZT

Increases AZT level by 35% in blood. No dose adjustment recommended.

- > Fatty foods
- Increases atovaquone level in blood. Atovaquone should be taken with food.
- Kaletra Decreases atovaquone level in blood. Atovaquone dose may have to be increased.
- > Rifabutin Should be used together with caution. May decrease atovaquone level in blood.
- Rifampin Should be used together with caution. Decreases atovaquone level by 50% in blood.
- > Ritonavir May decrease atovaquone level in blood.

Atovaquone plus ... (continued)

> TMP/SMX

Increases TMP/SMX blood level by 17% and 8%, respectively. No dose adjustment recommended.

Azithromycin (Zithromax) plus ...

- > AZT
 - May increase AZT level in blood. No dose adjustment recommended.
- Cyclosporine May increase cyclosporine level in blood.
- > ddl or ddl EC May increase ddI level in blood. No dose adjuctment recommanded
- adjustment recommended.
 > Digoxin
 - May increase digoxin level in blood.
- > Ergot derivatives May increase ergot drug level in blood.
- Food Azithromycin may be taken with or without food.
- > Hexobarbital May increase hexobarbital level in block
- May increase hexobarbital level in blood. > Midazolam

May increase midazolam level in blood. No dose adjustment recommended.

> Nelfinavir Should be used together with caution. Increases azithromycin level in blood. No dose adjustment recommended, but check for hearing and liver side effects.

- > Phenytoin
 - May increase phenytoin level in blood.
- > Rifabutin May increase rifabutin level in blood. No dose adjustment recommended.
- > Theophylline May increase theophylline level in blood. No dose adjustment recommended.
- > Terfenadine May increase terfenadine level in blood.
- > Warfarin May increase warfarin level in blood.

Birth control pills

See section on Ethinyl estradiol.

Cidofovir (Vistide) plus ...

> Adefovir

Should not be used together*. Increases risk of kidney toxicity.

- > Amikacin Should not be used together*. Increases risk of kidney toxicity.
- > Amphotericin B Should not be used together*. Increases risk of kidney toxicity.
- > Foscarnet Should not be used together*. Increases risk of kidney toxicity.

Cidofovir plus ... (continued)

- > Gentamicin Should not be used together*. Increases risk of kidney toxicity.
- Kanamycin Should not be used together*. Increases risk of kidney toxicity.
- > Pentamidine (IV) Should not be used together*. Increases risk of kidney toxicity.
- > Probenecid Needs to be taken together to decrease the risk of kidney toxicity.
- Streptomycin Should be used together with great caution. Increases risk of kidney toxicity.
- > Tenofovir Should not be used together*. May increase risk of side effects.
- > Tobramycin Should not be used together*. Increases risk of kidney toxicity.

Ciprofloxacin (Cipro) plus ...

- Antacids Needs to be taken 2 hours before or 6 hours after taking ciprofloxacin, otherwise
- decreases ciprofloxacin level in blood.Caffeine May increase caffeine level in blood.
- Cyclosporine May increase risk of elevated serum creatinine.
- > ddl or ddl EC Ciprofloxacin needs to be taken 2 hours before or 6 hours after ddl buffered tablets, otherwise ddl may decrease ciprofloxacin level in blood. Ciprofloxacin may be taken at the same time as ddl EC capsules.
- Food May be taken with or without food, but preferred time of dosing is 2 hours after a meal.
- > Iron supplements Needs to be taken 2 hours before or 6 hours after taking ciprofloxacin, otherwise decreases ciprofloxacin level in blood.
- > Methotrexate May increase blood level of methotrexate. If used together, check for methotrexate side effects.
- Probenecid Increases ciprofloxacin level in blood.
- Sucralfate Needs to be taken 2 hours before or 6 hours after taking ciprofloxacin, otherwise decreases ciprofloxacin level in blood.
- Theophylline Should not be used together.* Increases risk of serious side effects. If concurrent use cannot be avoided, check theophylline level and adjust dose.
- > Warfarin May increase warfarin level in blood. Check for warfarin toxicity.

Ciprofloxacin plus ... (continued)

> Zinc-containing multivitamins Needs to be taken 2 hours before or 6 hours after taking ciprofloxacin otherwise decreases ciprofloxacin level in blood.

Clarithromycin (Biaxin) plus ...

- > Alprazolam May increase midazolam blood level. Check for alprazolam side effects.
- Amprenavir Increases amprenavir level by 18% in blood. No dose adjustment recommended.
- > Anticoagulants May increase anticoagulants effect. Check prothrombin (coagulation) time.
- > Astemizole Should not be used together*. May increase risk of heart problems.
- Atazanavir Increases risk of side effects and may decrease clarithromycin effectiveness. Consider alternatives or reduce clarithromycin dose by at least 50%.
- Atorvastatin May increase risk of atorvastatin side effects.
- > AZT Decreases AZT level by 25% in blood. Consider taking at least 2 hours apart.
- > Carbamazepine Increases carbamazepine level in blood. Check carbamazepine level.
- > Cisapride Should not be used together*. Increases risk of side effects.
- Cyclosporine May increase cyclosporine level in blood.
- > Delavirdine Interaction is complex. Clarithromycin dose should be adjusted for people with kidney problems.
- Digoxin May increase digoxin level in blood. Check digoxin level.
- Disopyramide May increase risk of side effects. Check disopyramide level and QTc prolongation.
- Efavirenz Slightly increases efavirenz level but decreases clarithromycin level by 39% (increases 14-OH clarithromycin level by 34%) in blood. No dose adjustment currently recommended.
- > Ergot derivatives Should be used together with caution. May increase risk for serious side effects.
- Fluvastatin May increase risk of fluvastatin side effects.
- Indinavir Increases clarithromycin level by 53% and indinavir level by 29% in blood. No dose adjustment recommended.
- > Kaletra
 - Increases clarithromycin level in blood. Clarithromycin dose should be decreased for people with kidney dysfunction.

Clarithromycin plus ... (continued)

- > Loratadine
- May affect loratidine and clarithromycin levels in blood. No dose adjustment currently recommended.
- > Lovastatin
 - May increase risk of lovastatin side effects.
 - > Midazolam
 - May increase midazolam blood level. Check for midazolam side effects.
 - > Nevirapine Consider using alternatives to clarithromycin. Decreases clarithromycin level by 30% and increases nevirapine level by 26% in blood.
 - > Pimozide Should not be used together.* Increases risk of serious side effects.
- > Phenytoin May increase phenytoin level in blood.
- > Pravastatin May increase risk of pravastatin side effects.
- Quinidine May increase risk of side effects. Check quinidine level and QTc prolongation.
- > Rifabutin Should be used together with caution. May increase rifabutin level and decrease clarithromycin level in blood.
- > Rifampin Should be used together with caution. May decrease clarithromycin level in blood.
 - Ritonavir Increases clarithromycin level by 77% and increases ritonavir level by 12% in blood. Requires lower dose of clarithromycin and monitoring for kidney toxicity in people with a history of kidney problems.
 - > Saquinavir (Fortovase) Increases saquinavir level by 177% and increases clarithromycin level by 45% (but decreases 14-OH clarithromycin level by 24%) in blood. No dose adjustments currently recommended.
 - Sildenafil May increase sildenafil level in blood. Consider dose reduction.
 - Simvastatin May increase risk of simvastatin side effects.
 - St. John's Wort Should be used together with great caution. May decrease clarithromycin level in blood.
 - > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
 - > Theophylline Increases theophylline level in blood. Check theophylline level.
- Triazolam May increase triazolam level in blood. Check for triazolam side effects.
- > Trimetrexate May affect trimetrexate and/or clarithromycin level in blood.

Clarithromycin plus ... (continued)

- Vardenafil Should be used together with caution. May increase vardenafil level in blood.
- > Warfarin Check prothrombin (coagulation) time.

Clindamycin (Cleocin) plus ...

- > Erythromycin Should not be used together*. Causes erythromycin and clindamycin to be less effective.
- > Neuromuscular blocking agents May increase neuromuscular blocking effect.

Clofazimine (Lamprene) plus ...

> Dapsone May decrease effectiveness of clofazimine.

Combivir plus ...

Combivir is a single pill containing 2 anti-HIV drugs—AZT and 3TC. If you take Combivir as part of your regimen, refer separately to drug interactions for AZT and 3TC.

Cycloserine (Seromycin) plus ...

- Alcohol May increase risk of seizures.
 Ethiopoppide
- Ethionamide May increase risk of central nervous system related toxicity.
- > Isoniazid May increase risk of central nervous system related toxicity.

Dapsone plus ...

- Amprenavir May increase dapsone level in blood.
-) azt
- May increase risk of bone marrow toxicity. > Clofazimine
- May decrease effectiveness of clofazimine.
- May increase risk of peripheral neuropathy. > ddl

ddI buffered tablets and dapsone need to be taken 2 hours apart, otherwise dapsone has no activity.

-) Ganciclovir
- May increase risk of bone marrow toxicity. > Probenecid

Should be used together with caution. May increase dapsone level in blood and increase risk of side effects.

 > Pyrimethamine May increase risk of bone marrow toxicity.
 > Difabution

Rifabutin Should be used together with caution. May decrease dapsone level in blood.

Dapsone plus ... (continued)

- > Rifampin Should be used together with caution. Decreases dapsone level 7–10 times in blood.
- Saquinavir (Invirase) May increase dapsone level in blood. Check for dapsone toxicity.
- TMP/SMX Increases trimethoprim level by 1.5 times; increases dapsone level by 1.5 times in blood.

Delavirdine (Rescriptor) plus ...

- > Alprazolam Should not be used together*. Increases risk of side effects.
- > Amlodipine Should be used together with caution. May increase amlodipine level in blood. Check for toxicity.
- > Amphetamines Should be used together with caution. May increase amphetamine level in blood.
- > Amprenavir May increase amprenavir level in blood. No dose adjustment recommended.
- > Antacids Needs to be taken 1 hour apart, otherwise decreases delavirdine level by 44% in blood.
- > Astemizole Should not be used together*. Increases risk of side effects.
- Atorvastatin Should be used together with great caution. Use lowest possible dose of atorvastatin. May increase risk of side effects.
- > Barbiturates May decrease delavirdine level in blood.
- > Bepridil Should be used together with caution. May increase bepridil level in blood and side effects.
- Carbamazepine Should not be used together. May significantly decrease delavirdine level in blood.
- Cimetidine Should be used together with caution. May decrease delavirdine level in blood.
- Cisapride Should not be used together*. Increases risk of side effects.
- Clarithromycin Increases delavirdine level by 31% and increases clarithromycin level by 100% (but decreases 14-OH clarithromycin level by 75%) in blood. Clarithromycin dose should be adjusted for people with kidney problems.
- > Cyclosporine Should be used together with caution. May increase cyclosporine level in blood. Check cyclosporine level.
- > ddl or ddl EC Needs to be taken 1 hour apart, otherwise decreases delavirdine level in blood.

Delavirdine plus ... (continued)

> Dexamethazone

Should be used together with caution. May decrease delavirdine level in blood.

- Dihydroergotamine Should not be used together*. Increases risk of side effects.
- > Diltiazem Should be used together with caution. May increase diltiazem level in blood. Check for toxicity.
- > Ergonovine Should not be used together*. Increases risk of side effects.
- Ergotamine Should not be used together*. Increases risk of side effects.
- > Erythromycin May increase delavirdine level in blood.
- > Ethinyl estradiol May increase ethinyl estradiol level in blood. Impact of interaction is uncertain.
- Famotidine Should be used together with caution. May decrease delavirdine level in blood.
- > Felodipine Should be used together with caution. May increase felodipine level in blood. Check for toxicity.
- > Flecainide Should be used together with caution. Check flecainide level.
- > Fluoxetine Increases delavirdine level by 50% in blood.
- Fluvastatin Should be used together with great caution. Use lowest possible dose of fluvastatin. May increase risk of side effects.
- > Fosamprenavir

Should not be used together. May significantly decrease fosamprenavir level in blood.

- Indinavir Increases indinavir level 50–100% in blood. Indinavir may be reduced to 600mg 3 times a day.
- > Isradipine Should be used together with caution. May increase isradipine level in blood. Check for toxicity.
- > Itraconazole
- May increase delavirdine level in blood. > Kaletra
 - May increase lopinavir level in blood. Needs further study.
- > Ketoconazole Increases delavirdine level by 50% in blood.
- > Lansoprazole Should be used together with caution. May decrease delavirdine level in blood.
- > Lidocaine (systemic) Should be used together with caution. Check lidocaine level.
- > Loratadine

Should be used together with caution. May increase loratadine level in blood.

Delavirdine plus ... (continued)

> Lovastatin Should not be used together*. Increases risk of side effects.

> Methadone Increases methadone level in blood. May require lower dose of methadone.

- > Methylergonovine Should not be used together*. Increases risk of side effects.
- > Midazolam Should not be used together*. Increases risk of side effects.
- > Nelfinavir Increases nelfinavir level about 100% but decreases nelfinavir active metabolite by about 45%. Decreases delavirdine level by about 40% in blood. No data exist to guide dose adjustments.
- > Nicardipine Should be used together with caution. May increase nicardipine level in blood. Check for toxicity.
- > Nifedipine Should be used together with caution. May increase nifedipine level in blood. Check for toxicity.
- > Nisoldipine Should be used together with caution. May increase nisoldipine level in blood. Check for toxicity.
- > Nizatidine Should be used together with caution. May decrease delavirdine level in blood.
- > Omeprazole Should be used together with caution. May decrease delavirdine level in blood.
- > Phenobarbital Should not be used together. May significantly decrease delavirdine level in blood.
- > Phenytoin Should not be used together. May significantly decrease delavirdine level in blood.
- > Pimozide Should not be used together*. Increases risk of side effects.
- > Propafenone Should be used together with caution. Check propafenone level.
- Quinidine
 Should be used together with caution. Check quinidine level.
- Ranitidine Should be used together with caution. May decrease delavirdine level in blood.
- Rapamycin Should be used together with caution. May increase rapamycin level in blood. Check rapamycin level.
- Rifabutin Should not be used together*. Significantly decreases delavirdine level and increases rifabutin level in blood.
- > Rifampin

Should not be used together*, otherwise significantly decreases delavirdine level in blood.

Delavirdine plus ... (continued)

- > Ritonavir
 - Increases ritonavir level by about 70% in blood. No dose adjustment currently recommended.
- Saquinavir (Fortovase) Increases saquinavir level by 5 times in blood. Decrease saquinavir dose to 800mg 3 times a day.
- > Saquinavir (Invirase) Increases saquinavir level by 5 times in blood. May increase risk of gastrointestinal side effects and liver function tests. Decrease saquinavir dose to 800mg 3 times a day.
- Sildenafil Should be used together with caution. Increases sildenafil level in blood. Decrease sildenafil dose to no more than 25mg in a 48 hour period.
- Simvastatin Should not be used together*. Increases risk of side effects.
- St. John's Wort Should not be used together. May significantly decrease delavirdine level in blood.
- Tacrolimus Should be used together with caution. May increase tacrolimus level in blood. Check tacrolimus level.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > Triazolam Should not be used together*. Increases risk of side effects.
- > Vardenafil Should be used together with caution. May increase vardenafil level in blood.
- > Verapamil Should be used together with caution. May increase verapamil level in blood. Check for toxicity.
- Voriconazole Should be used together with caution. May increase delavirdine side effects and decrease effectiveness of voriconazole.
- > Warfarin Should be used together with caution. Check INR.

