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The Economics of HIV/AIDS in Low-Income Countries:

The Case for Prevention

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Abstract: There are two approaches to reducing the burden of sickness and death associated with HIV/AIDS: treatment and prevention. With limited resources, should the focus be on prevention or treatment? I discuss the range of prevention and treatment alternatives, examine their cost effectiveness, and consider various arguments that have been raised against the use of cost-effectiveness analysis in setting priorities for health. I conclude that promoting AIDS treatment using antiretrovirals in resource-constrained countries comes at a huge cost in terms of avoidable deaths that could be prevented through interventions that would substantially lower the scale of the epidemic.

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There are two approaches to reducing the burden of sickness and death associated with the human immunodeficiency virus (HIV) that leads to acquired immunodeficiency syndrome (AIDS): treatment and prevention. Treatment for HIV/AIDS involving antiretroviral therapy has been proven effective, even in resource-poor settings, in delaying the decline in the immune system, the onset of opportunistic infections, and death - and in extending the life expectancy of those infected by around four years (Ivers, Kendrick, and Doucette, 2005). Prevention measures to limit the transmission of AIDS include mass media campaigns, condom distribution, peer education of prostitutes, the prevention of mother-to-child transmission, voluntary counseling and testing, and diagnosis and treatment of other sexually transmitted diseases (since the symptoms of these diseases can make it easier for the HIV virus to spread). Despite large international aid flows for HIV/AIDS, the needs for prevention and treatment in low and middle income countries outstrip the resources available (Joint United Nations Program on HIV/AIDS, 2005). Thus, it becomes necessary to set priorities. The standard policy prescription is that in order that to maximize health, with a limited budget, funds should first be allocated to more costeffective interventions, and only then to interventions with lesser cost effectiveness. With limited resources, should the focus of efforts to combat HIV/AIDS be on prevention or treatment?

Antiretroviral treatment is complex and requires regular monitoring of adherence, efficacy, and side effects, with consequent changes in treatment regime when needed, requiring large inputs from local health services. Consequently, such treatment is expensive. Studies by Marseille Hofmann, and Kahn (2002) and Creese, Floyd, Alban and Guinness (2002) make highly favorable assumptions about the costs and benefits of highly active antiretroviral therapy, and they still conclude that a dollar spent on prevention is around 28 times more effective in reducing the burden of disease, as measured by illness and premature death. Thus, they conclude

that the limited resources available to combat HIV/AIDS in low-income countries should be targeted toward prevention rather than treatment. These results produced a rapid response at the XIV World AIDS Conference, held in Barcelona in 2002 as described by Boelaert, Van Damme, Meessen and Van der Stuyft (2002, several acronyms in the original are spelled out here):

At the Barcelona conference, Richard Feachem (Global Fund to Fights AIDS, Tuberculosis and Malaria), Gro Harlem Brundtland (World Health Organization) and Jeffrey Sachs voiced their disagreement with the conclusions of the cost-effectiveness analysis studies. All took unequivocal positions, stating, e.g. that 'The Global Fund will never hire such economists', 'prevention and treatment must go hand in hand', and 'it is wrong to accept that we have to choose between prevention or care, doing both is easily affordable'. All echoed the call of the activists: '10 billion dollars for the Global Fund, now!' and got the blessing of the international health establishment.

Among international health agencies, the consensus view is for a rapid expansion of treatment programs in developing countries. The World Health Organization has made the treatment of HIV/AIDS using antiretroviral therapies their priority, aiming to have three million people in developing and middle income countries in antiretroviral therapy by the end of 2005 (the "3 by 5" initiative). While this goal has not been achieved (there were around one million people under treatment in June 2005, (World Health Organization, 2005), the Joint United Nations Program on HIV/AIDS (UNAIDS, 2005) has a target of expanding treatment to around 9.8 million by 2010. Similarly, the United States through the President's Emergency Program for AIDS Relief, a five-year \$15 billion plan for to combat AIDS, will focus 80 percent of its

resources on treatment and care, with 20 percent going to prevention efforts. In addition, the World Bank encourages the use of its loans and grants for treatment of HIV/AIDS.

A strong case can be made for funding both prevention and treatment of HIV/AIDS. However, given the limited resources of low income countries and the fairly low levels of funding developed countries have made available in practice, choices have to be made about priorities. It is clear that the economists' method, cost-effectiveness analysis, is not the prime consideration in international organizations' policy making about HIV/AIDS in developing countries; indeed, it has been roundly rejected. I argue that this rejection is a mistake. If the goal is to maximize health benefits with the limited resources available, then cost effectiveness analysis is the right tool. Counterarguments come down either to putting forward goals other than maximizing health benefits with HIV/AIDS spending, or to pretending that resources are not scarce.

I begin with an overview of the effects of the AIDS epidemic in developing countries. I then discuss the range of prevention and treatment alternatives, and examine their cost effectiveness and the arguments that have been raised against such cost-effectiveness analysis, in terms of uncertainties in measurement, particularly when behavioral responses, feedbacks, and externalities are present. A central issue that arises is the ethical basis of cost effectiveness analysis in the public policy approach to HIV/AIDS in developing countries. While other criterion can be used for setting priorities, promoting AIDS treatment using antiretrovirals comes at a huge cost in terms of avoidable deaths that could be prevented by devoting resources to prevention interventions that would substantially lower the scale of the epidemic.

The Health, Economic, and Social burdens of HIV/AIDS

HIV/AIDS is a major cause of premature death and imposes a large disease burden around the world. Table 1 shows estimates of the number infected, the adult prevalence rate and the deaths from AIDS. There were about 40 million people in the world in 2005 infected with HIV, about 26 million of them in Africa, where the prevalence rate is 7.2 percent among the adult population aged 15 to 49. HIV/AIDS is the major health problem is Africa. A standard measure of the health costs and benefits is disability-adjusted life years (DALYs). About 18 percent of all disability-adjusted life-years lost due to health problems in Africa during 2002 were due to HIV/AIDS (World Health Organization, 2004). Most of the burden is due to early death. The disability weight (set by a panel of experts) associated with HIV is 0.135 while AIDS has a weight of 0.505, where 0 corresponds to no disability and 1 corresponds to near death The regional figures mask substantial variation, particularly in Africa where a number of countries have exceptionally high prevalence rates; Swaziland, Botswana, Lesotho, Zimbabwe, Namibia, and South Africa are estimated to have prevalence rates in excess of 20 percent. Nigeria, the most populous county in Africa has a relatively low rate but still has a large number of HIVinfected people.

