





HIV and medicines



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Zidovudine (AZT) in 1987









1987 – where were you?





HIV/AIDS in 1987















ART Development

Elvitegravir Dolutegravir e



29 ARVs have been developed (although we do not prescribe all of them)





What is antiretroviral therapy?



- Antiretroviral therapy (ART) is the use of HIV medicines to treat HIV infection. People on ART take a combination of HIV medicines (called an HIV *regimen*) every day.
- ART is recommended for everyone infected with HIV. ART <u>can't cure</u> HIV, but HIV medicines help people with HIV live longer, healthier lives.
- ART also reduces the risk of HIV transmission.





What is ART?

•ART stands for antiretroviral treatment. It is also called combination therapy or "HIV treatment".

What are ARVs?

•HIV drugs are called antiretrovirals (ARVs) because HIV is a type of virus called a retrovirus.

•ART nearly always includes at least three active drugs.

•Some pills contain more than one drug and some single pills contain a complete combination.





Antiretroviral (ARV) Medication



- 1996: combination therapy first shown to be effective = HAART
 - Highly
 - Active
 - Anti
 - Retroviral
 - Therapy



• 2017: 29 ARV medicines licensed in the UK

NOT A CURE





Then and now



Late 1990s



2017



One tablet taken once a day





UK Antiretrovirals 2017

•	NRTIs	Combination	Protease Inhibitors Combination
	 Lamivudine 	Kivexa, Triumeq, Combivir & Trizivir	– Lopinavir Kaletra
			– Atazanavir Evotaz
	 Abacavir 	Kivexa, Triumeq & Trizivir	– Darunavir Rezolsta, Symtuza
	 Tenofovir DF 	Truvada, Atripla, Eviplera & Stribild	Boosters
	 Tenofovir AF 	Descovy, Genvoya, Odefsey	- Ritonavir (low dose) Kaletra
	– Emtricitabine	Truvada, Descovy, Atripla,	– Cobicistat Evotaz, Rezolsta, Stribild, Genvoya, Symtuza
		Eviplera, Odefsey & Stribild	Integrase Inhibitors
	 Zidovudine 	Combivir & Trizivir	– Raltegravir
•	NNRTIS		– Elvitegravir Stribild, Genvoya
	 Efavirenz 	Atripla	– Dolutegravir Triumeq
	 Nevirapine 		CCR5 Entry Inhibitor
	 Etravirine 		 Maraviroc
	– Rilpivirine	Eviplera, Odefsey	Green text – available as a generic preparation
	Building healthier lives		Heart of England NHS



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Does ART really work?



- Yes. ART has reduced HIV-related deaths and illnesses in every country.
- More than 17 million people are now on treatment worldwide.
- ART works for adults and children, for women, men and transgender people. It works no matter how you were infected, whether this was sexually, through injecting drug use, at birth, or by blood or blood products.
- Taking drugs exactly as prescribed reduces the virus in your body to tiny amounts.
- Even though you will still be HIV positive, ART reduces the chance that you can transmit HIV.





HIV/Aids in the UK over 30 years

Number of diagnoses/deaths



ART WORKS



I'm living proof

BEFORE

I'm Joseph and I am HIV+. I nearly left it too late as HIV/A I was already sick when I went for a test. The health stayin care worker told me I had AIDS and she advised me on I'm how I could regain my strength with Anti-Retroviral Therapy or ART. I found out that ART is not just about drugs, it's about a way of living positively with VISIT YOUR LOCAL HOSPITAL OR VCT CENTRE NOW.

HIV/AIDS - by treating illnesses early, eating well and staying active. And look at me now! I'm just like I was before I got sick.

AFTER



VERNMENT OF KENT



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HIV Viral Load and CD4 Count



- HIV attacks and destroys the infection-fighting cells (called **CD4 cells**) of the immune system.
- The CD4 count in the blood falls.
- The amount of HIV in the blood (the HIV Viral Load) rises.
- Loss of CD4 cells makes it hard for the body to fight off infections and certain HIV-related cancers.





How do HIV medicines work?



- HIV medicines prevent HIV from multiplying (making copies of itself), which reduces the amount of HIV in the body. Having less HIV in the body gives the immune system a chance to recover.
- Even though there is still some HIV in the body, the immune system is strong enough to fight off infections and certain HIV-related cancers.
- By reducing the amount of HIV in the body, HIV medicines also reduce the risk of HIV transmission.





Blood test results after starting ART





After starting ART: •The HIV Viral Load falls

•CD4 count rises





When is it time to start taking HIV medicine?

- People infected with HIV should start ART as soon as possible.
- In people with the following conditions, it's especially important to start ART right away:
 - pregnancy
 - certain HIV-related illnesses and coinfections
 - early HIV infection (the period up to 6 months after infection with HIV.)