Didanosine (ddl, Videx) or (ddl EC, Videx EC) plus ...

EC stands for Enteric Coated.

- Alcohol Heavy alcohol intake may increase risk of pancreatitis.
- Allopurinol Should not be used together. Increases ddI level 113–312% in blood.
- > Amprenavir Should be taken 2 hours before or 1 hour after taking ddI tablets. ddI EC may be taken at the same time as amprenavir, but only on an empty stomach.

Didanosine plus ... (continued)

- **>** Atazanavir
- ddI buffered tablets can significantly decrease atazanavir level in blood. Take atazanavir (with food) either 2 hours before or 1 hour after buffered ddI tablets. Although ddI EC should not affect atazanavir level, they should be taken at least 1 hour apart. Atazanavir should be taken with food and ddI EC should be taken on an empty stomach.
- > Anticancer drugs May increase risk of peripheral neuropathy.
- > Azithromycin May increase AZT level in blood. No dose adjustment recommended.
- > Cimetidine
- Needs to be taken at least 2 hours apart.
- Ciprofloxacin Ciprofloxacin needs to be taken 2 hours before or 6 hours after ddI buffered tablets, otherwise ddI may decrease ciprofloxacin level in blood. Ciprofloxacin may be taken at the same time as ddI EC capsules.
- > Dapsone Needs to be taken 2 hours apart, otherwise dapsone has no activity.
- > ddC Should not be used together. Significantly increases risk of peripheral neuropathy.
- > Delavirdine Needs to be taken 1 hour apart, otherwise decreases delavirdine level in blood.
- **)** d4T

Should be used together with great caution. Increases risk of pancreatitis, especially in pregnant women.

- Food Decreases ddI level in blood. ddI and ddI EC should be taken on an empty stomach, at least 2 hours after eating.
- Ganciclovir
 Should be used with caution. Significantly increases ddI level in blood. May increase risk of pancreatitis.
- > Hydroxyurea Increases anti-HIV activity in test tubes. May increase risk of pancreatitis.
- Indinavir May decrease ddI and indinavir levels in blood. Indinavir should be taken 1 hour before or 2 hours after taking ddI.
- Itraconazole Needs to be taken 2 hours apart, otherwise ddI buffering agents may significantly decrease itraconazole level in blood.
- Xaletra ddI or ddI EC should be taken 1 hour before or 2 hours after taking Kaletra. ddI and ddI EC should be taken on an empty stomach, and Kaletra should be taken with a meal.
- > Ketoconazole Needs to be taken 2 hours apart, otherwise ddI buffering agents may significantly decrease ketoconazole level in blood.
- > Methadone Decreases ddI level by 60% in blood.

Didanosine plus ... (continued)

> Nelfinavir

Nelfinavir should be taken 2 hours before or after taking ddI or ddI EC. Nelfinavir should be taken with food, and ddI and ddI EC should only be taken on an empty stomach.

- > Pentamidine (IV) May increase risk of pancreatitis.
- Quinolones Quinolones need to be taken 2 hours before or 6 hours after ddI buffered tablets, otherwise ddI may decrease quinolone level in blood. Quinolones may be taken at the same time as ddI EC capsules.

Ribavirin Should not be used together*. Increases risk of pancreatitis and mitochondrial toxicity.

- Ritonavir Decreases ddI level by 13% in blood. No dose adjustment recommended, but take 2½ hours apart.
- > Saquinavir Saquinavir should be taken 2 hours before or after taking ddI or ddI EC. Saquinavir is best taken with food, and ddI and ddI EC should only be taken on an empty stomach.
- > Tenofovir

Should be used together with great caution. Increases ddI level by 44% in blood. May increase risk of ddI side effects.

> Tetracyclines Should not be used together*. May increase risk of pancreatitis.

Efavirenz (Sustiva) plus ...

> Amprenavir

Decreases amprenavir level by 36% and increases efavirenz level by 15% in blood. Amprenavir dose may need to be increased.

- Atazanavir Decreases atazanavir level in blood. Atazanavir should be boosted with ritonavir if used with efavirenz.
- Atorvastatin Decreases atorvastatin level in blood. Atorvastatin dose may need to be increased.
- Clarithromycin Slightly increases efavirenz level but decreases clarithromycin level by 39% (increases 14-OH clarithromycin level by 34%) in blood. No dose adjustment currently recommended.
- > Fluconazole Increases efavirenz level by 16% in blood. No dose adjustment currently recommended.
- Food Should be taken on an empty stomach. Increases efavirenz level 10–50% in blood.
- Fosamprenavir May significantly decrease fosamprenavir level in blood. Use only boosted fosamprenavir at 1,400mg + 300mg ritonavir once a day.
- Indinavir

Decreases indinavir level by 31% in blood. Indinavir dose should be increased to 1,000mg 3 times a day or 800mg twice daily if boosted with ritonavir.

Efavirenz plus ... (continued)

- > Kaletra
 - Decreases lopinavir level by about 25% (trough by about 33%). Kaletra dose should be_increased to 4 capsules twice a day.
 - Methadone Decreases methadone level by 52% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
 - > Nelfinavir Increases nelfinavir level by 20% and decreases efavirenz level by 22% in blood. No dose adjustments currently recommended.
 - > Nevirapine Decreases efavirenz level by 22% in blood. Efavirenz dose may need to be increased to 800mg once a day.
 - Rifabutin Decreases rifabutin level in blood. Increase rifabutin dose by 50% if taken daily or double the dose if taken 3 times a week.
- > Rifampin Decreases efavirenz level by 26% in blood. Impact of interaction is uncertain.
- > Ritonavir Should be used together with caution. Increases efavirenz level by 21% and ritonavir level by 18% in blood. Check for side effects.
- Saquinavir (Fortovase) Decreases efavirenz level by 12% and decreases saquinavir level by 62% in blood. Requires ritonavir boosting.
- St. John's Wort Should be used together with great caution. May decrease efavirenz level in blood.
- Simvastatin Decreases simvastatin level in blood. Simvastatin dose may need to be increased.
- Voriconazole Should not be used together. Increases risk of side effects.

Emtricitabine (FTC, Emtriva) plus ...

There are currently no known or suspected interactions between FTC and any other drugs.

Enfuvirtide (T20, Fuzeon) plus ...

There are currently no known or suspected interactions between enfuvirtide and any other drugs, although ritonavir can increase enfuvirtide level in blood somewhat.

Ethinyl estradiol (Ortho-Novum) plus ...

- > Amprenavir Should not be used together. May decrease ethinyl estradiol and norethindrone levels in blood. Use another form of contraception.
- Atazanavir Should be used together with caution. Increases ethinyl estradiol level in blood. Use lowest effective dose of ethinyl estradiol.
- > Delavirdine May increase ethinyl estradiol level in blood. Impact of interaction is uncertain.

Ethinyl estradiol plus ... (continued)

> Fluconazole

Increases ethinyl estradiol level by up to 38% in blood. No dose adjustments currently recommended.

- > Indinavir Should be used together with caution. Increases ethinyl estradiol level by 24% in blood.
- Kaletra Should not be used together. Decreases ethinyl estradiol level by 42% in blood. Use another form of contraception.
- > Nelfinavir Should not be used together. Decreases ethinyl estradiol level by 47% in blood. Use another form of contraception.
- > Nevirapine Should not be used together. Decreases ethinyl estradiol level in blood. Use another form of contraception.
- > Rifabutin Should not be used together. May decrease ethinyl estradiol level in blood. Use another form of contraception.
- Rifampin Should not be used together. May decrease ethinyl estradiol level in blood. Use another form of contraception.
- > Ritonavir
 Should not be used together. Decreases
 think of the based together and the based of the based

ethinyl estradiol level by 40% in blood. Use an another form of contraception.

Ethionamide (Trecator) plus ...

> Alcohol

Excessive alcohol consumption while on ethionamide may cause psychosis.

- Cycloserine May increase risk of central nervous system related toxicity.
-) ddC

May increase risk of peripheral neuropathy.

- Ethambutol May increase risk of side effects from other anti-TB drugs like ethambutol.
- > Isoniazid May increase risk of encephalopathy (dysfunction of brain). May increase isoniazid level in blood.
- > Pyrazinamide May increase risk of side effects from other anti-TB drugs like pyrazinamide.
- Rifabutin May increase risk of side effects from other anti-TB drugs like rifabutin.
- > Rifampin May increase risk of side effects from other anti-TB drugs like rifampin.
- Rifapentine May increase risk of side effects from other anti-TB drugs like rifapentine.

Fluconazole (Diflucan) plus ...

- > Amphotericin B
 - May interfere with the activity of amphotericin B.

Fluconazole plus ... (continued)

> Astemizole

Should not be used together*. Increases risk of side effects.

) AZT

Increases AZT level by 74% and may increase fluconazole level in blood. Check for toxicity.

- > Benzodiazepines May increase benzodiazepine level in blood.
- Cimetidine Decreases fluconazole level by 13% in blood. No dose adjustment recommended.
- Cisapride Should not be used together*. Increases risk of side effects.
- Cyclosporine Should be used together with caution. Increases cyclosporine level by 92% in blood. Check cyclosporine level and serum creatinine.
- Efavirenz Increases efavirenz level by 16% in blood. No dose adjustment currently recommended.
- > Ethinyl estradiol Increases ethinyl estradiol level by up to 38% in blood. No dose adjustments currently recommended.
- Glipizide Should be used together with caution. Increases glipizide level by 49% in blood. Closely check blood sugar for hypoglycemia.
- > Glyburide Should be used together with caution. Increases glyburide level by 44% in blood. Closely check blood sugar for hypoglycemia.
- > Hydrochlorothiazide Increases fluconazole level by 45% in blood.
- > Indinavir
- Decreases indinavir level by 19% in blood.
- > Levonorgestrel

Increases levonorgestrel level by 25% in blood.

- > Loratadine May affect loratadine and/or fluconazole level in blood.
- > Phenytoin Increases phenytoin level by 88% in blood.
- > Ranitidine May decrease fluconazole level in blood.
- > Rifabutin Should be used together with caution. May increase rifabutin level by up to 80% in blood. Increases risk of painful eye inflammation.
- > Rifampin Should be used together with caution. Decreases fluconazole level by 23% in blood. May require higher dose of fluconazole.
- Ritonavir Increases ritonavir level by 12% in blood. No dose adjustment recommended; however, check for liver toxicity.
- Saquinavir (Fortovase) May increase saquinavir level in blood. However, no dose adjustment recommended.
- Saquinavir (Invirase) May increase saquinavir level in blood. However, No dose adjustment recommended.

Fluconazole plus ... (continued)

> Tacrolimus

Should be used together with caution. May increase tacrolimus level in blood and increases risk of kidney toxicity.

- > Theophylline Should be used together with caution. Increases theophylline level by 21% in blood. Check theophylline level.
- > Tolbutamide Should be used together with caution. Increases tolbutamide level by 26% in blood. Closely check blood sugar for hypoglycemia.
- > Trimetrexate May affect trimetrexate and/or fluconazole level in blood.
- > Warfarin Should be used together with caution. Check for increases in prothrombin (coagulation) time.

Flucytosine (Ancobon) plus ...

- > Amphotericin B Increases antifungal activity in test tubes but increases risk of bone marrow and kidney toxicity.
-) AZT
- May increase risk of bone marrow toxicity. > Cvtosine arabinoside
- Should be used with caution. May decrease antifungal activity.
- Ganciclovir Should be used together with great caution. May increase risk of bone marrow toxicity.
- Interferon-alfa May increase risk of bone marrow toxicity.

Fosamprenavir (Lexiva) plus ...

- > Alprazolam Should be used together with caution. May require decrease in alprazolam dose.
- > Amiodarone Should be used together with great caution. May increase serious side effects. Check amiodarone blood concentrations if possible.
- Atorvastatin Fosamprenavir alone and with ritonavir increases atorvastatin level in blood. Lower atorvastatin dose to no more than 20mg/ day or use alternatives.
- > Bepridil Should be used together with great caution. Increases risk of serious side effects.
- Calcium channel blockers Should be used together with great caution. Check for side effects.
- Carbamazepine Should be used together with great caution. May significantly decrease fosamprenavir level in blood.
- Clorazepate Should be used together with caution. May require decrease in clorazepate dose.

Fosamprenavir plus ... (continued)

- **>** Cimetidine
- Should be used together with caution. May decrease fosamprenavir level in blood. No dose adjustment currently recommended.
- Cisapride Should not be used together*. Increases risk of serious side effects.
- Cyclosporine Should be used together with caution. May increase cyclosporine level in blood. Check cyclosporine level if used together.
- > Delavirdine Should not be used together. May significantly decrease fosamprenavir blood level.
- Dexamethazone Should be used together with great caution. May significantly decrease fosamprenavir level in blood.
- > Diazepam Should be used together with caution. May require decrease in diazepam dose.
- > Dihydroergotamine Should not be used together*. Increases risk of side effects.
- Efavirenz May significantly decrease fosamprenavir level in blood. Use only boosted fosamprenavir at 1,400mg + 300mg ritonavir once a day.
- > Ergonovine Should not be used together*. Increases risk of side effects.
- > Ergotamine Should not be used together*. Increases risk of side effects.
- Famotidine Should be used together with caution. May decrease fosamprenavir level in blood. No dose adjustment currently recommended.
- > Flecainide Should not be used together if boosting with ritonavir*. Increases risk of serious side effects.
- Flurazepam Should be used together with caution. May require decrease in flurazepam dose.
- Itraconazole Should be used together with great caution. Increases risk of serious side effects. Do not exceed 400mg of itraconazole if fosamprenavir used alone, or 200mg if boosted with ritonavir.
- Kaletra Significantly decreases fosamprenavir level in blood. Also increases risk of side effects.
- in blood. Also increases risk of side effects. Appropriate dose adjustments not yet established.
- Ketoconazole Should be used together with great caution. Increases risk of serious side effects. Do not exceed 400mg of ketoconazole if fosamprenavir used alone, or 200mg if boosted with ritonavir.
- > Lidocaine (systemic) Should be used together with great caution. May increase serious side effects. Check lidocaine blood concentrations if possible.

DRUG INTERACTIONS . AUGUST 2004 . PROJECT INFORM 1

Fosamprenavir plus ... (continued)

- > Lovastatin Should not be used together*. Increases risk of serious side effects.
- > Methylergonovine Should not be used together*. Increases risk of serious side effects.
- > Midazolam Should not be used together*. Increases risk of serious side effects.
- > Nizatidine Should be used together with caution. May decrease fosamprenavir level in blood. No dose adjustment currently recommended.
- > Phenobarbital Should be used together with great caution.
- > Phenytoin
- Should be used together with great caution. > Pimozide
- Should not be used together*. Increases risk of serious side effects.
- > Propafenone Should not be used together if boosting with ritonavir*. Increases risk of serious side effects.
- > Proton pump inhibitors Should be used together with great caution. May significantly decrease fosamprenavir level in blood.
- **)** Quinidine Should be used together with great caution. May increase serious side effects. Check quinidine blood concentrations if possible.
- > Ranitidine Should be used together with caution. Decreases fosamprenavir level by 51% in blood. No dose adjustment currently recommended.
- > Rapamycin Should be used together with caution. May increase rapamycin level in blood. Check rapamycin level if used together.
- > Rifabutin

Should be used together with great caution. Increases risk of neutropenia. Check for side effects and decrease rifabutin dose by at least 50% if fosamprenavir used alone or by 75% if boosted with ritonavir.