Prevalence rates are usually based on samples of pregnant women at clinics and these figures may overstate prevalence rates due to selection bias (Boerma, Ghys and Walker, 2003). Population-based surveys consistently find lower estimates of prevalence, leading to speculation that actual prevalence rates may be 25-40 percent lower than standard estimates (Halperin and Post, 2004).

In addition to the health impact, HIV/AIDS also has large potential impacts on the economy and society. Health can be considered as a form of human capital and health

interventions can boost school attendance (Bleakley, 2003; Miguel and Kremer, 2004) and worker productivity (for example, Basta et al. 1979; Savedoff and Schultz, 2000; Schultz, 2002). Shastry and Weil (2002) carry out a calibration based on microeconomic estimates of the effects of health on worker productivity to show that population health should have a substantial effect on national income levels, a conclusion that is supported by cross-country growth studies (Bhargava et al., 2001; Bloom, Canning and Sevilla, 2004).

Despite this evidence that population health in general may be associated with higher income levels, there is little evidence that HIV/AIDS has led to a substantial reduction in income per capita (Bloom and Mahal, 1997; Cuddington, 1993). Diseases with high adult morbidity (sickness), such as malaria, will reduce the productivity of workers and lower GDP per capita, while diseases with high child morbidity, such as malaria and hookworm may reduce school attendance. The effect of a disease whose burden is mainly in the form of higher mortality (death) is less clear. The loss of workers will lower output but their deaths will also lower total population, reducing both the numerator and the denominator in GDP per capita. Deaths from HIV/AIDS are concentrated among young adult women and middle-aged men and there was some evidence in the early stages of the epidemic that it was more prevalent among those with high socioeconomic status, though current evidence is more mixed (Wojcicki, 2005), and these selectivity effects could reduce GDP per capita.¹

¹ In a simple economic model, if there is a fixed factor, such as land, then a reduction in the population may increase the income per capita of the survivors (Young, 2004). Of course, in a more complex economic model with economies of scale and agglomeration, a reduction in population can also decrease income per capita of survivors.

While HIV/AIDS does not seem, so far, to have reduced income per capita substantially, income per capita is not a welfare measure. A more comprehensive welfare measure that included the suffering and death of its victims would show a large welfare reduction in societies with HIV/AIDS (Jamison, Sachs and Wang, 2001; Crafts and Haacker, 2004). In addition to the direct welfare effect of lower health, resources devoted to preventing and treating HIV/AIDS will reduce consumption of other goods. Estimating the size of this effect requires national AIDS accounts that trace the flow of funds spent on prevention and treatment, combining household, government and international donor expenditures. Table 2 reports preliminary results for total expenditures on HIV/AIDS from all sources from national AIDS accounts for a number of countries that suggest that in relatively low prevalence settings, the financial burdens have so far been small. The figures for Kenya indicate that in very high prevalence, low-income, settings spending on HIV/AIDS can consume a significant proportion of GDP.

While the macroeconomic effects of HIV/AIDS to date have been muted, the long run effects are potentially large. A shorter prospective lifespan may decrease incentives to invest in human capital (Kalemli-Ozcan, Ryder and Weil, 2000) and to save for the future (Bloom, Canning and Graham, 2003). The deaths of parents and the creation of a generation of AIDS orphans may lead to low levels of health and educational investments in these children and low productivity in the future (Bell, Devarajan and Gersbach, 2004). Many African countries have a tradition of fostering relatively large numbers of children to relations and some evidence that existing AIDS orphans can be absorbed into this system with relatively little hardship in terms on nutritional and educational outcomes (Ainsworth and Filmer, 2002), but this system may break down under the strain of greater numbers of orphans.

High rates of HIV/AIDS can have major effects on social relationships and institutions, although these effects can be difficult to quantify. The high level of stigma associated with HIV/AIDS can reduce trust in the community, while high mortality and the strains imposed by coping with extreme ill health before death can weaken institutions, such as families, community groups, firms, and government agencies. The weakening of these elements of social capital may have large long-run economic consequences (Haacker, 2004).

Epidemiology, Prevention and Treatment

Epidemiology and Transmission

The HIV virus creates a U-shaped path of infection and infectiousness. In the first few months after infection, the HIV virus replicates rapidly and may cause mild flu-like symptoms. In this stage the viral load is high, particularly in genital fluids. The emergence of an immune response, partly through CD4 cells that target the virus, leads to a rapid decline in the viral load to very low, though still detectable, levels. In this stage, which may last for 10 years or more, the disease has few clinical symptoms. The CD4 cells are a target of the virus and are gradually eliminated, however, and over time the immune response weakens. At some point (usually when the CD4 concentration has fallen from its normal level of around 1000 to below 200 per ml) the viral load again rises rapidly and is associated with a rapid decline in the overall immune system. At this point the body is open to opportunistic infections (in Africa, tuberculosis is common), usually leading to death within two years. People who have HIV are most infectious to others when their viral load is high, leading to a U-shaped rate of infectiousness. Based on a study of discordant couples (where one partner has the HIV virus, but the other does not) in Uganda, the

risk of causing a new infection is highest in the first two and a half months after infection (a transmission risk of 0.0082/coital act), and in the last twenty four months of life (0.0028/ coital act), with a low of 0.0007/ coital act in the middle period (Wawer et al., 2005). While these transmission rates per sex act are low, the cumulative probability of transmission from repeated unprotected sex with an infected partner quickly becomes large.

In developing countries, HIV/AIDS is transmitted mainly through heterosexual sex and through mother-to-child transmission before or during birth, or through breast feeding. High risk groups for heterosexual transmission are commercial sex workers and their clients. In many countries, male long distant truck driver and temporary work migrants provide a vector for infection into their local communities. In some countries, despite high infection rates in these high-risk groups, prevalence rates among the general population remain low. If each case in the general population infects less than one other person on average, initial infections tend to have a very limited impact on the general population. If, however, each new infection leads to more than one subsequent infection on average, then an epidemic occurs in the general population. These two different cases have implications for whether a country should focus its prevention efforts on the high-risk groups or target the general population.

An important issue is why HIV prevalence rates are so high in some African countries. Halperin and Epstein (2004) argue that while the number of sexual partners reported in Africa is similar to that found in other regions, the pattern of sexual activity, with concurrent long term relationships being common, as opposed to the serial monogamy found elsewhere may be a factor (one-off casual sex and commercial sex are common in many countries). A study of four Africa cities with very different levels of HIV (Auvert et al., 2001), comes to the conclusion that the driving force behind the differential rates of HIV prevalence is not differences in sexual

practices, such as the number of partners, but the rate of transmission of infection per sexual act, with high levels of transmission being associated with the presence of other, untreated, sexually transmitted diseases and, for men, being uncircumcised. Oster (2005) uses an epidemiological model to argue that the higher transmission rates in Africa can explain the difference between HIV prevalence in Africa and the United States, though she attributes a greater role to differences in sexual behavior and the timing of the epidemic in explaining differences across countries within Africa.