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What HIV medicines are included in an HIV regimen?

- There are many HIV medicines available for HIV regimens. The HIV medicines are grouped into drug classes according to how they fight HIV.
- A person's initial HIV regimen usually includes three HIV medicines from at least two different HIV drug classes.
- Selection of an HIV regimen depends on several factors, including possible side effects of HIV medicines and potential drug interactions between medicines.
- Because the needs of people with HIV vary, there are several HIV regimens to choose from.





5 different classes of ARVs



- 1. Nucleoside reverse transcriptase inhibitors (NRTIs, "nukes", "backbone") - zidovudine, abacavir, lamivudine, tenofovir, emtricitabine
- 2. Non-nucleoside reverse transcriptase inhibitors (NNRTIs, "non-nukes") efavirenz, nevirapine, etravirine, rilpivirine
- 3. Integrase inhibitors raltegravir, dolutegravir, elvitegravir
- 4. Protease inhibitors darunavir, atazanavir, lopinavir with a booster (ritonavir, cobicistat)
- 5. Fusion/entry inhibitors enfuvirtide, maraviroc











THREE active ARVs needed from at least 2 different drug classes

NRTI backbone – 2 drugs

• Abacavir + lamivudine (Kivexa)

OR



- Truvada
- (tenofovir disoproxil fumarate + emtricitabine)



OR

- Descovy
- (tenofovir alafenamide + emtricitabine)
 - NB 2 strengths depending on 3rd drug choice



Plus 3rd drug choice (1 drug)

NNRTI efavirenz

atazanavir /ritonavir atazanavir/cobicistat (*Evotaz*)





DNK

Examples of tablet regimens









Fixed Dose Combinations



One tablet to be taken once a day

- •Triumeq
- •Atripla
- •Eviplera



(tenofovir DF/emtricitabine/efavirenz)
] (tenofovir (DF)/emtricitabine/rilpivirine)

(abacavir/lamivudine/dolutegravir)

- •Odefsey
- •Stribild
- •Genvoya



(tenofovir (DF)/emtricitabine/elvitegravir (cobicistat)

(tenofovir (AF)/emtricitabine/rilpivirine

(tenofovir (AF)/emtricitabine/elvitegravir (cobicistat)





What does adherence mean?

- The correct dose :
 - number of tablets
 - volume of liquid
- At the correct time:
 - once or twice a day
 - Time of drug rounds on the wards
- In the correct way:
 - empty stomach (efavirenz)
 - with food (rilpivrine)
 - Avoid drug interactions
 - *e.g* vitamins/iron/indigestion remedies with integrase inhibitors.







"Drugs don't work if people don't take them"

Charles Everitt Koop







Taking every dose of your HIV treatment as prescribed will keep it working longer



People with undetectable HIV viral load (Below 400 copies)

 8 out of 10 people who took at least
 95% of their doses (dark green) reached their treatment goal (an undetectable viral load)

Your Life.

 Missing just a few doses increases your risk of treatment failure significantly

Source: Paterson DL, Swindles S, Mohr J et al. Ann Intern Med 2000; 133(1): 21-30

ADHERENCE (PATERSON)



unite is a programme from GlaxoSmithKline to help people living with HIV sustain their health and well-being in the long-term © GlaxoSmithKline HIVIPPI04/124221 - 20563377/MVL - May 2004

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Could you do it?

The same time each day.....











What Are the Key Barriers to Adherence?



Gifford AL, et al. J Acquir Immune Defic Syndr. 2000;23:386-395.













Drug resistance



- Drug resistance occurs when the virus changes its structure in a way that stops a drug from working. These changes are called drug mutations.
- Resistance only develops if you are on treatment or in the short period after stopping treatment.
- The risk of resistance increases when drug levels drop below a minimum active level. This usually only occurs if you miss doses or stop treatment. The more often you are late, the greater the chance of resistance.
- You can be infected (or reinfected) with drug resistant HIV.
- About 1 in 10 new infections in the UK have resistance to at least one drug or class of drug.





Drug interactions

A balance....





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Think about prescribed and non-prescribed drugs/remedies





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Where is this information?



- Liverpool HIV drug interactions website
 - www.hiv-druginteractions.org
 - Also available as an App (HIV iChart)
 - Or phone and speak to HIV pharmacists







Antimetrovirals and Regrantional Drop



HIV Pharmacy Service (Based at Hawthorn House)



Principal HIV Pharmacist Justine Barnes



Lead HIV Pharmacists: •Kate Gandhi (includes paediatrics) •Simone Gardner •Baldip Kaur

Rotational Senior Pharmacists: • Margaret Holmes • Shauna Henry

> Thank you Any questions?

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