- > Rifampin Should not be used together*. May decrease anti-HIV effect of fosamprenavir.
- > Ritonavir Increases fosamprenavir level in blood. Adjust dose to fosamprenavir 700mg + ritonavir 100mg twice a day or fosamprenavir 1,400mg + ritonavir 200mg once a day.
- Sildenafil Should be used together with great caution. May increase risk of side effects. Decrease sildenafil dose to 25mg every 48 hours and check for side effects.
- **)** Simvastatin Should not be used together*. Increases risk of serious side effects.
- > St. John's Wort Should not be used together. May significantly decrease fosamprenavir level in blood.

Fosamprenavir plus ... (continued)

- **>** Tacrolimus Should be used together with caution. May increase tacrolimus level in blood. Check tacrolimus level if used together.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > Triazolam Should not be used together*. Increases risk of serious side effects.
- > Tricyclic antidepressants Should be used together with great caution. May increase risk of side effects. Check tricyclics level if used together.
- Vardenafil Should be used together with great caution. May increase risk of side effects. Decrease vardenafil dose to 2.5mg/24 hours (if fosamprenavir used alone) or 2.5mg/72 hours (if boosted with ritonavir). Check for side effects.
- > Warfarin Should be used together with great caution. May increase risk of serious side effects.

Foscarnet (Foscavir) plus ...

- > Adefovir Should not be used together.* Increases risk of kidney toxicity.
- > Amikacin Should not be used together*. Increases risk of kidney toxicity.
- > Amphotericin B Should be used together with great caution. May decrease number of red blood cells and increases risk of kidney toxicity.
- > AZT

Should be used together with caution. May increase risk of anemia.

- > Cidofovir Should not be used together*. Increases risk of kidney toxicity.
- > ddC May increase risk of kidney toxicity and peripheral neuropathy. Check for toxicity.
- **)** Gentamicin Should not be used together*. Increases risk of kidney toxicity.
- > Kanamycin Should not be used together*. Increases risk of kidney toxicity.
- > Pentamidine (IV) Should be used with caution. May increase risk of kidney toxicity. Check closely.
- > Ritonavir Should be used with caution. Abnormal kidney function has been observed.
- > Streptomycin Should be used together with great caution. Increases risk of kidney toxicity.
- > Tenofovir Should be used together with great caution. May increase risk of kidney toxicity.

Foscarnet plus ... (continued)

> Tobramycin

Should not be used together*. Increases risk of kidney toxicity.

Ganciclovir (Cytovene, oral/IV) plus ...

> Acyclovir

People allergic to acyclovir should not take ganciclovir.

> Adriamycin

Should be used together with great caution. May increase risk of bone marrow toxicity.

- > Amphotericin B Should be used together with great caution. May increase risk of bone marrow toxicity.
- > Anticancer drugs May increase risk of bone marrow toxicity.
- > AZT

Increases AZT level in blood and may increase risk of neutropenia, anemia and bone marrow toxicity. Avoid combination or lower AZT dose.

- > Dapsone
- May increase risk of bone marrow toxicity. ddl or ddl EC
 - Should be used with caution. Significantly increases ddI level in blood. May increase risk of pancreatitis.
- **)** d4T

May increase risk of pancreatitis.

- > Food
 - Increases oral ganciclovir level in blood.
- > Flucytosine Should be used together with great caution. May increase risk of bone marrow toxicity.
- > Imipenem-cilastin

May increase risk of seizures. Check closely.

> Interferon-alfa May increase risk of bone marrow toxicity.

> Pentamidine Should be used together with great caution. May increase risk of bone marrow toxicity.

- > Probenecid Should be used together with caution. May increase ganciclovir level in blood and decrease ganciclovir clearance.
- > Tenofovir

Should be used together with great caution. May increase risk of side effects. Check for kidney toxicity.

- > TMP/SMX Should be used together with great caution. May increase risk of bone marrow toxicity.
- > Vinblastine Should be used together with great caution. May increase risk of bone marrow toxicity.
- > Vincristine Should be used together with great caution. May increase risk of bone marrow toxicity.

Indinavir (Crixivan) plus ...

> Amlodipine

Should be used together with great caution. May increase amlodipine level in blood.

- > Amprenavir Increases amprenavir level by 33% and decreases indinavir level by 38% in blood. No dose adjustment recommended.
- > Astemizole Should not be used together*. Increases risk of side effects.
- > Atorvastatin Should be used together with great caution. May increase atorvastatin level in blood.
- > Atazanavir Should not be used together*. Increases risk of side effects.
-) AZT

Increases AZT level 17–36% in blood. No dose adjustment recommended.

- Benzodiazepines May increase benzodiazepine level in blood.
- Bepridil Should be used together with great caution. May increase bepridil level in blood.
- Cisapride Should not be used together*. Increases risk of side effects.
- Clarithromycin Increases clarithromycin level by 53% and indinavir level by 29% in blood. No dose adjustment recommended.
-) ddl

May decrease ddI and indinavir levels in blood. Indinavir should be taken 1 hour before or 2 hours after taking ddI.

- Delavirdine Increases indinavir level 50–100% in blood. Indinavir may be reduced to 600mg 3 times a day.
- > Dihydroergotamine Should not be used together*. Increases risk of side effects.
- > Diltiazem Should be used together with great caution. May increase diltiazem level in blood.
- Efavirenz Decreases indinavir level by 31% in blood. Indinavir dose should be increased to 1,000mg 3 times a day or 800mg twice daily if boosted with ritonavir.
- > Ergonovine Should not be used together*. Increases risk of side effects.
- > Ergotamine Should not be used together*. Increases risk of side effects.
- > Ethinyl estradiol Should be used together with caution. Increases ethinyl estradiol level by 24% in blood.
- Fatty foods Decreases indinavir level in blood. Indinavir should be taken 1 hour before eating or 2 hours after eating a large or fatty meal. Indinavir may be taken with a small snack such as juice and dry toast.

Indinavir plus ... (continued)

- > Felodipine
- Should be used together with great caution. May increase felodipine level in blood.
- > Fluconazole Decreases indinavir level by 19% in blood.
- Fluvastatin Should be used together with great caution. May increase fluvastatin level in blood.
- Grapefruit juice Decreases indinavir level by 26% in blood.
- Isradipine
 Should be used together with great caution.
 May increase isradipine level in blood.
- Itraconazole May affect indinavir and/or itraconazole level in blood. Decrease dose of indinavir to 600mg 3 times a day when combined with itraconazole 200mg twice daily.
- > Kaletra
 - Increases indinavir trough level by about 4fold in blood. Currently under study; however, indinavir dose should be reduced to no more than 600mg twice a day if used together.
- > Ketoconazole Increases indinavir level by 68% in blood. Decrease indinavir dose to 600mg 3 times a day.
- > Lovastatin Should not be used together*. Increases risk of side effects.
- > Methylergonovine Should not be used together*. Increases risk of side effects.
- > Midazolam Should not be used together*. Increases risk of side effects.
- > Nelfinavir

Increases indinavir level by 51% and increases nelfinavir level by 83% in blood. There are limited data to support a dose of indinavir 1,200mg + nelfinavir 1,250mg twice a day.

- > Nevirapine Decreases indinavir level by 27% in blood. Indinavir dose should be increased to 1,000mg 3 times a day or 800mg twice daily if boosted with ritonavir.
- Nicardipine
 Should be used together with great caution. May increase nicardipine level in blood.
- > Nifedipine
 - Should be used together with great caution. May increase nifedipine level in blood.
- > Nimodipine Should be used together with great caution. May increase nimodipine level in blood.
- > Nisoldipine
 Should be used together with great caution. May increase nisoldipine level in blood.
- > Nitrendipine
 Should be used together with great caution. May increase nitrendipine level in blood.
- > Norethindrone

Should be used together with caution. Increases norethindrone level in blood by 14%.

Indinavir plus ... (continued)

- > Quinidine
- Increases indinavir level by 10% in blood.
- Rifabutin Increases rifabutin level by 204% and decreases indinavir level by 32% in blood. Decrease dose of rifabutin by half the standard dose and increase indinavir dose to 1,000mg 3 times a day.
- > Rifampin Should not be used together*. May increase rifampin level in blood.
- > Rifampicin
 Should not be used together. Decreases indinavir level by 80% in blood, even when boosted with ritonavir.
- Ritonavir Significantly increases indinavir level in blood. Adjust dose to indinavir 800mg + ritonavir 200mg twice a day or indinavir 400mg + ritonavir 400mg twice a day.
- Saquinavir (Fortovase) Increases saquinavir level 3.6–6.2 times in blood.
- Saquinavir (Invirase) Increases saquinavir level 4.6–7.2 times in blood.
- > Sildenafil Should be used with caution. Increases indinavir and sildenafil levels in blood.
- Simvastatin Should not be used together*. Increases risk of side effects.
- St. John's Wort Should not be used together. Decreases indinavir level by 57% in blood.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > TMP/SMX Should be used together with caution. Increases trimethoprim level by 31% in blood.
- > Triazolam Should not be used together*. Increases risk of side effects.
- Vardenafil Should be used together with caution. Increases vardenafil level in blood. Dose of vardenafil should not exceed 2.5mg/day.
- > Verapamil Should be used together with great caution. May increase verapamil level in blood.

Interferon-alfa (Intron-A, Roferon-A) plus ...

- > Anticancer drugs
- May increase risk of bone marrow toxicity. > AZT
- Increases anti-HIV activity in test tubes and may increase risk of bone marrow toxicity. Lower AZT dose by 50–75%.
- > Flucytosine
- May increase risk of bone marrow toxicity. > Ganciclovir
- May increase risk of bone marrow toxicity.

Interferon-alfa plus ... (continued)

 > Pentamidine May increase risk of bone marrow toxicity.
 > Pyrimethamine

May increase risk of bone marrow toxicity.

Isoniazid (INH) plus ...

- > Acetaminophen Should not be used together*. Increases risk of side effects.
- > Alcohol Daily or heavy alcohol use may increase risk of isoniazid-associated hepatitis.
- Carbamazepine Should be used together with great caution. Increases carbamazepine level in blood. Check for toxicity and adjust carbamazepine dose accordingly.
- Corticosteroids Increases isoniazid metabolism and decreases isoniazid level in blood. Isoniazid dose may require adjustment.
- Cycloserine May increase risk of central nervous system toxicity.
-) ddC

May increase risk of peripheral neuropathy.

- Ethionamide May increase risk of encephalopathy (dysfunction of the brain) and may increase isoniazid level in blood.
- Food Should not be used together. Decreases isoniazid level in blood.
- Itraconazole Should be used together with great caution. May decrease itraconazole level in blood.
- > Ketoconazole Should not be used together*. Decreases ketoconazole level in blood.
- > Phenytoin Should be used together with caution. Increases phenytoin level in blood. Check for toxicity and adjust dose of phenytoin accordingly.
- Rifampin Should be used together with caution. May increase risk of liver toxicity.
- Sulfonylureas May increase risk of high blood sugar level.
- > Theophylline Should be used together with great caution. Increases level of theophylline in blood. Check theophylline level closely.

Itraconazole (Sporanox) plus ...

- Alprazolam Should be used together with caution. May increase level of alprazolam in blood. Check for side effects.
- > Amphotericin B May interfere with activity of amphotericin B.

Itraconazole plus ... (continued)

- > Amprenavir May increase itraconazole and amprenavir levels in blood. Check for side effects.
- Antacids
 Decreases itraconazole level in blood. Should
 be taken 2 hours apart.

 Astemizole
- Should not be used together*. Increases risk of side effects.
- > Atorvastatin Should not be used together*. Increases risk of side effects.
- > Busulfan Should be used together with caution. Increases busulfan level in blood. Check for toxicity.
- Carbamazepine Should be used together with caution. Increases carbamazepine level and decreases itraconazole level in blood. Check for side effects.
- Cimetidine Decreases itraconazole level in blood. May need to be taken at least 2 hours apart.
- Cisapride Should not be used together*. Increases risk of side effects.
- Cola beverage Should be used together with caution. Increases itraconazole level 75% in blood.
- Cyclosporine Should be used together with caution. Increases cyclosporine level in blood and may increase risk of side effects.
- > ddl Needs to be taken 2 hours apart, otherwise ddI buffering agents may significantly decrease itraconazole level in blood.
- > Delavirdine May increase delavirdine level in blood.
- Diazepam Should be used together with caution. May increase level of diazepam in blood. Check for side effects.
- Digoxin Should be used together with caution. Increases digoxin level in blood. Check for side effects.
- > Dofetilide Should not be used together*. Increases dofetilide level in blood.
- Felodipine Should be used together with great caution. Increases risk of felodipine side effects. Check closely.
- Fluvastatin Should not be used together*. Increases risk of side effects.
- Food May increase itraconazole level in blood.
- Fosamprenavir Should be used together with great caution. Increases risk of serious side effects. Do not exceed 400mg of itraconazole if fosamprenavir used alone, or 200mg if boosted with ritonavir.

Itraconazole plus ... (continued)

> Glipizide

May increase risk of low blood sugar level.

- > Glyburide
 - May increase risk of low blood sugar level. > Indinavir
 - May affect itraconazole and/or indinavir level in blood. Reduce dose of indinavir to 600mg 3 times a day when combined with itraconazole 200mg twice daily.
 - > Isoniazid

Should be used together with great caution. May decrease itraconazole level in blood.

> Kaletra

Increases itraconazole level in blood. Dose of itraconazole should not exceed 200mg/ day.

- > Lovastatin Should not be used together*. Increases risk of side effects.
- Midazolam (oral) Should not be used together*. Increases risk of side effects.
- > Nevirapine Should be used together with caution. May decrease itraconazole level in blood. Itraconazole dose may need to be increased.
- > Nicardipine Should be used together with great caution. Increases risk of nicardipine side effects. Check closely.
- > Nifedipine Should be used together with great caution. Increases risk of nifedipine side effects. Check closely.

> Nimodipine Should be used together with great caution. Increases risk of nimodipine side effects. Check closely.

- > Nisoldipine Should be used together with great caution. Increases risk of nisoldipine side effects. Check closely.
- > Nitrendipine Should be used together with great caution. Increases risk of nitrendipine side effects. Check closely.
- > Omeprazole Decreases itraconazole level in blood.
- > Phenobarbital Should be used together with caution. Decreases itraconazole level and may increase phenobarbital level in blood.
- > Phenytoin Should be used together with caution. Decreases itraconazole level and may increase phenytoin level in blood.
- > Pimozide Should not be used together*. Increases risk of side effects.
- Quinidine Should not be used together*. Increases risk of side effects.
- > Ranitidine

Decreases itraconazole level in blood.