Prevention Options

A number of prevention measures can dramatically lower the transmission of HIV/AIDS. One set of measures focuses on reducing the rate of HIV transmission through non-sexual pathways, from mothers to children, and through blood transfusions. A second set of prevention measures focus on sexual behavior, promoting changes in sexual activity like abstinence, reducing the number of partners, and condom use though mass media campaigns, school-based or peer education, condom distribution and voluntary counseling and testing. A third set of prevention measures targets transmission rates per unprotected sex act, through treating other sexually transmitted diseases and encouraging male circumcision.

Without intervention, 15-30 percent of children born to HIV positive (that is, those who have the virus) mothers will be infected either before or during birth and a further 10-20 percent will be infected via breast feeding. Mother to child transmission can be dramatically lowered by giving HIV positive mothers a single dose of nevirapine to the mother at the onset of labor and a single dose to the infant at birth (Hashimoto, Kapiga and Murata, 2002). This policy may reduce transmission to below 3 percent at birth. The issue of breast feeding is more difficult, because an

argument can be made that in situations of high child mortality the benefits to children of breast feeding by HIV-positive mothers (without the addition of any foods) for the first six months outweigh the risks from possible HIV transmission (Coutsoudis et al., 2002). Breast milk confers direct health benefits to children compared to formula milk and additional health problems can also arise when there is a lack of clean water to make up the formula milk from powder. Using HIV-infected blood in a transfusion produces almost certain transmission of the disease to the recipient, but it is now easy to test blood products before use.

Reducing the transmission of the HIV virus that occurs through sexual activity can be done either by reducing certain kinds of sexual activity or by making sexual activity less likely to transmit disease. Voluntary counseling and testing for HIV is widely available in developing countries and has a role in prevention strategies. Rapid tests based on finger prick blood spots or mouth swab are cheap and widely. Due to the likely effect of learning one's HIV status, counseling services are required, both before and after testing. Voluntary counseling and testing can play a role in prevention by reducing risky sexual behavior in those who are HIV positive, particularly if there is joint counseling of couples.

The effects of mass media national campaigns to influence behavior are difficult to estimate since a control group is usually not available. The ABC -- Abstain, Be Faithful, Use Condoms -- initiative in Uganda, combined with a high level of political commitment to HIV prevention, was successful in significantly reducing the prevalence of AIDS (Cohen, 2004). In mass efforts such as this, it is difficult to ascribe success to individual components (these is a debate about the relative importance of condoms in the ABC strategy) but they do provide evidence that broad-based and well supported efforts at behavior change can be effective prevention strategies. The impact of mass media campaigns tends to be short (Vidanapathirana,

Abramson, Forbes and Fairley, 2005) and there needs to be an ongoing effort. These mass campaigns can be aided by condom distribution, and more targeted HIV education programs aimed at youth in and out of school.

A number of interventions can reduce transmission rates for unprotected sex. HIV transmission is higher when the presence of other sexually transmitted diseases gives rise to open sores that allow entry by the virus to the blood stream – such untreated sores are common in sub-Saharan Africa. Treatment of these sexually transmitted disease is simple and cheap and can prevent the spread of HIV. While some of these interventions have been successful, the effect of such programs appears to be highly dependent on circumstances (Grosskurth et al., 1995; Gilson et al., 1997; Sangani, Rutherford and Wilkinson, 2005)

Observational evidence suggests that HIV rates are higher among uncircumcised men in Africa, and that the countries with highest rates HIV also have the highest proportions of uncircumcised men (Caldwell and Caldwell,1996). The foreskin is rich in white blood cells which may act as an entry point for HIV. Recent evidence from a randomized trial appears to support the beneficial effects of male circumcision in reducing HIV transmission (Cohen, 2005).

Treatment

Highly active antiretroviral therapy for HIV/AIDS patients, while effective in most patients, is difficult and requires a high level of medical supervision. Patients should have a CD4 test (CD4 cells are part of the immune system that help fight infections and low counts indicate a reduced immune response) or viral load (the number of HIV viruses found per milliliter of blood) to find if therapy is appropriate and at intervals to see if it is proving effective. World

Health Organization guidelines are for antiretroviral therapy to begin when the CD4 count is below 200.

Patients usually visit a clinical once a month to get a new supply of drugs, to discuss their symptoms and to take medical tests. The drugs used in treatment are toxic and patients must be monitored for side effects as well as efficacy. In addition, effectiveness depends on high levels of adherence (at least 85 to 90 percent). Counseling and follow-up to ensure adherence are required, especially since many people with HIV/AIDS suffer from depression which may affect adherence. Antiretroviral therapy treatments can interact with drugs for other illness, a serious problem since often half of those entering treatment with a CD4 count of 200 are very likely to have opportunistic infections that require treatment.

HIV mutates rapidly, and while many mutations are less virulent, resistance to antiretrovirals emerges; antiretroviral therapy usually consists of a cocktail of three drugs to combat the emergence of resistance. Once resistance to the first line of antiretroviral therapy emerges, patients can be treated with second- or third-line drug regimes. Medeiros, Diaz and Filho (2002) find in a Brazilian study that the average duration of benefits from first line antiretroviral therapy was only 14.1 months. This finding suggests that a need for second line antiretrovirals will quickly emerge in Africa, but while drug costs of first line treatment are now low, with generic drugs being available at a cost of around \$175 per year, the costs of the second line drugs can be ten times higher.

In addition to antiretrovirals, other measures can be used to prevent and treat opportunistic infections. The major opportunistic infection, and leading cause of death for those with AIDS in Africa, is tuberculosis. All HIV-positive patients, particularly those with latent tuberculosis, can take a drug to prevent its emergence. Multivitamin supplements can also reduce

the effects of opportunistic infections (Fawzi et al., 2004). Direct treatment of opportunistic infections in HIV positive patients can be effective (for example, Grant, Kaplan and Cock, 2001). In particular, the standard treatment of tuberculosis (directly observed treatment, short course) is highly effective in general and can be equally effective in HIV- positive patients (Davies et al., 1999). HIV-positive patients with tuberculosis can also be given drugs to prevent the onset of pneumonia, which can dramatically reduce mortality rates.

The Cost-effectiveness Bottom Line

Many high and middle income countries provide both prevention and treatment. Based on the high valuations of health found in cross-country estimates of willingness to pay to avoid a small chance of death (Aldy and Viscusi, 2003), Jamison and Sachs (2001) argue that countries should set the cut-off for cost effective health interventions at around three times GDP per capita. This implies that countries with a gross national product per capita in excess of \$500 should almost certainly be undertaking all of the prevention interventions outlined here, as well as firstline antiretroviral therapy. For example, Brazil is successfully providing universal access to antiretroviral therapy (Teixeira, Vitoria and Barcarlo, 2004).

The tough issue arises in the least developed countries in which resource constraints are so tight that neither sufficient prevention nor treatment are being carried out. Given the resource constraints, cost effectiveness analysis is a way of setting priorities. Table 3 sets out the range cost effectiveness estimates that have been used in four papers that compare interventions. The figures in columns A, B and C calculate cost effectiveness figures using estimates of health benefits and costs found in particular studies, Marseille, Hofmann and Kahn (2002), Creese,

Floyd, Alban and Guinness (2002), and Masaki, Green, Greig, Walsh, and Potts (2003) respectively.

A problem with cost effectiveness studies for prevention efforts against an opportunistic disease is that the overall benefits depend on the epidemiology of the disease. The effectiveness of prevention efforts tend to be higher when incidence rates are high, because more instances of transmission can be prevented. We also need to estimate not only the number of infections directly avoided by the target group, but the extent to which each infection can cause subsequent infections in a multiplier process. Changing the behavior of the population may also have feedback effects by changing risk and incentives for individuals.

A full cost effectiveness analysis therefore has to be embedded in an epidemiological model. Oster (2005) does this to examine the effect of treating sexually transmitted diseases on the course of the epidemic. This approach requires modeling different sub-populations; for example, single men and women, married men and women, and commercial sex workers and their clients are often modeled as distinct subgroups, but others groups may also be modeled depending of their importance in a particular setting, like intravenous drug users and urban versus rural dwellers. The model needs to include rates of sexual interaction and other types of transmission within and between groups, and calibrate the rate of transmission rates of HIV in different sexual activities.

Epidemiological models usually find multiplier effects, so that individual actions that reduce personal risk also reduce subsequent risk to others. However the interactions in such models are complex and may even reverse the direct effects of a prevention intervention. For example, Kremer and Morcom (1998) show that if HIV-negative people abstain from sexual activity, this can increase the proportion of HIV- positive individuals among the sexually active,

thus increasing the chances of infection for others who remain sexually active. This negative externality particularly affects those who are highly sexually active who are more likely to get, and then to spread, the disease. Abstinence by those who would otherwise have a small number of partners can have the apparently paradoxical effect of increasing prevalence rates, despite reducing overall sexual activity.

The final two columns of Table 3 give cost effectiveness estimates based on Hogan, Baltussen, Hayashi, Lauer, and Salomon (2006). The effects of each intervention are embedded in a common epidemiological model. This allows for interactions between interventions, and the possibility of complex feedback effects. The first column gives the cost effectiveness of each intervention on its own (if no other interventions were used) while the second gives the incremental cost effectiveness of each intervention based on the optimal sequencing of interventions, introduced as ranked by their cost effectiveness.

The interventions in Table 3 are ordered roughly in order of cost effectiveness. This ranking suggests that mass media messages are highly cost effective at changing behavior and reducing the transmission of HIV. Peer education, condom distribution, and treatment of sexually transmitted diseases for commercial sex workers are also highly cost effective. The treatment of tuberculosis is highly cost effective in those who are HIV positive as well as the general population. These interventions costs between \$1 and \$20 per DALY gained.

Blood transfusion safety and the prevention of mother to child transmission through single dose nevirapine are next in the ranking both with costs less than \$50 per DALY gained. Then comes the diagnosis and treatment of sexually transmitted diseases, voluntary counseling and testing, and condom distribution to medium and high risk women, which probably have costs less than \$100 per DALY gained and are always estimated at less than \$300 per DALY.

Similarly preventive therapy against tuberculosis for those who are HIV-positive costs less than \$300 per DALY gained. Prevention of mother to child transmission through using formula milk rather than breast feeding has a wide range of estimated costs and is dependent on the trade off between avoiding HIV infection and the extra risks to the child that come from using formula milk rather than breast feeding. School-based HIV/AIDS education has low cost effectiveness due to the relatively small effects on behavior it seems to bring about.

The least cost effective interventions are treatment using antiretroviral therapy. The studies reported in columns A, B and C have found that first line antiretroviral therapy has a cost in excess of \$1,000 per DALY loss averted (the figure of \$350 in the first column reflects only drug costs at the time and not the cost to the health system of providing antiretroviral therapy). The downward trend in drug prices means the figure today is probably somewhat smaller, perhaps between \$500 and \$600 as reported in the most recent study shown in Table 3.

Second-line drugs remain expensive in low income countries making this the least cost effective intervention we consider (the figure \$1977 in the fourth column is the average cost effectiveness of a policy of providing both first and second line antiretroviral therapy - second line could never be provided on its own - while the figure \$5175 in the fifth column is the marginal cost effectiveness of moving to second line after using first-line antiretrovirals).

Table 4 shows the results the costs and loss of disability-adjusted life-years averted for three different policies in 20 high mortality countries of Africa. The first policy is using all the prevention interventions with cost effectiveness calculated in the final column of table 3, with the exception of school-based HIV/AIDS education. These highly cost effective prevention interventions would avoid the loss of 30.9 million disability-adjusted life-years at a cost of \$859 per year. The alternative policy of providing first line antiretroviral therapy would avoid the loss

of 2.4 million disability-adjusted life-years at a cost of \$1350 million. Providing both first and second line antiretroviral therapy would raise the health gains only marginally, but at a large additional cost.

Focusing on prevention before turning to antiretroviral therapy will bring large health gains mainly in the form of infections avoided. Table 4 shows that a focus on prevention in the high mortality countries of Africa will reduce new HIV infections in the region by around 3.53 million per year, as compared with an estimated reduction of 40,000 a year with antiretroviral therapy. If the need for both treatment and prevention could be met, or if there were no resources available at all, priority setting would not be required. UNAIDS (2005) estimates that \$8.9 billion will be available for spending on HIV/AIDs in low and middle income countries in 2005. Because this amount is insufficient to meet all needs, redirecting this flow towards prevention could have significant effects.

There is natural uneasiness about having to make such a "tragic choice" between prevention and treatment (Calabresi and Bobbit, 1978). Such tragic choices are inevitable in low income countries where resources are so tightly constrained, highlighting the need for policies to support economic growth and poverty reduction to make such choices less necessary in the future.

Is Cost-Effectiveness Analysis Trustworthy for Priority Setting?

A number of arguments have been put forward for an emphasis on highly active antiretroviral therapy over prevention, even in the face of the evidence which suggests it is less

cost effective than prevention (for example, Boelaert et al., 2002; Farmer, 2003; Mukherjee et al., 2003). We consider the arguments in turn.

The Number Infected with AIDS is So High

One argument is that the scale of the HIV/AIDS epidemic is so great, and so threatening to overall social and economic development, that it constitutes an emergency that requires a direct response (Stabinski et al., 2003). HIV/AIDS is the biggest single health problem in Africa, accounting for 12 percent of the disability-adjusted years of life lost, followed by malaria, respiratory infections, and injuries (11, 10 and 9 percent of total disability-adjusted years of life lost, respectively (World Health Organization, 2004). Thus, it may seem obvious that health care priorities should be set according to the size of the burden imposed by the disease, and HIV/AIDS has the largest burden.

Using the burden of disease as a priority setting tool, however, makes an elementary mistake. Spending money focused on the largest health problem, or in proportion to the size of the problem, equalizes spending across needs, but does not maximize health produced. For example, if nothing could be done to reduce the burden of HIV/AIDS, then spending anything on it at all would be foolish. Maximizing health outcomes requires that at the margin the cost effectiveness of interventions be equalized. We should use the most cost-effective health interventions until they are exhausted (no more cases left to prevent or treat) or cost effectiveness falls due to diminishing returns, for example due to the difficulty in identifying rare cases, and then move on to the next most cost effective intervention. This logic argues that there is no need to know the total burden of each disease for priority setting, only the marginal cost-effectiveness of interventions (Williams, 1999)..

One argument against this marginal view of priority setting is that certain health care policies may have fixed costs (Murray and Lopez, 2000). For example, most of the costs of research and development are fixed. In the simplest possible case, where research leads to a breakthrough cure with no marginal cost per patient, the expected cost effectiveness of research is simply the cost of the research times the probability of success, divided by total burden of the disease in disability-adjusted life-years (on the grounds that this burden can then be avoided if the research is successful). Development of an effective HIV/AIDS vaccine would bring enormous benefits because of the many millions of HIV infections that could be avoided, which justifies large investments even if the probability of success is low.

There may also be fixed costs in the implementation of interventions. If only a small number of interventions are feasible due to institutional constraints, it may be better to focus on a few reasonably cost-effective interventions for high burden diseases that will produce a large total health gain rather than implement a similar number of highly cost effective but small-scale interventions.

In the case of HIV treatment and prevention, however, it is difficult to argue that a focus on antiretroviral treatment, while being less cost effective, will produce a greater total health gain than a focus on prevention. For example, Ainsworth and Teokul (2000) accept the idea that there are administrative as well as financial constraints and that we should focus on limited number of interventions, but they still argue for priority to be given to changing the behavior of the groups most at risk, treating opportunistic infections in those with HIV, and preventing the potential economic and social consequences of AIDS, for example through programs to ensure the welfare of orphans.

Nothing in the consideration of the effects of the scale of the HIV/AIDS epidemic would lead a policy-maker to put greater emphasis on treatment rather than prevention. Indeed, the enormous scale of the epidemic and its social and economic consequences suggests the importance of spending available resources as effectively as possible. Prevention efforts will tend to reduce the scale of the disease while treatment focuses on its symptoms. As shown by Table 4, promoting treatment before widespread prevention efforts denies the very interventions that can reduce the scale of the disease.

Cost-effectiveness Analysis Overstates Returns to Prevention and Understates Returns to Treatment

The idea of a disability-adjusted life year, though commonly used in cost effectiveness studies, may be called into question as a measure of health (for example, Anand and Hanson, 1997). The disability-adjusted life-year puts different weight on different disability states, and weights a year of life at different ages differently, based on the recommendations of expert committees, and discounts future disability-adjusted life years at a rate of 3 percent a year. All of these methods raise questions. Cost-effectiveness analysis also requires a number of judgment calls by the researcher, such as how to allocate overhead costs among several activities. However, most of the burden of HIV/AIDS is in the form of premature death, making it relatively insensitive to how disability is weighted. The other factors seem unlikely to produce a bias in favor of prevention.

With regard to the reliability of cost-effectiveness estimates of prevention efforts, the results on the cost effectiveness of mother to child transmission through single dose nevirapine are high and fairly uniform across a range of studies. The effectiveness of treatment of sexually

transmitted diseases, on the other hand, seems to be dependent on context (Sangani, Rutherford and Wilkinson, 2004). For interventions such as information, education, and counseling, shortrun behavior change can be observed, but studies do not usually provide convincing evidence of effects on long-run behavior or disease incidence (Grassly, 2001; Walker, 2003). Instead, longrun estimates are usually constructed through an epidemiological model that estimates the effect of the behavior change on prevalence rates.

The case for prevention through behavior change largely rests on the experience of Uganda (the ABC mass media campaign) and Thailand (where efforts concentrated on commercial sex workers), where a high level of political commitment, and multiple channels of communication, translated into successful efforts to reduce the spread of HIV/AIDS. There is the question of how replicable these success stories are in other countries.

Overall, the evidence for the cost effectiveness of prevention is clear-cut for the nonsexual pathways, mother to child transmission and blood transfusion safety. The effects of interventions that change behavior, or change transmission rates, need to be calculated in an epidemiological model that calculates how incidence and prevalence rates respond to the change. An epidemiological model can always be questions, because it is typically built on many parameters, but such models can still generate reliable results.

The cost effectiveness of treatment could also change if full account is taken of measurement problems and externalities. One important issue is that the effectiveness of antiretroviral therapy is highly dependent on keeping individual adherence to the therapy regimen above 85 percent, since below this level of adherence, the efficacy of the treatment falls off quickly. While adherence levels have been high in small-scale research studies in Africa,

there is a real concern that rapid expansion of treatment provision will lead to lower levels of counseling and support, and a fall-off in adherence (Gill et al., 2005).

The case for prevention is based, to a large extent, on the externality that each infection leads to added transmission to others, so that the public benefits of prevention efforts outweigh the private benefits. Treatment may also have externalities that are not included in cost benefit analysis. One such benefit is that making HIV/AIDS a treatable illness may reduce the stigma associated with it (Castro and Farmer, 2005), though evidence on this is so far meager. Another possible externality is that when treatment reduces the viral loads in those who have HIV/AIDS, it also can reduce the possibility of infecting sex partners -- though this effect is counterbalanced by some extent the longer lifespan and additional sexual activity for transmission of those in treatment.

Possible externalities on behavior may also arise. Since the availability of treatment raises the possibility that testing may lead to treatment, testing for HIV may become more common, allowing people who learn that they are HIV-positive to change their sexual behavior. However, the current World Health Organization recommended treatment regime in low income countries is to begin antiretroviral therapy at a CD4 count of 200, at which time opportunistic infections are likely to be present. Under this standard, early testing for HIV is unlikely to lead to earlier treatment than waiting for AIDS to manifest through opportunistic infections, and then seeking testing and treatment.

A major possible negative externality is that because the availability of treatment lowers the cost of being infected, people may become more risk taking. There is evidence of such an effect in the United States (Goldman, Lakdawalla and Sood, 2004) as well as evidence of widespread reductions in condom use in Kenya in response to (incorrect) announcements of

cures for AIDS being found (Over et al., 2004). Treatment may also have another negative externality, in the form of the development and transmission of drug resistant strains of HIV, though existing evidence on this suggests that it may not be a major concern in Africa in the foreseeable future (Blower et al., 2005).

Other externalities from HIV/AIDS are caused by the reduction in care to other members of the family when an AIDS victim dies. The death of a father or mother may reduce family income or the provision of child care, and may have enormous repercussions on children's development. But these externalities also apply to HIV/AIDs cases avoided through prevention methods and so are unlikely to tip the balance.

An important possible negative externality of a focus on treatment is that it absorbs scarce health sector workers from other activities, including cost effective treatments of other diseases. In the long run the supply of doctors and nurses can respond to the additional demand, but in the short run the high level of funding for AIDS treatment may reduce health sector workers available for other activities (Over, 2005). The shortage of even basic medical facilities and personnel in the poorest countries of Africa, particularly in the rural areas, make it difficult to see how antiretroviral treatment can be rolled out to provide close to universal coverage in the next few years. Many low income countries have difficulty providing very simple, and highly cost effective, medical services such as vaccinations for children, which makes it seem implausible that they can provide a national network of AIDS treatment facilities.

Another possible externality to treatment interventions is that "learning by doing" can increase future cost effectiveness. Even modest gains in cost effectiveness would increase the number of countries for whom antiretroviral therapy would become an affordable intervention. For example, Farmer et al. (2001) experiment with basing initiation of antiretroviral therapy on

the failure of conventional treatments for opportunistic infections, rather than laboratory tests such as CD4 counts. Badri et al. (2004) suggest that while using symptoms as a guide to initiating antiretroviral therapy reduces the number of eligible patients, relative to using a CD4 count, it does not affect mortality rates. However, most current treatment programs explicitly exclude research as a goal.

Evidence on these positive and negative externalities is not conclusive and more research into their magnitudes is required. My reading of the existing evidence is that the risk of negative externalities of treatment – that is, widespread treatment leading to higher infection rates – is at least as high as the possibility of positive externalities from treatment. Thus, using positive externalities as an argument for a large-scale treatment would require new evidence of a large net positive spillover effect. Until then, given the very large gap between the cost-effectiveness estimates for prevention and treatment, it is not likely that correcting measurement errors will reverse the ranking.

Ethical Arguments

Three common ethical arguments are put forward for emphasizing antiretroviral treatment: the need to save those whose lives are immediately at risk, the presence of a common bond with AIDS patients, and the need treat people equally globally, and.²

One common argument in favor of highly active antiretroviral therapy in developing countries is the "rule of rescue" (Jonson, 1986), which argues that people feel a psychological imperative to save those whose lives they see to be in immediate danger, irrespective of opportunity cost. While the rule of rescue has undoubtedly been a strong force in making

 $^{^{2}}$ There are also a number of arguments that could be summarized under the heading of "political economy", such as that treating some people will raise political pressure for further treatment and prevention. This issues are not dealth with here but seem unlikely to have effects large enough to lead to a support of treatment over prevention.

antiretroviral therapy a priority (Feachem, 2002), it raises difficult issues. McKie and Richardson (2003) examine the conflict between the rule of rescue and cost-effectiveness analysis. In the cost-effectiveness analysis of treatment and prevention of HIV/AIDS, those who are ill with full-blown AIDS do not receive a higher weight than those who could be prevented from getting an HIV infection in the first place (though the later gains are discounted by time). In contrast, the rule of rescue gives absolute priority to currently identifiable victims at the expense of a larger number of potential victims who have not yet been identified. It is difficult to argue that being identifiable gives extra moral weight to a person's claim on resources.

The rule of rescue implies a commitment to treat a future AIDS case, even if we do not wish to support efforts to prevent that same person becoming infected today, when prevention is a much cheaper way overcoming the problem and when the person would prefer not to be infected, which seems bizarre. The rule of rescue also suggests that antiretroviral therapy would still need to be compared in cost-effectiveness to other treatments for life-threatening conditions. There are many highly cost effective life-saving interventions in developing countries -- for example treatment of tuberculosis, and basic vaccinations for children -- which cost around \$20 per loss of disability-adjusted life year averted.

One reason for an emphasis on treatment is that there exists a global community of people concerned about HIV/AIDS, and people in this community feel a common bond with those with HIV/AIDS, wherever they may be. I once spoke for more cost-effective use of resources at a conference and a more politically knowledgeable participant whispered to me afterwards that if we did not use the funds for AIDS treatment, there would be no funds at all. While it might be more efficient in social welfare terms to concentrate resources on prevention, the group whose welfare we are considering may be *defined* as those with HIV, so that the

feeling of community and obligation only occurs after infection. Funding for HIV/AIDS treatment could be viewed as a form of charitable giving (similar, say, to giving by alumni to their college or university). Such charitable giving is always subject to the critique that the funds might have been spent with greater effect elsewhere, but such giving is still a desirable form of redistribution, raising the welfare of both recipient and giver.

Both the rule of rescue and common bond arguments essential deny that our goal should be maximizing overall population health, instead the focus should be on a particular group. While private charity may focus disproportionately on some groups, there is a strong ethical counterargument the public policy should give more equal weights to people, as is done in cost benefit analysis.

A third ethical argument is that the same standard of health care should apply around the world. Since antiretroviral therapy is generally available in high-income countries, this argument holds that it would be unethical to deny such treatment to those in low-income countries. It could be argued that taking ethical claims to our national borders, and then stopping, shows a lack of moral stamina. However, ethics frequently draws some distinctions between the imperatives that apply to smaller communities with stronger ties, such as within the nuclear family or a nation-state, and imperatives that relate to fairness and sharing with weaker ties, as in the global community. Even weak ties provide a strong ethical argument that the international community should help provide basic goods, such as health care, that are essential elements of creating capabilities in developing countries. But this does not suggest complete equality across countries, and they do not imply a priority to AIDS treatment over prevention.

One way of interpreting the argument that the standard of health care should be the same around the world is that we should not prioritize between treatment and prevention interventions,

instead the international community should make sufficient funds available to do both. But the argument that there should not be a resource constraint comes up against the hard reality that there is a resource constraint in low-income countries, and antiretroviral treatment does have a cost in terms of prevention efforts foregone.

Conclusion

Developed, middle-income, and even many low-income countries around the world can finance both prevention efforts and antiretroviral treatment for HIV/AIDS, but countries with very low incomes face binding resource limitations. In these countries, the issue is to what extent the international community should fund antiretroviral therapy. Private charity can of course choose to focus disproportionate attention on any particular group to which it feels a strong connection. However, if the goal is to maximize the health benefits produced, developing country governments and international institutions should focus their health spending first on the prevention of HIV transmission, before moving on to treatment. The opportunity cost of emphasizing HIV/AIDS treatment over prevention in a resource-constrained environment is measured in millions of lives needlessly lost.

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References

Aldy Joseph E. and W. Kip Viscusi 2003, The Value of a Statistical Life: A Critical Review of Market Estimates throughout the World, with, *Journal of Risk and Uncertainty*, 27: 5-76.

Ainsworth, Martha, and Deon Filmer, 2002, Poverty, AIDS and Children's Schooling: A Targeting Dilemma, Policy Research Working Paper No. 2885, Washington: World Bank.

Ainsworth Martha and Waranya Teokul, 2000, Breaking the silence: setting realistic priorities for AIDS control in less-developed countries, *The Lancet*, 356, 55-60.

Anand S, Hanson K, 1997, Disability-adjusted life years: a critical review, *Journal of Health Economics*, 16:685-702.

Auvert, B.; Buve, A.; Ferry, B.; Carael, M.; Morison, L.; Lagarde, E.; Robinson, N. J.; Kahindo, M.; Chege, J.; Rutenberg, N.; Musonda, R.; Laourou, M.; Akam, E.; for the Study Group on the Heterogeneity of HIV Epidemics in African Cities, 2001, Ecological and individual level analysis of risk factors for HIV infection in four urban populations in sub-Saharan Africa with different levels of HIV infection. *AIDS*, .15 Supplement 4:S15-S30.

Badri, Motasim; Bekker, Linda-Gail; Orrell, Catherine; Pitt, Jennifer; Cilliers, Francois; Wood, Robin. 2004. Initiating highly active antiretroviral therapy in sub-Saharan Africa: an assessment of the revised World Health Organization scaling-up guidelines. *AIDS*, 18(8):1159-1168.

Basta, S. S., M. S. Soekirman, D. Karyad, and N. S. Scrimshaw. 1979. Iron deficiency anemia and the productivity of adult males in Indonesia. *American Journal of Clinical Nutrition*, 32:916-25.

Bell Clive, Shantayanan Devarajan, and Hans Gersbach , 2004, Thinking About the Long-Run Economic Costs of AIDS, in *The Macroeconomics of HIV/AIDS*, Editor: Markus Haacker, International Monetary Fund, Washington D.C.

Bhargava, A. Jamison, D., Lau, L., and Murray, C. 2001. Modeling the Effects of Health on Economic Growth, *Journal of Health Economics*, 20: 423-440.

Bleakley, H. (2003), "Disease and Development: Evidence from the American South." *Journal of the European Economic Association*, Vol. 1, pp. 376–386.

Bloom D.E., Canning D. and Graham B., 2003, Longevity and Life Cycle Savings, *Scandinavian Journal of Economics*, 105: 319-338.

Bloom D.E., Canning D. and Sevilla J., 2004, "The Effect of Health on Economic Growth: A Production Function Approach", *World Development*, Vol.32, pp1-13.

Bloom David E. and Ajay Mahal, 1997, Does the AIDS Epidemic Threaten Economic Growth?' *Journal of Econometrics*, 105–124.

Blower, Sally, Bodine, Erin, Kahn, James, McFarland, Willi. 2005. The antiretroviral rollout and drug-resistant HIV in Africa: insights from empirical data and theoretical models. *AIDS*, 19(1):1-14.

Boelaert M., Van Damme W, Meessen B, Van der Stuyft P. 2002, The AIDS crisis, costeffectiveness and academic activism. *Tropical Medicine and International Health.*, 7(12):1001-2.

Boerma J Ties, Peter D Ghys and Neff Walker, 2003, Estimates of HIV-1 prevalence from national population-based surveys as a new gold standard, *The Lancet*, 362; 1929-1931.

Calabresi G. and P. Bobbit, 1978 Tragic Choices, New York, Norton & Co.

Caldwell John C. and Pat Caldwell, 1996, The African Aids Epidemic, *Scientific American*, March: 62-68.

Castro A and Farmer P. 2005. Understanding and addressing AIDS-related stigma: from anthropological theory to clinical practice in Haiti, *American Journal of Public Health*. 95(1):53-9.

Cohen J., 2005, Male Circumcision Thwarts HIV Infection, Science, 309: 860.

Cohen S. 2004, Beyond slogans: lessons from Uganda's experience with ABC and HIV/AIDS. *Reproductive Health Matters*, 12(23):132-5.

Coutsoudis, A., A. E. Goga, N. Rollins et al. 2002, Free formula milk for infants of HIV-infected women: Blessing or curse? *Health Policy and Planning*, 17(2):154-60.

Crafts Nicholas and Markus Haacker, 2004, Welfare Implications of HIV/AIDS, in *The Macroeconomics of HIV/AIDS*, Editor: Markus Haacker, International Monetary Fund, Washington D.C.

Creese Andrew, Katherine Floyd, Anita Alban and Lorna Guinness, 2002, Cost-effectiveness of HIV/AIDS interventions in Africa: a systematic review of the evidence, *The Lancet*, 359, 1635-1642.

Cuddington, John T., 1993, "Modeling the Macroeconomic Effects of AIDS, with an Application to Tanzania," *World Bank Economic Review*, Vol. 7, No. 2, pp. 173–89.

Davies, Geraint R., Connolly, Cathy, Sturm, A. Wim, McAdam, Keith P.W.J., Wilkinson, David 1999, Twice-weekly, directly observed treatment for HIV-infected and uninfected tuberculosis patients: cohort study in rural South Africa. *AIDS*, 13(7):811-817.

Farmer, Paul, 2003, AIDS as a global emergency, *Bulletin of the World Health Organization*, 81(10).

Farmer Paul, Fernet Léandre, Joia S Mukherjee, Marie Sidonise Claude, Patrice Nevil, Mary C Smith-Fawzi, Serena P Koenig, Arachu Castro, Mercedes C Becerra, Jeffrey Sachs, Amir Attaran, Jim Yong Kim, *2001*, Community-based approaches to HIV treatment in resource-poor settings, *The Lancet*, 358: 404-409.

Fawzi WW, Msamanga GI, Spiegelman D, Wei R, Kapiga S, Villamor E, Mwakagile D, Mugusi F, Hertzmark E, Essex M, Hunter DJ. 2004 A randomized trial of multivitamin supplements and HIV disease progression and mortality. *New England Journal of Medicine*, 351(1):23-32.

Feachem Richard (2002), speech to the XIV International AIDS Conference, Barcelona, available at http://www.theglobalfund.org/en/media_center/speeches/

Gill, Christopher J; Hamer, Davidson H; Simon, Jonathon L; Thea, Donald M; Sabin, Lora L 2005 No room for complacency about adherence to antiretroviral therapy in sub-Saharan Africa. *AIDS*, 19(12):1243-1249.

Gilson L, Mkanje R, Grosskurth H, Mosha F, Picard J, Gavyole A, Todd J, Mayaud P, Swai R, Fransen L, Mabey D, Mills A, Hayes R. 1997 Cost-effectiveness of improved treatment services for sexually transmitted diseases in preventing HIV-1 infection in Mwanza Region, Tanzania. *The Lancet*, 350(9094):1805-9.

Goldman Dana, Darius Lakdawalla, and Neeraj Sood, 2004, HIV Breakthroughs and Risky Sexual Behavior, NBER Working Paper No. 10516.

Grant AD, Kaplan JE, De Cock KM, 2001 Preventing opportunistic infections among human immunodeficiency virus-infected adults in African countries. *American Journal of Tropical Medicine and Hygiene*, 65(6):810-21.

Grassly NC, Garnett GP, Schwartlander B, Gregson S, Anderson RM. The effectiveness of HIV prevention and the epidemiological context. 2001, *Bulletin of the World Health Organization*, 79:1121-32.

Grosskurth H, Mosha F, Todd J, Mwijarubi E, Klokke A, Senkoro K, Mayaud P, Changalucha J, Nicoll A, ka-Gina G, et al. 1995, Impact of improved treatment of sexually transmitted diseases on HIV infection in rural Tanzania: randomised controlled trial. *The Lancet*, 346(8974):530-6.

Haacker Markus, 2004, HIV/AIDS: The Impact on the Social Fabric and the Economy, in *The Macroeconomics of HIV/AIDS*, Editor: Markus Haacker, International Monetary Fund, Washington D.C.

Hashimoto H, Kapiga SH, Murata Y. 2002 Mass treatment with nevirapine to prevent mother-tochild transmission of HIV/AIDS in sub-Saharan African countries. *Journal of Obstetric and Gynecological Research*, 28(6):313-9. Halperin Daniel T. and Helen Epstein, 2004, Concurrent sexual partnerships help to explain Africa's high HIV prevalence: implications for prevention, *The Lancet*, 364; 4-6.

Halperin Daniel T and Glenn L Post, 2004, Global HIV prevalence: the good news might be even better, *The Lancet*, 364: 1035-1036.

Hogan Daniel R, Rob Baltussen, Chika Hayashi, Jeremy A Lauer and Joshua A Salomon, 2006, Cost effectiveness analysis of strategies to combat HIV/AIDS in developing countries, *British Medical Journal*, 331;1431-1437.

Ivers, Louise C., David Kendrick, and Karen Doucette, 2005, Efficacy of Antiretroviral Therapy Programs in Resource-Poor Settings: A Meta-analysis of the Published Literature, *Clinical Infectious Diseases*, 41: 217–224.

Jamison, Dean, Sachs, Jeffery, and Wang, J., The Effect of the AIDS Epidemic on Economic Welfare in Sub-Saharan Africa, by D.T. Jamison, J.D. Sachs, and J. Wang. World Health Organization Commission on Macroeconomics and Health, Working Paper WG1:13, December 2001.

Joint United Nations Program on HIV/AIDS, 2005, *Resource needs for an expanded response to AIDS in low- and middle-income countries*, UNAIDS, Geneva.

Joint United Nations Program on HIV/AIDS, 2004, National Spending for HIV/AIDS, UNAIDS, Geneva.

Joint United Nations Program on HIV/AIDS, 2005, *Global summary of the HIV and AIDS epidemic in 2005*, UNAIDS, Geneva.

Jonsen, A.R., 1986, Bentham in a box: technology assessment and health care allocation, *Law, Medicine and Health Care*, 14:172–174.

Kalemli-Ozcan S., H.E. Ryder and D.N. Weil 2000. Mortality decline, human capital investment, and economic growth, *Journal of Development Economics*, 62:1-23.

Kremer Michael and Charles Morcom, 1998 "On the Effect of Changing Sexual Activity on HIV Prevalence," *Mathematical Biosciences*, 151(1): 99-122.

Marseille Elliot, Paul B Hofmann and James G Kahn 2002, HIV prevention before ARV in sub-Saharan Africa, *The Lancet*, 359: 1851-1856.

Masaki Emi, Russell Green, Fiona Greig, Julia Walsh, and Malcolm Potts, 2003, "Cost-Effectiveness of HIV Interventions for Resource Scarce Countries: Setting Priorities for HIV/AIDS" Bay Area International Group. <u>http://repositories.cdlib.org/big/hiv_prev</u>

McKie J. and Richardson J. 2003, The rule of rescue, *Social Science and Medicine*, 56(12):2407-19.

Medeiros R, Diaz RS, Filho AC. 2002 Estimating the length of the first antiretroviral therapy regiment durability in Sao Paulo, Brazil. *Brazilian Journal of Infectious Disease*, 6(6):298-304.

Miguel Edward and Michael Kremer (2004), Worms: Identifying Impacts on Education and Health in the Presence of Treatment Externalities, *Econometrica*, 72: 159-217.

Mukherjee J S, P E Farmer, D Niyizonkiza, L McCorkle, C Vanderwarker, P Teixeira and J Y Kim, 2003 Tackling HIV in resource poor countries, *British Medical Journal* 327;1104-1106.

Murray, Christopher J.L. and Alan D. Lopez, 2000, Progress and directions in refining the Global Burden of Disease approach: a response to Williams, *Health Economics* 9(1): 69-82.

Oster Emily, 2005, Sexually Transmitted Infections, Sexual Behavior and the HIV/AIDS Epidemic, *Quarterly Journal of Economics*, 120: 467-515.

Over Mead, 2004, Impact of the HIV/AIDS Epidemic on the Health Sectors of Developing Countries, in *The Macroeconomics of HIV/AIDS*, Editor: Markus Haacker, International Monetary Fund, Washington D.C.

Over Mead, Peter Heywood, Julian Gold, Indrani Gupta, Subhash Hira, and Elliot Marseille, 2004, *HIV/AIDS Treatment and Prevention in India: Modeling the Cost and Consequences*, Human Development Network, Health, