

# HIV Drug Resistance Last updated: September 15, 2021

Together, we can change the course of the HIV epidemic...one woman at a time.

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# What Is Resistance?

HIV drugs are designed to keep the amount of HIV virus in your body under control by *preventing it from reproducing* (making copies of itself)

- HIV may change (**mutate**) as it makes copies of itself
  - Changes allow virus to overcome effects of a drug and keep reproducing
  - When this happens, HIV has developed resistance to that drug
- Resistance is a major challenge in HIV treatment:
  - Reduces treatment options



#### What Is Resistance?

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- The best way to *prevent resistance* is to stick closely (adhere) to HIV drug regimen
- With good adherence, resistance is less likely to develop
- Keeps more treatment options open for the future



What Causes Resistance?

- After entering a CD4 cell (disease-fighting white blood cell), HIV makes many new copies of itself, which then infect other CD4 cells
  - This happens very quickly; HIV can make billions of new viruses every day
- When making new viruses, HIV must copy its genetic information
- Copying happens so fast that mistakes (mutations) happen randomly
  - Some mutations are harmless
  - Other mutations can cause big problems and allow a virus to reproduce even when it is exposed to certain HIV drugs



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### What Causes Resistance?

- If a drug does not work against a mutated virus, that virus will reproduce rapidly
  - This causes viral load to go up
  - May have to change drugs to get HIV back under control
- Main reason to use a combination of HIV drugs is to block reproduction at several points in HIV's lifecycle
  - Combination of drugs aimed at several different targets is much better at preventing HIV reproduction than one alone
- With less reproduction:
  - Viral load is lower
  - Mutations and resistance are less likely to occur



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# Cross-Resistance and Sequencing

*Cross-resistance:* When mutations cause resistance not just to one drug, but to an *entire class of drugs* 

- There are seven classes of HIV drugs:
  - Nucleoside/nucleotide reverse transcriptase inhibitors (NRTIs)
  - Non-nucleoside reverse transcriptase inhibitors (NNRTIs)
  - Protease Inhibitors (PIs)
  - Entry inhibitors
  - Integrase inhibitors
  - Boosting agents
  - Post-attachment inhibitor



# Cross-Resistance and Sequencing

- Cross-resistance is *more likely to happen in some classes* than others
  - NRTIs
  - NNRTIs
  - May limit choices when picking new combination of HIV drugs
- Health care providers think about future HIV regimen options in case current regimen does not work
  - This is called **sequencing** treatment



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# How Do I Know if I Have Drug Resistance?

- Resistance is common, can be transmitted with the virus
  - People who just acquired HIV already have drug-resistant HIV
  - Have fewer HIV treatments to choose from
- People living with HIV who have taken HIV drugs are more likely to have resistant virus, fewer drug choices
- Regular viral load tests are best way to tell if you have drug resistance
  - If HIV drugs are working well, your viral load should be "undetectable"
- Have a drug resistance test if you are taking HIV drugs and:
  - Your viral load does not become undetectable
  - Your viral load goes up after you have been taking the drugs for a while



# **Resistance Testing**

- US Department of Health and Human Services guidelines recommend drug resistance testing for people who:
  - Just acquired HIV
  - Are starting HIV care
  - Have never been on HIV drugs and are planning to start
  - Are on HIV drugs and see their viral load go up (usually over 1,000 copies/mL)
  - Have started HIV drugs but viral load is not coming down to undetectable
  - Are pregnant and living with HIV (in certain circumstances)
- Resistance testing is *not usually recommended* for:
  - People who have stopped HIV drugs for four weeks or more
  - People with a viral load below 500 copies



## Ways to Test for Resistance

#### Genotype test:

- Uses HIV from your blood to check for mutations associated with drug resistance
- Preferred test for those:
  - Who are new to HIV treatment
  - Whose HIV drugs fail to bring their viral load down enough
  - Who are pregnant with detectable virus while on treatment

#### Phenotype test:

- Exposes your virus to many HIV drugs in a test tube
  - Determines which drugs still work against your HIV
  - Preferred test for people with complex drug-resistance patterns



## Ways to Test for Resistance

#### Virtual phenotype test:

- Genotype test that goes one step further
  - Takes your genotype and finds similar genotypes in a database
  - Uses their phenotypic test results to predict which drugs will be effective against your virus (and its mutations)

Resistance tests are helpful when choosing a drug regimen, *but* 

- Tests are only a guide
- Other factors (past medications, side effects, adherence) should be considered as well



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# **Preventing Resistance**

The best way to avoid resistance is to *take medications daily as prescribed* 

- Important not to skip doses
- Take medications at the same time every day
- Good adherence
- Remember, good adherence is the best way to prevent resistance
  - Follow your medication schedule
  - The virus will not reproduce as quickly
  - If not reproducing, cannot make changes that lead to resistance



#### Learn More!

- To learn more, please read the full fact sheet on this topic:
  - <u>Resistance</u>
- For more fact sheets and to connect to our community of women living with HIV, visit:
  - www.thewellproject.org
  - www.facebook.com/thewellproject
  - <u>www.twitter.com/thewellproject</u>