Itraconazole plus ... (continued)

- > Rifabutin Should not be used together. May increase rifabutin level and may decrease itraconazole level in blood.
- Rifampin Should not be used together. Decreases rifampin level and may decrease itraconazole level in blood.
- Ritonavir Should be used together with caution. May significantly increase itraconazole level in blood.
- > Saquinavir (Fortovase) Increases saquinavir level in blood. However, no dose adjustment recommended.
- Saquinavir (Invirase) Increases saquinavir level in blood. However, no dose adjustment recommended.
- Simvastatin Should not be used together*. Increases risk of side effects.
- > Sirolimus Should be used together with caution. Increases sirolimus level in blood.
- Tacrolimus Should be used together with caution. Increases tacrolimus level in blood.
- > Tedalafil Should be used together with caution. Increases tedalafil level in blood. Tedalafil dose should not exceed 10mg in a 72 hour period.
- Testosterone May decrease testosterone level in blood.
- > Tolbutamide May increase risk of low blood sugar level.
- Trazodone Increases trazodone level in blood. Trazodone doe may need to be reduced.
- > Triazolam Should not be used together*. Increases risk of side effects.
- > Trimetrexate May affect trimetrexate and/or itraconazole level in blood.
- Vardenafil Should be used together with caution. Increases vardenafil level in blood. If taking 400mg itraconazole per day, dose of vardenafil should not exceed 2.5mg/day. If taking 200mg itraconazole per day, dose of vardenafil should not exceed 5mg/day.
- > Verapamil Should be used together with great caution. Increases risk of verapamil side effects. Check closely.
- > Vinblastine Should be used together with caution. Increases vinblastine level in blood. Check for side effects.
- > Vincristine Should be used together with caution. Increases vincristine level in blood. Check for side effects.

Itraconazole plus ... (continued)

- **)** Warfarin
 - Should be used together with caution. May increase prothrombin (coagulation) time and increase anticoagulant effects. Check warfarin dose closely.

Kaletra (lopinavir/ritonavir) plus ...

Kaletra is a single pill containing 2 anti-HIV drugs, lopinavir and ritonavir. References below indicate which drug contained in Kaletra is specifically altered in combinations.

> Amiodarone

May increase amiodarone level in blood. Checking amiodarone level in blood is recommended.

- > Amprenavir Decreases lopinavir level by 15% and increases amprenavir trough level about 2-fold in blood. Amprenavir dose may be reduced to 750mg twice a day or less. However, optimal dose is unclear.
- > Astemizole Should not be used together. Increases risk of side effects.
- Atorvastatin Should be used together with great caution. Increases active atorvastatin level about 6fold and atorvastatin metabolite by 2.5 times in blood and May increase risk of side effects.
- > Atovaquone Decreases atovaquone level in blood. Atovaquone dose may have to be increased.
- > Bepridil May increase bepridil level in blood. Checking bepridil level in blood is recommended.
- Carbamazepine Should be used together with great caution. May decrease lopinavir level in blood.
- Cisapride Should not be used together. Increases risk of side effects.
- Clarithromycin Increases clarithromycin level in blood. Clarithromycin dose should be decreased for people with kidney dysfunction.
- Cyclosporine May increase cyclosporine level in blood. Checking cyclosporine level in blood is recommended.
- > ddl or ddl EC ddI or ddl EC should be taken 1 hour before or 2 hours after taking Kaletra. ddI and ddI EC should be taken on an empty stomach, and Kaletra should be taken with a meal.
- > Delavirdine May increase lopinavir level in blood. Needs further study.
- Dexamethasone Should be used together with great caution. May decrease lopinavir level in blood.
- > Dihydroergotamine Should not be used together. Increases risk of side effects.
- > Disulfiram/metronidazole Should be used together with Kaletra oral solution with caution. Alcohol in Kaletra oral solution may increase risk of side effects.

Kaletra plus ... (continued)

- Efavirenz
- Decreases lopinavir level by about 25% (trough by about 33%). Kaletra dose should be increased to 4 capsules twice a day.
- Frgonovine Should not be used together. Increases risk of side effects.
- > Ergotamine Should not be used together. Increases risk
- of side effects.
 > Ethinyl estradiol Should not be used together. Decreases ethinyl estradiol level by 42% in blood. Use another form of contraception.
- > Felodipine Should be used together with caution. May increase felodipine level in blood. Check for side effects.
- > Food Increases lopinavir level in blood. Kaletra should be taken with food.
- > Fosamprenavir Significantly decreases fosamprenavir level in blood. Also increases risk of side effects. Appropriate dose adjustments not yet established.
- Indinavir Increases indinavir trough level by about 4 times in blood. Currently under study; however, indinavir dose should be reduced to no more than 600mg twice a day if used together.
- Itraconazole Increases itraconazole level in blood. Itraconazole dose should not exceed 200mg/day.
- > Ketoconazole Increases ketoconazole level by 3 times in blood. Ketoconazole dose should not exceed 200mg/day.
- Lidocaine (systemic) May increase lidocaine level in blood. Checking lidocaine level in blood is recommended.
- > Lovastatin Should not be used together. Increases risk of side effects.
- > Methadone Decreases methadone level by 53% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Methylergonovine Should not be used together. Increases risk of side effects.
- > Midazolam Should not be used together. Increases risk of side effects.
- > Nelfinavir Decreases lopinavir level in blood. Kaletra dose may need to be increased. Optimal dose is under study.
- > Nevirapine Decreases lopinavir level in blood. Kaletra dose should be increased to 4 capsules twice a day.
- > Nicardipine
- Should be used together with caution. May increase nicardipine level in blood. Check for side effects.

Kaletra plus ... (continued)

> Nifedipine Should be used together with caution. May increase nifedipine level in blood. Check for side effects.

- > Phenobarbital Should be used together with great caution. May decrease lopinavir level in blood.
- > Phenytoin Should be used together with great caution. May decrease lopinavir level in blood.
- > Pimozide Should not be used together. Increases risk of side effects.
- > Pravastatin Increases pravastatin level in blood. No dose adjustment recommended.
- > Quinidine May increase quinidine level in blood. Checking for quinidine level in blood is recommended.
- Rapamycin May increase rapamycin level in blood. Checking for rapamycin level in blood is recommended.
- > Rifabutin Increases rifabutin level by 5.7-fold. Rifabutin dose should be changed to 150mg every other day or 3 times a week. Check for side rifabutin effects.
- > Rifampin Should not be used together. Decreases lopinavir level by 75% in blood.
- > Saquinavir (Fortovase) Increases saquinavir trough level by about 3-fold. Saquinavir dose may be reduced to 800mg twice a day. Optimal dose is under study.
- Sildenafil Should be used together with caution. Increases sildenafil level in blood. Sildenafil dose should be reduced to no more than 25mg every 48 hours.
- Simvastatin Should not be used together. Increases risk of side effects.
- St. John's Wort Should not be used together. May decrease lopinavir level in blood.
- > Tacrolimus May increase tacrolimus level in blood. Checking for tacrolimus level in blood is recommended.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > Tenofovir Increases tenofovir level by about 30% and decreases lopinavir level 15% in blood. No dose adjustment currently recommended.
- Triazolam Should not be used together. Increases risk of side effects.
- Vardenafil Should be used together with caution. May increase vardenafil level in blood.

Kaletra plus ... (continued)

- > Warfarin
- May affect warfarin level in blood. Check INR.

Ketoconazole (Nizoral) plus ...

- > Alcohol May increase risk of nausea, vomiting and low blood pressure.
- Alprazolam Should be used together with caution. May increase level of alprazolam in blood.
- > Amphotericin B May interfere with the activity of amphotericin B.
- > Amprenavir Increases amprenavir level by 31% and increases ketoconazole level by 44% in blood. Impact of interaction is uncertain.
- Antacids Should be taken 2 hours apart, otherwise may decrease ketoconazole level in blood.
- Astemizole Should not be used together*. Increases risk of side effects.
- Cimetidine
 Should be taken 2 hours apart, otherwise
 may decrease ketoconazole level in blood.
- Cisapride Should not be used together*. Increases risk of side effects.
- Clonazepam Should be used with caution. May increase clonazepam level in blood.
- Corticosteroids May decrease corticosteroid level in blood.
- > Cyclosporine
- Should be used together with caution. May increase cyclosporine level in blood. Check cyclosporine level and adjust dose accordingly.
- > ddl Should be taken 2 hours apart, otherwise ddI buffering agents may significantly decrease ketoconazole level in blood.
- Delavirdine Increases delavirdine level by 50% in blood.
- > Diazepam Should be used together with caution. May increase diazepam level in blood.
- > Digoxin May increase digoxin level in blood. Digoxin level should be carefully checked.
- Famotidine Should be taken 2 hours apart, otherwise may decrease ketoconazole level in blood.
- Fosamprenavir Should be used together with great caution. Increases risk of serious side effects. Do not exceed 400mg of ketoconazole if fosamprenavir used alone, or 200mg if boosted with ritonavir.
- Glipizide May increase risk of low blood sugar level.

Ketoconazole plus ... (continued)

- > Glyburide
- May increase risk of low blood sugar level.
- > Indinavir
- Increases indinavir level by 68% in blood. Decrease indinavir dose to 600mg 3 times/day.
- Isoniazid Should not be used together*. Decreases ketoconazole level in blood.
- > Kaletra
 - Increases ketoconazole level by 3 times in blood. Ketoconazole dose should not exceed 200mg/day.
- > Loratadine

Increases loratadine level by 300% in blood.

> Methylprednisolone

Should be used together with caution. May increase methylprednisolone level in blood. Check blood level of cyclosporine and adjust dose accordingly.

> Miconazole

Should be used together with great caution. May lead to severe hypoglycemia (low blood sugar).

- > Midazolam Should not be used together*. Increases midazolam level in blood.
- > Nelfinavir Increases nelfinavir level by 35% in blood. No dose adjustment recommended.
- > Nevirapine Should not be used together. Significantly decreases ketoconazole level in blood.
- > Omeprazole Should be used together with caution. Decreases ketoconazole and increases omeprazole level in blood.
- > Phenytoin Should be used together with caution. May affect ketoconazole and phenytoin levels in blood. Check phenytoin and ketocon-

azole levels and adjust dose accordingly.

> Prednisolone May increase prednisolone level in blood.

- Ranitidine Should be taken 2 hours apart, otherwise may decrease ketoconazole level in blood.
- > Rifabutin May decrease ketoconazole level in blood.
- Rifampin Should not be used together*. Significantly decreases ketoconazole level in blood.
- Ritonavir Should be used together with caution. Significantly increases ketoconazole level in blood. Ketoconazole dose should not exceed 200mg/day.
- Saquinavir (Fortovase) Significantly increases saquinavir level in blood. However, no dose adjustment recommended.
- > Saquinavir (Invirase) Significantly increases saquinavir level in blood. However, no dose adjustment recommended.

Ketoconazole plus ... (continued)

> Tacrolimus

Should be used together with caution. May increase tacrolimus level in blood. Check tacrolimus level.

- > Tedalafil Should be used together with caution. Increases tedalafil level in blood. Tedalafil dose should not exceed 10mg in a 72 hour period.
- > Testosterone May decrease testosterone level in blood, especially when ketoconazole is used at higher doses.
- > Tolbutamide May increase risk of low blood sugar level.
- > Trazodone Increases trazodone level in blood. Trazodone dose may need to be reduced.
- Triazolam Should not be used together*. Increases triazolam level in blood.
- > Trimetrexate May affect trimetrexate and/or ketoconazole level in blood.
- > Vardenafil

Should be used together with caution. Increases vardenafil level in blood. If taking ketoconazole 400mg/day, vardenafil dose should not exceed 2.5mg/day. If taking ketoconazole 200mg/day, vardenafil dose should not exceed 5mg/day.

> Warfarin Should be used together with caution. May increase prothrombin (coagulation) time and increase anticoagulant effects. Check warfarin dose closely.

Lamivudine (3TC, Epivir) plus ...

> Abacavir

Decreases 3TC level by 15% in blood. No dose adjustment recommended.

- > ddC Should not be used together. Decreases anti-HIV activity.
- > Nelfinavir Increases 3TC level by 10% in blood. No dose adjustment recommended.
- > TMP/SMX Increases 3TC level by 44% in blood. No dose adjustment recommended. Check for 3TC side effects.

Methadone plus ...

- > Abacavir May decrease methadone level in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
-) AZT

Increases AZT level by 43% in blood. Check for AZT side effects.

- > ddl or ddl EC Decreases ddI level by 60% in blood.
- > Efavirenz

Decreases methadone level by 52% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.

Methadone plus ... (continued)

> Kaletra

Decreases methadone level by 53% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.

- > Nelfinavir Decreases methadone level in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Nevirapine Decreases methadone level by about 60% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Rifabutin May decrease effectiveness of methadone. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Rifampin May decrease effectiveness of methadone. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- Ritonavir Decreases methadone level by about 40% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.

Nelfinavir (Viracept) plus ...

> 3TC

Increases 3TC level by 10% in blood. No dose adjustment recommended.

- > Amiodarone Should not be used together*. Increases risk of side effects.
- > Amprenavir Increases nelfinavir level by 15% in blood. No dose adjustment recommended.
- > Astemizole

Should not be used together*. Increases risk of side effects.

- > Atorvastatin Should be used together with great caution. May increase atorvastatin level in blood.
- > Azithromycin Should be used together with caution. Increases azithromycin level in blood. No dose adjustment recommended, but check
- for hearing and liver side effects.> AZT Decreases AZT level by 35% in blood. No
- dose adjustment recommended.
 Carbamazepine Should be used together with caution. May decrease nelfinavir level in blood.
- Cisapride
 Should not be used together*. Increases risk of side effects.
- > ddl or ddl EC Nelfinavir should be taken 2 hours before or after taking ddI or ddI EC. Nelfinavir should be taken with food, and ddI and ddI EC should only be taken on an empty stomach.

Nelfinavir plus ... (continued)

- > Dihydroergotamine
- Should not be used together*. Increases risk of side effects.
- > Delavirdine

Increases nelfinavir level by about 100% but decreases nelfinavir active metabolite by about 45% in blood. Decreases delavirdine level by about 40%. No data exist to guide dose adjustments.

- > Ergonovine Should not be used together*. Increases risk of side effects.
- > Ergotamine

Should not be used together*. Increases risk of side effects.

- Ethinyl estradiol Should not be used together. Decreases ethinyl estradiol level by 47% in blood. Use another form of contraception.
- > Fluvastatin May interact somewhat with nelfinavir. No

dose adjustment currently recommended.

- Food Increases nelfinavir level in blood. Nelfinavir should be taken with food.
- Indinavir
 Indinavir
 Increases indinavir level by 51% and increases nelfinavir level by 83% in blood.

increases nelfinavir level by 83% in blood. Increase indinavir dose to 1,000mg every 8 hours or boost indinavir with ritonavir.

- Kaletra Decreases lopinavir level in blood. Kaletra dose may need to be increased. Optimal dose is under study.
- > Ketoconazole Increases nelfinavir level by 35% in blood. No dose adjustment recommended.
- > Lovastatin Should not be used together. Increases lovastatin level in blood.
- > Methadone Decreases methadone level in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Methylergonovine Should not be used together*. Increases risk of side effects.
- > Midazolam Should not be used together*. Increases risk of side effects.
- > Nevirapine Modestly decreases nelfinavir level in blood. No dose adjustment recommended.
- > Norethindrone Should be used together with caution. Decreases norethindrone level by 18% in blood.
- > Phenobarbital Should be used together with caution. May decrease level of nelfinavir in blood.
- > Phenytoin Should be used together with caution. May increase rate of metabolism of phenytoin. Check phenytoin level and adjust dose accordingly.

DRUG INTERACTIONS . AUGUST 2004 . PROJECT INFORM

16

Nelfinavir plus ... (continued)

> Pimozide Should not be used together*. Increases risk of side effects.

> Quinidine Should not be us

Should not be used together*. Increases risk of side effects.

- Rifabutin Should be used together with caution. Increases rifabutin and decreases nevirapine level in blood. Reduce rifabutin dose by one-half and use nevirapine at 1,250mg twice a day.
- > Rifampin Should not be used together*. Decreases nelfinavir level by 82% in blood.
- Ritonavir Increases ritonavir level by 9% and increases nelfinavir level by 152% in blood. Alternate doses under study include nelfinavir at 500–750mg + ritonavir 400mg twice a day.
- > Saquinavir (Fortovase) Increases nelfinavir level 18% and increases saquinavir level 4 times in blood. Saquinavir dose may be reduced to 800mg 3 times a day or 1,200mg twice a day.
- Saquinavir (Invirase) Increases saquinavir level by 4 times and increases nelfinavir level by 18% in blood. Saquinavir dose may be reduced to 800mg 3 times a day or 1,200mg twice a day.
- Sildenafil Should be used together with caution. Increases sildenafil level in blood. Sildenafil dose should be reduced to no more than 25mg every 48 hours.
- Simvastatin Should not be used together. Increases simvastatin level in blood.
- St. John's Wort Should not be used together. May decrease nelfinavir level in blood.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- > Triazolam Should not be used together*. Increases risk of side effects.
- Vardenafil Should be used together with caution. May increase vardenafil level in blood.

Nevirapine (Viramune) plus ...

- > Amiodarone Should be used together with caution. May decrease amiodarone level in blood.
- Amoxicillin May increase risk of rashes and Stevens Johnson Syndrome. Requires careful monitoring.
- > Amprenavir May decrease amprenavir level in blood. If boosting amprenavir with ritonavir, increase ritonavir to 200mg twice a day.

Nevirapine plus ... (continued)

- Carbamazepine
 Should be used together with caution. May decrease carbamazepine level in blood.
- Cimetidine Should be used together with caution. May slightly increase nevirapine level in blood.
- Cisapride Should be used together with caution. May decrease cisapride level in blood.
- Clarithromycin Decreases clarithromycin level by 30% and increases nevirapine level by 26% in blood. Consider using alternatives to clarithromycin.
- Clonazepam Should be used together with caution. May decrease clonazepam level in blood.
- > Cyclophosphamide Should be used together with caution. May decrease cyclophosphamide level in blood.
- Cyclosporin Should be used together with caution. May decrease cyclosporin level in blood.
- > Dihydroergotamine Should be used together with caution. May decrease dihydroergotamine level in blood.
- > Dicumarol Should not be used together*. May increase dicumarol level in blood.
- > Diltiazem Should be used together with caution. May decrease diltiazem level in blood.
- > Disopyramide Should be used together with caution. May decrease disopyramide level in blood.
- > Efavirenz Decreases efavirenz level by 22% in blood. Efavirenz dose may need to be increased to 800mg once a day.
- > Ergonovine Should be used together with caution. May decrease ergonovine level in blood.
- > Ergotamine Should be used together with caution. May decrease ergotamine level in blood.
- > Erythromycin Should be used together with great caution. May increase risk of liver toxicity.
- Ethinyl estradiol Should not be used together. Decreases ethinyl estradiol level in blood. Use another form of contraception.
- > Ethosuximide Should be used together with caution. May decrease ethosuximide level in blood.
- Fentanyl Should be used together with caution. May decrease fentanyl level in blood.
- Indinavir Decreases indin
- Decreases indinavir level by 27% in blood. Indinavir dose should be increased to 1,000mg 3 times a day or 800mg twice daily if boosted with ritonavir.

Nevirapine plus ... (continued)

> Itraconazole

Should be used together with caution. May decrease itraconazole level in blood. Itraconazole dose may need to be increased.

- Kaletra Decreases Kaletra level in blood. Kaletra dose should be increased to 4 capsules twice a day.
- Ketoconazole Should not be used together. Significantly decreases ketoconazole level in blood.
- > Lidocaine (systemic) Should be used together with caution. May decrease lidocaine level in blood.
- > Methadone Decreases methadone level by about 60% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Methylergonovine Should be used together with caution. May decrease methylergonovine level in blood.
- > Nelfinavir
 Modestly decreases nelfinavir level in blood. No dose adjustment recommended.
- > Nifedipine Should be used together with caution. May decrease nifedipine level in blood.
- > Norethindrone Decreases norethindrone level in blood. Use another form of contraception.
- > Prednisone May increase risk of ray
- May increase risk of rash.
- > Rifabutin Should be used together with caution. May significantly affect rifabutin level in blood.
- Rifampin Should not be used together. Decreases nevirapine level in blood.
- Ritonavir Decreases ritonavir level by 11% in blood. No dose adjustment recommended.
- Saquinavir (Fortovase) Decreases saquinavir level by 24% in blood. Saquinavir may require ritonavir boosting.
- Saquinavir (Invirase) Decreases saquinavir level by 27% in blood. Saquinavir may require ritonavir boosting.
- Sirolimus Should be used together with caution. May decrease sirolimus level in blood.
- > Steroids
- May increase rate of metabolism of steroids. > St. John's Wort
- Should not be used together. May decrease nevirapine level in blood.
- > Tacrolimus Should be used together with caution. May decrease tacrolimus level in blood.
- TMP/SMX Should not be initiated together for the first 4-6 weeks. May increase risk of rashes.

Nevirapine plus ... (continued)

> Voriconazole

Should be used together with caution. May increase nevirapine side effects and decrease effectiveness of voriconazole.

> Warfarin Should be used together with great caution. Interaction is complex. Warfarin level in blood should be checked carefully.

Oral contraceptives

See section on Ethinyl estradiol.

Pentamidine (Pentam) plus ...

- > Adefovir
- Should not be used together*. Increases risk of kidney toxicity.
- > Amphotericin B Should be used together with great caution. May increase risk of kidney toxicity.
- > Anticancer drugs May increase risk of anemia.
-) AZT
 - May increase risk of anemia.
- > Cidofovir Should not be used together*. Increases risk of kidney toxicity.

> ddC Should be used with great caution. May increase risk of peripheral neuropathy and pancreatitis.

- > ddl or ddl EC May increase risk of pancreatitis.
- **)** d4T
 - May increase risk of pancreatitis.
- > Foscarnet Should be used with caution. May increase risk of kidney toxicity. Check closely.
- > Ganciclovir Should be used together with great caution. May increase risk of bone marrow toxicity.
- > Interferon-alfa May increase risk of bone marrow toxicity.
- > Tenofovir Should not be used together*. May increase risk of side effects.

Probenecid (Benemid, ColBenemid) plus ...

- > Acetaminophen Should be used together with caution. Increases acetaminophen level in blood.
- > Acyclovir Increases acyclovir level in blood. Impact of interaction is uncertain.
- > Aspirin Should be used together with caution. Decreases activity of probenecid.
-) AZT

Should be used together with caution. Increases AZT level by 106% in blood. Check for AZT side effects.

> Benzodiazepines Increases benzodiazepine level in blood.

Probenecid plus ... (continued)

) Burnetadine

Should not be used together. Probenecid decreases the effectiveness of bumetadine.

- > Captopril Should be used together with caution. May increase captopril level in blood.
- > Cidofovir Needs to be taken together to decrease risk of kidney toxicity.
- > Cilastatin Should be used together with caution. May increase cilastatin level in blood and increase risk of side effects.
- > Ciprofloxacin Should be used together with caution. May increase ciprofloxacin level in blood and increase risk of side effects.
- > Cisplatin Should be used together with caution. May increase cisplatin level in blood and increase risk of side effects.
- **)** Clofibrate Should be used together with caution. May increase clofibrate level in blood and increase risk of side effects.
- > Dapsone Should be used together with caution. May increase dapsone level in blood and increase risk of side effects.
- > ddC Significantly increases ddC level in blood. Check for toxicity and consider lower ddC dose.
- **>** Famotidine Should be used together with caution. Significantly increases famotidine level in blood.
- > Furosemide May increase risk of inner ear poisoning (ototoxicity).
- **)** Ganciclovir Should be used together with caution. May increase ganciclovir level in blood and increase risk of side effects.
- > Indomethacin Should be used together with caution. Increases indomethacin level in blood. May allow for lower indomethacin dose.
- > Imipenem Should be used together with caution. May increase imipenem level in blood and increases risk of side effects.
- > Ketamine Should be used together with caution. Increases ketamine level in blood.
- > Ketoprofen Should not be used together. Increases ketoprofen level in blood and increases risk of side effects.
- > Lorazepam Should be used together with caution. Increases lorazepam level in blood.
- > Meclofenamate Increases meclofenamate level in blood.

Probenecid plus ... (continued)

- **>** Methotrexate
- Should be used together with caution. Increases methotrexate level in blood and increases risk of side effects.
- > Naproxen Should be used together with caution. Increases naproxen level in blood.
- > Penicillin Should be used together with caution. Increases penicillin level 2-4 times in blood and increases risk of side effects.
- > Pyrazinamide Should be used together with caution. Decreases activity of probenecid.
- > Rifabutin Should be used together with caution. May increase rifabutin level in blood and increase risk of side effects.
- > Rifampin Should be used together with caution. May increase rifampin level in blood and increase the risk of side effects.
- > Sulfonamide Increases sulfonamide level in blood.
- > Sulindac Increases sulindac level in blood and decreases activity of probenecid.
- > Theophylline
 - May increase theophylline level in blood.
- > Thiopental Should be used together with caution. Increases thiopental level in blood.

Pyrimethamine (Daraprim) plus ...

) AZT

Should be used together with caution. May increase risk of bone marrow toxicity.

- > Dapsone
- May increase risk of bone marrow toxicity.
- > Ethionamide May increase risk of side effects from other anti-TB drugs like pyrazinamide.
- > Interferon-alfa
 - May increase risk of bone marrow toxicity.
- > Lorazepam May increase risk of liver toxicity.
- > Sulfonamides Should be used together with caution. May increase risk of bone marrow toxicity.
- > TMP/SMX Should be used together with caution. May increase risk of bone marrow toxicity.

Rifabutin (Mycobutin) plus ...

- > Amprenavir Decreases amprenavir level by 15% and increases rifabutin level by 193% in blood. Requires lower rifabutin dose.
- > Anticoagulants Should be used together with caution. May decrease effectiveness of anticoagulants. Check INR.

18

Rifabutin plus ... (continued)

- > Atovaquone Should be used together with caution. Decreases atovaquone level by 50% in blood.
-) AZT

May decrease AZT level in blood.

> Barbiturates May decrease effectiveness of barbiturates.

Beta-blockers May decrease the effectiveness of betablockers. Check response to treatment and adjust dose accordingly.

- > Buspirone May decrease effectiveness of buspirone. Check symptoms and adjust dose accordingly.
- Clarithromycin Use together with caution. May increase rifabutin level and decrease clarithromycin level in blood.
- Clofibrate May decrease effectiveness of clofibrate. Check response to treatment and adjust dose accordingly.
- Chloramphenicol May decrease the effectiveness of chloramphenicol. Check response to treatment and adjust dose accordingly.

 Corticosteroids May significantly decrease corticosteroid level in blood and require higher doses of corticosteroids.

- Cyclosporine May decrease cyclosporine level in blood. Check cyclosporine blood level and adjust dose accordingly.
- > Dapsone Should be used together with caution. May decrease dapsone level in blood.
- > Delavirdine Should not be used together*. Significantly decreases delavirdine level and increases rifabutin level in blood.
- > Diazepam May decrease effectiveness of diazepam. Check response to treatment and adjust dose accordingly.
- > Digoxin

Should be used together with caution. May significantly decrease effectiveness of digoxin. Check digoxin level in blood, control of heart rhythm and other cardiac symptoms.

- > Disopyramide May decrease effectiveness of disopyramide. Check response to treatment and adjust dose accordingly.
- > Doxycycline May decrease effectiveness of doxycycline. Check response to treatment and adjust dose accordingly.
- > Efavirenz Decreases rifabutin level in blood. Increase

rifabutin dose by 50% if taken daily or double the dose if taken 3 times a week.

Estrogen Should be used together with caution. May decrease effectiveness of estrogen.

Rifabutin plus ... (continued)

- Ethinyl estradiol Should not be used together. May decrease ethinyl estradiol level in blood. Use another form of contraception.
- Ethionamide Ethionamide May increase risk of side effects from other anti-TB drugs like rifabutin.
- Fluconazole Should be used together with caution. May increase rifabutin level by up to 80% in blood. Increases risk of painful eye inflammation.
- Fosamprenavir Should be used together with great caution. Increases risk of neutropenia. Check for side effects and lower rifabutin dose by at least 50% if fosamprenavir used alone or by 75% if boosted with ritonavir.
- > Glipizide

Should be used together with caution. May decrease glipizide level in blood. Check blood sugar level and adjust dose accordingly.

> Glyburide

Should be used together with caution. May decrease glyburide level in blood. Check blood sugar level and adjust dose accordingly.

> Indinavir

Increases rifabutin level by 204% and decreases indinavir level by 32% in blood. Reduce rifabutin dose by half the standard dose and increase indinavir dose to 1,000mg 3 times a day.

- Itraconazole Should not be used together. May increase rifabutin level and may decrease itraconazole level in blood.
- Xaletra

Increases rifabutin level by 5.7-fold in blood. Rifabutin dose should be changed to 150mg every other day or 3 times a week. Check for rifabutin side effects.

- > Ketoconazole Should be used together with caution. May decrease ketoconazole level in blood.
- > Methadone May decrease effectiveness of methadone. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Mexiletine May decrease effectiveness of mexiletine. May require lower rifabutin dose.
- > Midazolam Should not be used together. Significantly decreases midazolam level in blood. Use alternative to midazolam.
- > Nelfinavir Should be used together with caution. Increases rifabutin and decreases nelfinavir levels in blood. Reduce rifabutin dose by one-half and use nelfinavir at 1,250mg twice a day.
- > Nevirapine Should be used together with caution. May significantly affect rifabutin level in blood.
- > Norethindrone Should not be used together. May decrease norethindrone level in blood. Use another form of contraception.

Rifabutin plus ... (continued)

> Opiates

May decrease effectiveness of opiate drugs. Check pain control and response to treatment and adjust dose accordingly.

> Phenytoin Should be used together with caution. May decrease phenytoin level in blood. Check phenytoin level in blood and seizure activity. Adjust dose accordingly.

- > Probenecid Should be used together with caution. May increase rifabutin level in blood and increase risk of side effects.
- > Progesterone

Should be used together with caution. May decrease effectiveness of progesterone.

> Quinidine

May decrease quinidine level in blood. Check response to treatment and adjust dose accordingly.

> Ritonavir Should be used together with caution. Increases rifabutin level by 4-fold in blood. Increases risk of side effects. Rifabutin dose should be reduced to no more than 300mg

dose reduction may be necessary.
> Saquinavir (Fortovase) Should be used together with great caution. Decreases saquinavir level by 43% in blood. Consider alternative to rifabutin.

once a week or 150mg every 3 days. Further

Saquinavir (Invirase)

Should be used together with great caution. Decreases saquinavir level by 40% in blood. Consider alternative to rifabutin.

- > Tolbutamide Should be used together with caution. May decrease tolbutamide level in blood. Check
- blood sugar level and adjust dose accordingly.
 Theophylline May decrease theophylline level in blood.

Check theophylline level in blood and adjust dose accordingly.

- Triazolam Should not be used together. Significantly decreases triazolam level in blood. Use alternative to triazolam.
- > Trimetrexate Should be used with caution. May affect trimetrexate and/or rifabutin level.
- > Verapamil Should not be used together. May significantly decrease effectiveness of verapamil.
- > Zolpidem May decrease effectiveness of zolpidem. Check for response to treatment and adjust dose accordingly.

Rifampin (Rifadin) plus ...

> Amprenavir

Should not be used together*. Significantly decreases amprenavir level in blood.

Anticoagulants Should be used to

Should be used together with caution. May decrease effectiveness of anticoagulants. Check INR.

Rifampin plus ... (continued)

- > Atovaquone Should be used together with caution. Decreases atovaquone level by 50% in blood.
- Atazanavir Should not be used together. Rifampin decreases level of most protease inhibitors in blood.
-) AZT
- Decreases AZT level by 47% in blood. May require higher AZT dose.
- > Barbiturates May decrease effectiveness of barbiturates.
- Clarithromycin Use together with caution. May decrease clarithromycin blood level.
- Clofibrate May decrease effectiveness of clofibrate. Check response to treatment and adjust dose accordingly.
- Chloramphenicol May decrease the effectiveness of chloramphenicol. Check response to treatment and adjust dose accordingly.
- Corticosteroids May significantly decrease corticosteroid level in blood and require higher doses of corticosteroids.
- > Cyclosporine May decrease cyclosporine level in blood. Check cyclosporine blood level and adjust dose accordingly.
- > Dapsone Should be used together with caution. Decreases dapsone level 7–10 times in blood.
- > Delavirdine Should not be used together*, otherwise significantly decreases delavirdine level in blood.
- > Diazepam May decrease effectiveness of diazepam. Check response to treatment and adjust dose accordingly.
- Digoxin Should be used together with caution. May significantly decrease effectiveness of digoxin. Check digoxin level in blood, control of heart rhythm and other cardiac symptoms.
- > Disopyramide May decrease effectiveness of disopyramide. Check response to treatment and adjust dose accordingly.
- > Doxycycline May decrease effectiveness of doxycycline. Check response to treatment and adjust dose accordingly.
- Efavirenz Decreases efavirenz level by 26% in blood. Impact of interaction is uncertain.
- > Estrogen
- Should be used together with caution. May decrease effectiveness of estrogen.
- Ethinyl estradiol Should not be used together. May decrease ethinyl estradiol level in blood. Use another form of contraception.

Rifampin plus ... (continued)

- > Ethionamide
 - May increase risk of side effects from other anti-TB drugs like rifampin.
- Fluconazole Should be used together with caution. Decreases fluconazole level by 23% in blood. May require higher fluconazole dose.
- Food May decrease rifampin level in blood.
 Fosamprenavir Should not be used together*. May decrease
- anti-HIV activity of fosamprenavir.
 > Glipizide Should be used together with caution. May decrease glipizide level in blood. Check blood
- sugar level and adjust dose accordingly.
 Glyburide
 Should be used together with caution. May decrease glyburide level in blood. Check blood sugar level and adjust dose accordingly.
- > Halothane Should not be used together*. May increase risk of liver toxicity.
- Indinavir Should not be used together*. May increase rifampin level in blood.
- > Isoniazid Should be used together with caution. May increase risk of liver toxicity.
- > Itraconazole Should not be used together. Decreases rifampin level and may decrease itraconazole level in blood.
- > Ketoconazole Should not be used together*. Significantly decreases ketoconazole level in blood.
- > Kaletra Should not be used together. Decreases lopinavir level by 75% in blood.
- Methadone
 May decrease effectiveness of methadone. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Mexiletine May decrease effectiveness of mexiletine. May require lower rifabutin dose.
- > Midazolam Should not be used together. Significantly decreases midazolam level in blood. Use alternative to midazolam.
- > Nelfinavir Should not be used together*. Decreases nelfinavir level by 82% in blood.
- > Nevirapine Should not be used together. Decreases nevirapine level in blood.
- > Norethindrone Should not be used together. May decrease norethindrone level in blood. Use another form of contraception.
- > Opiates May decrease effectiveness of opiate drugs. Check pain control and response to treatment and adjust dose accordingly.

Rifampin plus ... (continued)

- > Phenytoin
- Should be used together with caution. May decrease phenytoin level in blood. Check phenytoin level in blood and seizure activity. Adjust dose accordingly.
- > Probenecid Should be used together with caution. May increase rifampin level in blood and increase risk of side effects.
- > Progesterone Should be used together with caution. May decrease effectiveness of progesterone.
- > Quinidine May decrease quinidine level in blood. Check response to treatment and adjust dose accordingly.
- Ritonavir Should not be used together. Decreases ritonavir level by 35% in blood. Consider using rifabutin instead.
- > Saquinavir (Fortovase) Should not be used together. Decreases saquinavir level by 84% in blood.
- Saquinavir (Invirase) Should not be used together. Decreases saquinavir level by 80% in blood.
- > Theophylline May decrease theophylline level in blood. Check theophylline level in blood and adjust dose accordingly.
- Triazolam Should not be used together. Significantly decreases triazolam level in blood. Use alternative to triazolam.
- > Trimetrexate Should be used together with caution. May affect trimetrexate and/or rifampin level in blood.
- > Verapamil
- Should not be used together. May significantly decrease effectiveness of verapamil.
- > Zolpidem May decrease effectiveness of zolpidem. Check for response to treatment and adjust dose accordingly.

Ritonavir (Norvir) plus ...

- > Alfentanil Should be used with caution. May significantly increase alfentanil level in blood. May require lower alfentanil dose.
- > Alprazolam Should be used together with caution. Although ritonavir decreases alprazolam level in blood, it may also prolong sedation.
- > Amiodarone Should not be used together*. Increases risk of side effects.
- > Amitriptyline Should be used together with great caution. May increase amitriptyline level in blood. May require lower amitriptyline dose.

Ritonavir plus ... (continued)

> Amlodipine

Should be used together with caution. May significantly increase amlodipine level in blood. May require lower amlodipine dose.

> Amoxapine

Should be used together with great caution. May increase amoxapine level in blood. May require lower amoxapine dose.

> Amprenavir

Increases amprenavir level by 70% in blood. Alternate dosing is amprenavir 600mg + ritonavir 100mg twice a day, or amprenavir 1,200mg + ritonavir 200mg once daily.

- > Astemizole Should not be used together*. Increases risk of side effects.
- Atazanavir Substantially increases atazanavir level in blood. If combined, use atazanavir 300mg once a day (with food) + ritonavir 100mg once a day.
- Atorvastatin Should be used together with great caution. Ritonavir + saquinavir increases atorvastatin level by 343% in blood. Use lowest dose of atorvastatin or consider using pravastatin.
- Atovaquone May decrease atovaquone level in blood.
-) AZT

Decreases AZT level by 25% in blood. No dose adjustment recommended.

- Bepridil Should not be used together*. Increases risk of side effects.
- Bromocriptine Should be used together with caution. May significantly increase bromocriptine level in blood. May require lower bromocriptine dose.
- > Buproprion Should be used together with caution. Increases risk of side effects.
- > Buprenorphine Should be used with caution. May significantly increase buprenorphine level in blood. May require lower buprenorphine dose.
- > Butorphanol Should be used with caution. May significantly increase butorphanol level in blood. May require lower butorphanol dose.
- Carbamazepine Should be used together with caution. May significantly increase carbamazepine level in blood. May require lower carbamazepine dose.
- > Chlorpromazine Should be used together with caution. May increase chlorpromazine level in blood. Check for side effects.
- Cisapride Should not be used together*. Increases risk of side effects.
- Citalopram Should be used together with caution. May increase citalopram level in blood. May require lower citalopram dose.

Ritonavir plus ... (continued)

- Clarithromycin Increases clarithromycin level by 77% and increases ritonavir level by 12% in blood. Requires lower dose of clarithromycin and checking for kidney toxicity in people with a history of kidney problems.
- Clofibrate
- May decrease clofibrate level in blood.
- Clomipramine Should be used together with caution. May increase clomipramine level in blood. May require dose reduction of clomipramine.
- Clonazepam Should be used together with caution. May increase clonazepam level in blood. May require dose reduction of clonazepam.
- Clorazepate Should be used together with caution. Increases risk of side effects.
- Clotrimazole
 May increase risk of side effects.
- Clozapine Should not be used together*. Increases risk of side effects.
- Codeine Should be used with caution. May decrease codeine level in blood. May require lower codeine dose.
- Cyclosporine May significantly increase cyclosporine level in blood.
- > ddl or ddl EC Decreases ddI level by 13% in blood. No dose adjustment recommended, but take 2½ hours apart.
- > Delavirdine Increases ritonavir level by about 51% in blood. May require lower ritonavir dose.
- Desipramine Should be used together with caution. Increases desipramine level by 145% in blood. Check desipramine level in blood and adjust dose accordingly.
- > Dexamethasone Should be used together with caution. May significantly increase dexamethasone level in blood. May require lower dexamethasone dose.
- > Diazepam Should be used together with caution. Increases risk of side effects.
- > Dihydroergotamine Should not be used together*. Increases risk of side effects.
- Diltiazem Should be used together with caution. May significantly increase diltiazem level in blood. May require lower diltiazem dose.
- > Diphenoxylate May decrease diphenoxylate level in blood.
- Disopyramide Should be used together with caution. May increase disopyramide level in blood and increase heart (cardiac) and brain/nerve (neurological) side effects. May require lower disopyramide dose.

Ritonavir plus ... (continued)

> Divalproex

Should be used together with caution. May decrease divalproex level in blood. A higher divalproex dose may be necessary.

- > Doxepin Should be used together with caution. May increase doxepin level in blood. May require lower doxepin dose.
- > Dronabinol

Should be used together with caution. May increase dronabinol level in blood. May require lower dronabinol dose.

> Efavirenz

Should be used together with caution. Increases efavirenz level by 21% and ritonavir level by 18% in blood. Check for side effects.

> Encainide Should not be used together

Should not be used together*. Increases risk of side effects.

> Enfuvirtide

Ritonavir used at the full dose increases enfuvirtide level by 22% in blood. Ritonavir at a boosting dose of 100mg twice a day increases enfuvirtide level by 14% in blood No dose adjustment recommended.

- > Ergonovine Should not be used together*. Increases risk of side effects.
- > Ergotamine Should not be used together*. Increases risk of side effects.
- > Erythromycin Should be used together with caution. May increase erythromycin level in blood. May require lower erythromycin dose.
- Escitalopram Should be used together with caution. May increase escitalopram level in blood. May require lower escitalopram dose.
- Estazolam Should be used together with caution. Increases risk of side effects.
- Ethinyl estradiol Should not be used together. Decreases ethinyl estradiol level by 40% in blood. Use another form of contraception.
- > Ethosuximide Should be used together with caution. May increase ethosuximide level in blood. May require lower ethosuximide dose.
- > Etoposide

Should be used with great caution. May increase etoposide level in blood and increase risk of side effects.

> Felodipine

Should be used together with caution. May significantly increase felodipine level in blood. May require lower felodipine dose.

- Fentanyl Should be used together with caution. May significantly increase fentanyl level in blood. May require lower fentanyl dose.
- > Flecainide Should not be used together*. Increases risk of side effects.

Ritonavir plus ... (continued)

> Flucitasone propionate Should not be used together. Increases risk of side effects.

- Fluconazole Increases ritonavir level by 12% in blood. No dose adjustment recommended; however, check for liver toxicity.
- Fluoxetine Should be used together with great caution. May increase risk of serious side effects. May require lower fluoxetine dose.
- Flurazepam Should be used together with caution. Increases risk of side effects.
- > Fluvastatin Should be used together with caution. May increase fluvastatin level in blood. Check for toxicity.
- > Fluvoxamine Should be used together with caution. May increase fluvoxamine level in blood. May require lower fluvoxamine dose.
- > Food Increases ritonavir level in blood.
- Fosamprenavir Increases fosamprenavir level in blood. Adjust dose to fosamprenavir 700mg + ritonavir 100mg twice a day or fosamprenavir 1,400mg + ritonavir 200mg once a day.
- Foscarnet Should be used with caution. Abnormal kidney function has been observed.
- > Haloperidol Should be used together with caution. May increase haloperidol level in blood. Check for side effects.
- > Hydrocodone Should be used with caution. May significantly increase hydrocodone level in blood. May require lower hydrocodone dose.
- > Hydromorphone Should be used with caution. May significantly increase hydromorphone level in blood. May require lower hydromorphone dose.
-) Imipramine
 - Should be used together with caution. May increase imipramine level in blood. May require lower imipramine dose.
- Indinavir Ritonavir significantly increases indinavir level in blood. Adjust dose to indinavir 800mg + ritonavir 200mg twice a day or indinavir 400mg + ritonavir 400mg twice a day.
- Isradipine Should be used together with caution. May significantly increase isradipine level in blood. May require lower isradipine dose.
- Itraconazole Should be used together with caution. May significantly increase itraconazole level in blood.
- > Ketoconazole

Should be used together with caution. Significantly increases ketoconazole level in blood. Ketoconazole dose should not exceed 200mg/day.

Ritonavir plus ... (continued)

- > Ketoprofen
- May increase ketoprofen level in blood.
- May increase ketorolac level in blood. > Lamotrigine Should be used together with caution. May decrease lamotrigine level in blood. A higher lamotrigine dose may be necessary.
- Lidocaine Should be used together with caution. May significantly increase lidocaine level in blood. May require lower lidocaine dose.
- > Loperamide Should not be used together. Decreases the effectiveness of loperamide.
- > Loratadine Should be used together with caution. May increase loratadine level in blood. Check for side effects.
- > Lorazepam May increase lorazepam level in blood.
 > Lovastatin
- Should not be used together. May significantly increase lovastatin level in blood.
- Maprotiline Should be used together with caution. May increase maprotiline level in blood. Check for side effects.
- > Mefloquine Should be used with great caution. May significantly decrease ritonavir level in blood.
- > Meperidine Decreases meperidine level in blood, but leads to increased concentrations of the active metabolite normeperidine over time. Long-term use in combination at higher meperidine doses is not recommended.
- > Methadone Decreases methadone level by about 40% in blood. Check for methadone withdrawal symptoms and adjust methadone dose accordingly.
- > Methamphetamine Caution is warranted. May increase methamphetamine level in blood.
- > Methylergonovine Should not be used together*. Increases risk of side effects.
- > Metoclopramide May decrease metoclopramide level in blood.
- > Metoprolol Should be used together with caution. May increase metoprolol level in blood. May require lower metoprolol dose.
- > Metronidazole Should be used together with caution. May increase risk of side effects (ritonavir liquid only).
- Mexiletine Should be used together with caution. May increase mexiletine level in blood and increase heart (cardiac) and brain/nerve (neurological) side effects. May require lower mexiletine dose.

Ritonavir plus ... (continued)

- > Miconazole
- Should be used together with caution. May increase miconazole level in blood. Check for side effects.
- > Midazolam Should not be used toget

Should not be used together*. Increases risk of side effects.

- > Morphine Should be used together with caution. May decrease morphine level in blood. Check for pain management and possible withdrawal symptoms and adjust dose accordingly.
- > Naproxen
 - $\dot{\mathrm{May}}$ increase naproxen level in blood.
- > Nefazadone

Should be used together with caution. May significantly increase nefazadone level in blood and increase cardiac and neurological side effects. May require dose reduction of nefazadone.

> Nelfinavir

Increases ritonavir level by 9% and increases nelfinavir level by 152% in blood. Other dose under study includes nelfinavir 500–750mg + ritonavir 400mg twice a day.

- > Nevirapine Decreases ritonavir level by 11% in blood. No dose adjustment recommended.
- > Nicardipine Should be used together with caution. May significantly increase nicardipine level in blood. May require lower nicardipine dose.
- > Nifedipine Should be used together with caution. May significantly increase nifedipine level in blood. May require lower nifedipine dose.
- > Nimodipine Should be used together with caution. May significantly increase nimodipine level in blood. May require lower nimodipine dose.
- > Nisoldipine Should be used together with caution. May significantly increase nisoldipine level in blood. May require lower nisoldipine dose.
- > Nitrendipine Should be used together with caution. May significantly increase nitrendipine level in blood. May require lower nitrendipine dose.
- > Nortriptyline Should be used together with caution. May increase nortriptyline level in blood. May require lower nortriptyline dose.
- > Ondansetron Should be used together with caution. May increase ondansetron level in blood. Check for side effects.
- Oxazepam May increase oxazepam level in blood.
- > Oxycodone Should be used with caution. May significantly increase oxycodone level in blood. May require lower oxycodone dose.
- > Paclitaxel

Should be used together with caution. May increase paclitaxel level in blood. Check for side effects.

Ritonavir plus ... (continued)

> Paroxetine

Should be used together with caution. May increase paroxetine level in blood. May require lower paroxetine dose.

- > Penbutolol Should be used together with caution. May increase penbutolol level in blood. May require lower penbutolol dose.
- > Pentazocine Should be used with caution. May significantly increase pentazocine level in blood. May require lower pentazocine dose.
- > Perphenazine Should be used together with caution. May increase perphenazine level in blood. May require lower perphenazine dose.
- > Phenytoin Should be used together with caution. May decrease phenytoin level in blood. An increased dose of phenytoin may be necessary.
- > Pimozide Should not be used together*. Increases risk of side effects.
- > Pindolol Should be used together with caution. May increase pindolol level in blood. Check for side effects.
- > Piroxicam Should not be used together*. Increases risk of side effects.
- > Pravastatin No clinically significant interactions with ritonavir as sole protease inhibitor. However, ritonavir + saquinavir decreases pravastatin level by 47% in blood.
- > Prednisone Should be used together with caution. May increase prednisone level in blood. May require lower dexamethasone dose.
- > Propafenone Should not be used together*. Increases risk of side effects.
- > Propoxyphene Should be used together with caution. Increases risk of side effects.
- > Protriptyline Should be used together with caution. May increase protriptyline level in blood. May require lower protriptyline dose.
- > Quinine Should be used together with caution. May significantly increase quinine level in blood. May require lower quinine dose.
- > Quinidine Should not be used together*. Increases risk of side effects.
- > Rapamycin Should be used together with caution. May significantly increase rapamycin level in blood. May require lower rapamycin dose.
- > Rifabutin
- Should be used together with caution. Increases rifabutin level by 4-fold in blood. Increases risk of side effects. Rifabutin dose should be reduced to no more than 300mg once a week or 150mg every 3 days. Further dose reduction may be necessary.

Ritonavir plus ... (continued)

- > Rifampin
- Should not be used together. Decreases ritonavir level by 35% in blood. Consider using rifabutin instead.
- > Risperidone Should be used together with caution. May increase risperidone level in blood. May require lower risperidone dose.
- > Saquinavir (Fortovase) Increases saquinavir level by 1,587% in blood. Adjust doses to saquinavir 1,000mg + ritonavir 100mg twice a day.
- > Saguinavir (Invirase) Increases saquinavir level by about 20 times in blood. Adjust doses to saquinavir 1,000mg + ritonavir 100mg twice a day.
- > Sertraline Should be used together with caution. May significantly increase sertraline level in blood. May require lower sertraline dose.
- > Sildenafil Should be used with caution. Increases sildenafil level by 11 times in blood. Sildenafil dose should not exceed 25mg within a 48 hour period.
- Simvastatin Should not be used together. Ritonavir + saquinavir increases simvastatin level by 3,059% in blood.
- > St. John's Wort Should not be used together. May significantly decrease ritonavir level in blood.
- > Tacrolimus Should be used together with caution. May significantly increase tacrolimus level in blood. May require lower tacrolimus dose.
- > Tamoxifen Should be used together with caution. May significantly increase tamoxifen level in blood. Check for side effects.
- > Tedalafil Should be used together with caution. Increases tedalafil level in blood. Tedalafil dose should not exceed 10mg in a 72 hour period.
- > Temazepam May increase temazepam level in blood.
- > Theophylline Decreases theophylline level by 43% in blood. Check theophylline level in blood and adjust dose accordingly.
- > Thioridazine Should be used together with caution. May increase thioridazine level in blood. May require lower thioridazine dose.
-) Timolol Should be used together with caution. May increase timolol level in blood. May require lower timolol dose.
- > Tipranavir Increases tipranavir level 12 times and decreases ritonavir level by 75-80% in blood. Other doses now under study.
- > Tramadol Should be used together with caution. May increase tramadol level in blood. May require lower tramadol dose.

Ritonavir plus ... (continued)

> Trazodone

Increases trazodone level in blood. Trazodone dose may need to be reduced.

- > Triazolam Should not be used together*. Increases risk of side effects.
- > Trimipramine

Should be used together with caution. May increase trimipramine level in blood. May require lower trimipramine dose.

- > Vardenafil Should be used together with caution. Increases vardenafil level in blood. Vardenafil dose should not exceed 2.5mg in a 72 hour period.
- > Venlafaxine Should be used together with caution. May
- increase venlafaxine level in blood. May require lower venlafaxine dose. > Verapamil

Should be used together with caution. May significantly increase verapamil level in blood. May require lower verapamil dose.

- > Vinblastine Should be used together with caution. May increase vinblastine level in blood. Check for side effects.
- > Vincristine Should be used together with caution. May increase vincristine level in blood. Check for side effects.
- > Voriconazole Should not be used together. Increases risk of side effects.

> Warfarin Should be used together with caution. May decrease warfarin level in blood. A higher warfarin dose may be necessary; however, closely check INR.

> Zolpidem Should be used together with caution. Increases risk of side effects.

Saguinavir (Fortovase, sqc) plus ...

SGC stands for soft gel capsule.

- > Amprenavir Decreases amprenavir level by 32% and saquinavir level by 19% in blood. No dose adjustment likely required.
- > Astemizole Should not be used together*. Increases risk of side effects.
- > Atazanavir Increases saquinavir level by 5-6 fold in blood. Dose changes now under study.
- > Atorvastatin Should be used together with great caution. Ritonavir + saquinavir increases atorvastatin level by 343% in blood.
- **)** Carbamazepine Should be used together with great caution. May decrease saquinavir level in blood.
- > Cisapride

Should not be used together*. Increases risk of side effects.

Saquinavir sgc plus ... (continued)

Clarithromycin Increases saquinavir level by 177% and increases clarithromycin level by 45% (but decreases 14-OH clarithromycin level by 24%) in blood. No dose adjustments currently recommended.

- Dihydroergotamine Should not be used together*. Increases risk of side effects.
- Delavirdine Increases saquinavir level by 5 times in blood. Reduce saquinavir dose to 800mg 3 times/day.
- > Dexamethasone Should be used together with caution. May decrease saquinavir level in blood.

> ddl or ddl EC Saquinavir should be taken 2 hours before or after taking ddI or ddI EC. Saquinavir is best taken with food, and ddI and ddI EC should only be taken on an empty stomach.

- > Efavirenz Decreases efavirenz level by 12% and saquinavir level by 62% in blood. Requires ritonavir boosting.
- > Ergonovine Should not be used together*. Increases risk of side effects.
- > Ergotamine Should not be used together*. Increases risk of side effects.
- > Fluconazole May increase saquinavir level in blood. However, No dose adjustment recommended.
- Food High fat meal increases saquinavir level by 02% in blood Should be taken within 2
- 93% in blood. Should be taken within 2 hours of a meal.
- Garlic supplements Should be used together with great caution. Decreases saquinavir level by 51% in blood.
- > Grapefruit juice Increases saquinavir level in blood.

Indinavir Increases saquinavir level 3.6–6.2 times in blood.

- > Itraconazole Increases saquinavir level in blood. However, No dose adjustment recommended.
- Kaletra
 Kaletra
 Increases saquinavir trough level by about
 3-fold. Saquinavir dose may be reduced to
 200 mg truice a day. Ontimed dosing is

800mg twice a day. Optimal dosing is under study.
> Ketoconazole Significantly increases saquinavir level in blood. However, no dose adjustment

- recommended.
 Lovastatin Should not be used together*. Increases risk of side effects.
- > Methylergonovine Should not be used together*. Increases risk of side effects.

Saquinavir sgc plus ... (continued)

> Miconazole

May increase saquinavir level in blood. However, no dose adjustment recommended.

- > Midazolam Should not be used together*. Increases risk of side effects.
- > Nelfinavir Increases nelfinavir level 18% and increases saquinavir level 4 times in blood. Saquinavir dose may be reduced to 800mg 3 times a day or 1,200mg twice a day.
- > Nevirapine Decreases saquinavir level by 24% in blood. Saquinavir may require ritonavir boosting.
- > Phenobarbital Should be used together with great caution. May decrease saquinavir level in blood.
- > Phenytoin Should be used together with great caution. May decrease saquinavir level in blood.
- > Pravastatin Ritonavir + saquinavir decreases pravastatin level by 47% in blood.
- > Rifabutin Should be used together with great caution. Decreases saquinavir level by 40% in blood. Consider alternative to rifabutin.
- Rifampin Should not be used together. Decreases saquinavir level by 84% in blood.
- Ritonavir Increases saquinavir level by 1,587% in blood. Adjust doses to saquinavir 1,000mg + ritonavir 100mg twice a day.
- Sildenafil
 Should be used together with caution.
 Increases sildenafil level by 210% in blood.
 Start sildenafil at no more than 25mg.
- Simvastatin Should not be used together*. May significantly increase simvastatin level in blood and increase risk of side effects. Ritonavir + saquinavir (soft gel) increases simvastatin level by 3,059% in blood.
- St. John's Wort Should not be used together. May decrease saquinavir level in blood.
- > Tedalafil Should be used together with caution. May increase tedalafil level in blood.
- Triazolam Should not be used together*. Increases risk of side effects.
- > Vardenafil Should be used together with caution. May increase vardenafil level in blood.

Saquinavir (Invirase, hgc) plus ...

HGC stands for hard gel capsule.

> Astemizole Should not be used together*. Increases risk of side effects.

Saquinavir hgc plus ... (continued)

> Atorvastatin

Should be used together with great caution. Ritonavir + saquinavir (soft gel) increases atorvastatin level by 343%.

- > Bepridil Should be used together with caution. May increase bepridil level in blood. Check for side effects.
- Carbamazepine Should not be used together. May decrease saquinavir level in blood.
- Clindamycin Should be used with caution. May increase clindamycin level in blood. Check for side effects.
- > Dapsone

May increase dapsone level in blood. Check for dapsone side effects.

- > ddl or ddl EC Saquinavir should be taken 2 hours before or after taking ddI or ddI EC. Saquinavir is best taken with food, and ddI and ddI EC should only be taken on an empty stomach.
- > Delavirdine Increases saquinavir level by 5 times in blood. May increase risk of GI side effects and LFTs. Reduce saquinavir dose to 800mg 3 times a day.
- > Dexamethasone Should be used together with caution. May decrease saquinavir level in blood.
- > Diltiazem Should be used together with caution. May increase diltiazem level in blood. Check for side effects.
- > Felodipine Should be used together with caution. May increase felodipine level in blood. Check for side effects.
- > Fluconazole May increase saquinavir level in blood. However, No dose adjustment recommended.
- > Food Increases saquinavir level in blood. Should be taken within 2 hours of a meal.
- > Grapefruit juice
 - Increases saquinavir level in blood.
- Indinavir
- Increases saquinavir level 4.6–7.2 times in blood.
- Itraconazole Increases saquinavir level in blood. However, No dose adjustment recommended.
- > Ketoconazole Significantly increases saquinavir level in blood. However, no dose adjustment recommended.
- Lovastatin Should not be used together*. Increases risk of side effects.
- Miconazole May increase saquinavir level in blood. However, No dose adjustment recommended.

Saquinavir hgc plus ... (continued)

> Nelfinavir

Increases saquinavir level by 4 times and increases nelfinavir level by 18% in blood. Saquinavir dose may be reduced to 800mg 3 times a day or 1,200mg twice a day.

- > Nevirapine Decreases saquinavir level 24% in blood. Saquinavir may require ritonavir boosting.
- > Nicardipine Should be used together with caution. May increase nicardipine level in blood. Check for side effects.
- > Nifedipine Should be used together with caution. May increase nifedipine level in blood. Check for side effects.
- > Nimodipine Should be used together with caution. May increase nimodipine level in blood. Check for side effects.
- > Nisoldipine Should be used together with caution. May increase nisoldipine level in blood. Check for side effects.
- > Nitrendipine Should be used together with caution. May increase nitrendipine level in blood. Check for side effects.

> Phenobarbital Should not be used together. May decrease saquinavir level in blood.

- > Phenytoin Should not be used together. May decrease saquinavir level in blood.
- > Pravastatin Ritonavir + saquinavir (soft gel) decreases pravastatin level by 47% in blood.
- > Quinidine Should be used with caution. May increase quinidine level in blood. Check for side effects.
- > Rifabutin Should be used together with great caution. Decreases saquinavir level by 40% in blood. Consider alternative to rifabutin.
- > Rifampin Should not be used together. Decreases saquinavir level by 84% in blood.
- Ritonavir Increases saquinavir level by about 20 times in blood. Adjust doses to saquinavir 1,000mg + ritonavir 100mg twice a day.
- Simvastatin Should not be used together*. May significantly increase simvastatin level in blood and increase risk of side effects. Ritonavir + saquinavir (soft gel) increases simvastatin level by 3,059% in blood.
- St. John's Wort Should not be used together. May decrease saquinavir level in blood.
- > Triazolam Should be used with caution. May increase triazolam level in blood. Check for side effects.

Stavudine (d4T, Zerit) plus ...

- > AZT
 - Should not be used together. Decreases anti-HIV activity.
- > ddl or ddl EC Should be used together with great caution. Increases risk of pancreatitis, especially in pregnant women.
- > ddC
- May increase risk of peripheral neuropathy and pancreatitis.
- S Ganciclovir May increase risk of pancreatitis.
- Hydroxyurea Increases anti-HIV activity in test tubes. May increase risk of pancreatitis.
- > Pentamidine (IV) May increase risk of pancreatitis.
- Ribavirin Should not be used together. Decreases anti-HIV activity in test tubes.

Sulfadiazine plus ...

- **)** Glipizide
- Should be used together with caution. May decrease glipizide level in blood. Check blood sugar level and adjust dose accordingly.
- > Glyburide Should be used together with caution. May decrease glyburide level in blood. Check blood sugar level and adjust dose accordingly.
- > Tolbutamide Should be used together with caution. May decrease tolbutamide level in blood. Check blood sugar level and adjust dose accordingly.

Tenofovir (Viread) plus...

- > Abacavir
- Should be used together with great caution. Although blood level not affected, tenofovir + abacavir speeds the development of resistance to both.
- > Adefovir Should not be used together*. Increases risk of kidney toxicity.
- Acyclovir Should be used together with caution. May increase risk of kidney toxicity.
- > Amikacin Should not be used together*. May increase risk of side effects.
- Amphotericin B Should not be used together*. May increase risk of side effects.
- > Atazanavir Increases tenofovir level by 24% and decreases atazanavir level by up to 40% in blood. Atazanavir should be boosted with ritonavir if used with tenofovir.
- > Cidofovir Should not be used together*. May increase risk of side effects.

Tenofovir plus ... (continued)

> ddl or ddl EC

Should be used together with great caution. Increases ddI level by 44% in blood. May increase risk of ddI side effects.

- Food (high fat meal) Increases tenofovir level by 40% in blood. Tenofovir should be taken with food.
- Foscarnet Should be used together with great caution. May increase risk of kidney toxicity.
- Ganciclovir Should be used together with great caution. May increase risk of side effects. Check for kidney toxicity.
- > Gentamycin Should not be used together*. May increase risk of side effects.
- > Kaletra Increases tenofovir level by about 30% and decreases lopinavir level 15% in blood. No dose adjustment currently recommended.
- Pentamidine (IV)
 Should not be used together*. May increase risk of side effects.
- > Tobramycin Should not be used together*. May increase risk of side effects.
- Valacyclovir Should be used together with caution. May increase risk of kidney toxicity.
- > Valganciclovir Should not be used together*. May increase risk of side effects.

TMP/SMX (Bactrim, Septra) plus ...

) 3TC

Increases 3TC level by 44% in blood. No dose adjustment recommended. Check for 3TC side effects.

> Anticancer drugs May decrease number of red blood cells (anemia) and neutrophils (neutropenia).

> AZT Should be used together with caution. May decrease number of red blood cells (anemia) and neutrophils (neutropenia).

- > Dapsone Increases trimethoprim level by 1.5 times and increases dapsone level by 1.5 times in blood.
- > ddl or ddl EC Slightly decreases trimethoprim level and increases ddI level in blood. No dose adjustment recommended.
- > Diuretics May decrease number of platelets in elderly people.
- Ganciclovir
 Should be used together with great caution. May increase risk of bone marrow toxicity.
- Indinavir Should be used together with caution. Increases trimethoprim level by 31% in blood.

25

TMP/SMX plus ... (continued)

- > Methotrexate Should be used with caution. May increase methotrexate level in blood.
- > Nevirapine Should not be started together for the first 4-6 weeks. May increase risk of rash.
- > Phenytoin Should be used with caution. May increase phenytoin level in blood. Check for side effects.
- > Pyrimethamine Should be used together with caution. May increase risk of bone marrow toxicity.
- > Warfarin Should be used with caution. May increase prothrombin (coagulation) time. Check INR.

Trimetrexate (Neutrexin) plus ...

- > Acetaminophen May affect trimetrexate and/or acetaminophen level in blood.
- Cimetidine May affect trimetrexate and/or cimetidine level in blood.
- Clarithromycin May affect trimetrexate and/or clarithromycin level in blood.
- > Erythromycin May affect trimetrexate and/or erythromycin level in blood.
- > Fluconazole May affect trimetrexate and/or fluconazole level in blood.
- > Itraconazole May affect trimetrexate and/or itraconazole level in blood.
- > Ketoconazole May affect trimetrexate and/or ketoconazole level in blood.
- > Ranitidine May affect trimetrexate and/or ranitidine level in blood.
- > Rifabutin Should be used together with caution. May affect trimetrexate and/or rifabutin level in blood.
- > Rifampin Should be used together with caution. May affect trimetrexate and/or rifampin level in blood.

Trizivir plus ...

Trizivir is a single pill containing 3 anti-HIV drugs, AZT, 3TC and abacavir. If you take Trizivir as part of your regimen, refer separately to drug interactions for AZT, 3TC and abacavir.

Zalcitabine (ddC, Hivid) plus ...

) 3TC

Should not be used together. Decreases anti-HIV activity of both drugs.

Zalcitabine plus ... (continued)

> Aminoglycosides

May increase ddC level in blood and increase risk of peripheral neuropathy. Check for toxicity.

- > Amphotericin B May increase ddC level in blood and increase risk of peripheral neuropathy. Check for toxicity.
- Antacids Decreases ddC level by 25% in blood. Take at least 2 hours apart.
- Anticancer drugs May increase risk of peripheral neuropathy.
- Chloramphenicol May increase risk of peripheral neuropathy.
 Cimetidine
 - Significantly increases ddC level in blood. Check for toxicity; consider lower ddC dose.
- Cisplatin May increase risk of peripheral neuropathy.
 Dapsone
- May increase risk of peripheral neuropathy.
- > ddl or ddl EC Should not be used together. Significantly increases risk of peripheral neuropathy.
- > Disulfiram May increase risk of peripheral neuropathy.
 > Doxorubicin

May decrease anti-HIV activity of ddC.

- > d4T May increase risk of peripheral neuropathy and pancreatitis.
- > Ethionamide May increase risk of peripheral neuropathy.
- Food May decrease ddC level in blood.
- > Foscarnet
- May increase risk of kidney toxicity and peripheral neuropathy. Check for toxicity.
- > Glutethimide May increase risk of peripheral neuropathy.
- > Hydralazine May increase risk of peripheral neuropathy.
 > lodoquinol
- May increase risk of peripheral neuropathy.
- May increase risk of peripheral neuropathy. > Metoclopramide
- Decreases ddC level by 10% in blood.Metronidazole
- May increase risk of peripheral neuropathy.
- Nitrofurantoin May increase risk of peripheral neuropathy.
- Pentamidine (IV) Should be used with great caution. May increase risk of peripheral neuropathy and pancreatitis.
- > Phenytoin
- May increase risk of peripheral neuropathy. > Probenecid
- Significantly increases ddC level in blood. Check for toxicity; consider lower ddC dose.

Zalcitabine plus ... (continued)

- > Ribavirin
- May increase risk of peripheral neuropathy.
- May increase risk of peripheral neuropathy.

Zidovudine (AZT, Retrovir) plus ...

- > Abacavir Increases AZT level by 10% in blood. No dose adjustment recommended.
- > Amphotericin B May increase risk of bone marrow toxicity.
- Amprenavir Increases AZT level by 31% and increases amprenavir level by 13% in blood. No dose adjustment recommended.
- Anticancer drugs May increase risk of bone marrow toxicity.
- > Atovaquone Increases AZT level by 35% in blood. No dose adjustment recommended.
- > Azithromycin May increase AZT level in blood. No dose adjustment recommended.
- Clarithromycin Decreases AZT level by 25% in blood. Consider taking at least 2 hours apart.
- > Dapsone
- May increase risk of bone marrow toxicity. > Doxorubicin
- Should not be used together. Decreases effectiveness and increases toxicity.
- > d4T Should not be used together. Decreases anti-HIV activity.
- Fluconazole Increases AZT level by 74% and may increase fluconazole level in blood. Check for toxicity.
- > Flucytosine May increase risk of bone marrow toxicity.
- Food Fatty foods may decrease AZT level by 57% in blood.
- > Foscarnet Should be used together with caution. May increase risk of anemia.
- > Ganciclovir Increases AZT level in blood and may increase risk of neutropenia, anemia and bone marrow toxicity. Avoid combination or use lower AZT dose.
- Indinavir Increases anti-HIV activity in test tubes. Increases AZT level by 17–36% in blood. No dose adjustment recommended.
- Interferon-alfa Increases anti-HIV activity in test tubes and may increase risk of bone marrow toxicity. Lower AZT dose by 50–75%.
- > Methadone Increases AZT level by 43% in blood. Check for AZT toxicity.
- Nelfinavir Decreases AZT level by 35% in blood. No dose adjustment recommended.

DRUG INTERACTIONS . AUGUST 2004 . PROJECT INFORM

26

Zidovudine plus (continued)	Zidovudine plus (continued)	Zidovudine plus (continued)
 > Pentamidine	 > Ribavirin	 > TMP/SMX
May increase risk of anemia. > Phenytoin	Should not be used together. Decreases	Should be used together with caution. May
May decrease phenytoin and AZT levels in	anti-HIV activity in test tubes. > Rifabutin	decrease number of red blood cells and
blood. No dose adjustment recommended. > Probenecid	May decrease AZT level in blood. > Rifampin	neutrophils thus increasing risk of anemia. > Valproic Acid
Increases AZT level by 106% in blood.	Decreases AZT level by 47% in blood. May	Increases AZT level by 80% in blood.
Adjusting AZT dose may be necessary. > Pyrimethamine	require increase in AZT dose. > Ritonavir	Check for AZT toxicity. > Vinblastine
Should be used together with caution. May	Decreases AZT level by 25% in blood. No	May increase risk of bone marrow toxicity. > Vincristine
increase risk of bone marrow toxicity.	dose adjustment recommended.	May increase risk of anemia.

Glossary of terms:

Anemia

A decrease in red blood cells. Can lead to fatigue, weakness, dizziness, and shortness of breath.

Anti-coagulant A drug that thins blood and is used to treat a number of heart conditions.

ECG

Electrocardiogram.

INR (International Normalization Ratio) A standardized way to measure various clotting factors in the blood.

NSAIDS Non-steroidal anti-inflammatory drugs.

Neutropenia

A decrease in blood cells called neutrophils that are helpful in fighting off bacterial infections.

Pancreatitis An inflammation of the pancreas. Can be life-threatening.

Peripheral neuropathy A disorder affecting the nerves of primarily the feet and hands. Symptoms may include numbness, tingling or burning sensations, pain, abnormal reflexes, weakness and partial paralysis.

Prothrombin time

A type of test to measure various clotting factors in blood.

QTc prolongation

QTc is a measurement of the pace of electrical activity of the lower chambers of the heart, which causes them to contract and relax. When the pace is slowed down (prolonged), people can experience abnormal heart rhythms, and in extreme cases spasms or a stopping of the heart.

Stevens Johnson Syndrome

Dilation of blood capillaries that results in redness and lesions all over the skin. Eyes and mouth may become swollen leading to inability to eat. Sometimes fatal.

Systemic

Throughout the body. As opposed to medicine that stays in one part of the body.

Toxicity

Side effects; the degree to which a drug can cause side effects.

Interpreting directions:

Should be used together with caution The risk of side effects from a combination is possible, but not highly common.

Should be used together with great caution

The risk of side effects from a combination is possible, and though not common may be serious if they occur.

Should not be used together

The risk of side effects from a combination is too high for the two medications to be used together. An asterisk (*) following this phrase means the side effects may be life-threatening.

Anti-HIV Medications + Street Drugs: Some Cocktails Don't Mix

Reprinted from Notes from the Underground, Winter 1998-99, #38.

For most drugs to be effective and not kill you, they need to be metabolized [broken down] by the liver or kidneys. These organs have limited resources and a set number of chemicals which accomplish this task. Because of this, certain drugs, whether they're HIV medications or recreational drugs, can affect how other drugs act. This is called a *drug interaction*— and some of them can be deadly.

There hasn't been much research on how street drugs and HIV medications interact because there is little financial incentive for the pharmaceutical companies to do the work, and because the government believes "just say no," is the only way to deal with drugs. Certainly, your best bet is not to take street drugs at all if you are taking HIV medications. Bus some interactions are deadlier than others.

Ecstasy (X, MDMA)

There has already been one death in England which resulted from a regular dose of ecstasy (MDMA, X) taken with ritonavir. Ritonavir acts to slow down the liver enzyme that breaks down X—so it makes the dose 5–10 times stronger. In addition, between 3–10% of the white population (the figure for other populations is not known) have a deficiency in this enzyme, which may be why some people overdose on what may be a safe dose for others. If you are taking any protease inhibitor [ritonavir, nelfinavir, indinavir, saquinavir (Fortovase)] or NNRTI [nevirapine delavirdine or efavirenz], X can be extremely dangerous. Of these, ritonavir and delavirdine seem to be the most dangerous, while nevirapine and efavirenz may be less so-although because effects in the test tube have sometimes been opposite to those seen in the body, this is hard to predict.

If you do take X with a protease inhibitor, wait as long as possible after taking the protease inhibitor to take the X, and be sure to have someone with you who knows what you've done in case you have difficulties. These overdoses are often not reversible, so it's really better not to mix these drugs!

Recent research has found that X damages serotonin neurons, so avoid it if you have a family or personal history of depression or anxiety disorders.

Alcohol

ddI can increase the risk of pancreatitis (intense stomach pain that feels like it's

going all the way through to your back. So, if you're using alcohol regularly, don't use ddI. There are other nucleosides to choose from.

Occasional and light use of alcohol is not known to interact with other HIV medications; however, chronic, heavy use can be destructive to the liver. This can be dangerous because the way drugs are broken down can be hurt. More drugs will stay in your system for the most part, which is likely to cause overdoses and worse side effects. Alcohol can cause dehydration; so be sure to drink lots of water to help your body deal with any alcohol you drink.

Marijuana

Protease inhibitors may increase THC levels (the active ingredient in marijuana)—so smaller doses may make you more stoned. This is also true of the synthetic version (Marinol) used in the treatment of weight loss. Since THC overdose is impossible, this interaction is not dangerous.

Sedatives

The sedatives Halcion (triazolam), Valium (diazepam), Ambien (zolpidem) and Versed (midazolam) can also be deadly if mixed with protease inhibitors. Norvir has the largest negative effect. At high doses these drugs can stop your breathing. Ativan (lorazepam), Serax (oxazepam) and Restoril (temazepam) are safer with Norvir, and may actually be weakened by it.

Barbiturates

Crixivan may increase blood levels of phenobarbital (Luminal), making overdose more likely. Other protease inhibitor interactions are also possible.

Cocaine (coke, blow)

There are no known interactions between cocaine and HIV medications, but in the test tube, cocaine doubles the speed at which the virus reproduces, meaning it may speed up how sick you get. Heroin (smack,

brown, junk, China White) Ritonavir seems to reduce heroin levels by 50% making overdose less likely. However, this drug and the other protease inhibitors have sometimes been known to have opposite effects (they cut methadone levels in real life, while test tube experiments predicted they would increase them), so caution is in order. Some synthetics sold as heroin (fentanyl, alpha-methyl-fentanyl) are potent in tiny doses and could be deadly if mixed with another drug.

GHB (gamma-hydroxy-butyrate, grievous bodily harm, liquid X GHB is potentially dangerous with Norvir and other protease inhibitors.

Amyl nitrite (amyl nitrate/poppers) Glutathione is used by the liver to process amyl nitrite, and high glutathione is linked with survival. If using amyl nitrite cuts glutathione, it could lead to disease progression.

LSD (acid) No known interactions.

Ketamine (Special K)

When combined with ritonavir, special K can lead to "chemical hepatitis," an unpleasant inflammation of the liver resulting in jaundice. A New York HIV doctor has seen two cases of it. Both went away in several weeks. But anything which damages the liver can be a serious problem for people living with HIV.

Amphetamines

(dexedrine, amphetamine, methamphetamine, crystal meth) Ritonavir is predicted to increase amphetamine levels in the blood by a factor of 2–3. The other protease inhibitors should have less of an impact, but strange opposite results are always possible.

Ritalin

Norvir and other similar drugs can either strengthen Ritalin's effects or make it weaker.

Beware!

Interactions not listed could be deadly. Street drugs are often not what they are sold as, they are frequently cut with substances that may interact with drugs themselves and their potency can vary wildly, even in the same batch. With the lack of research in this area, it's better to avoid potential interactions if at all possible.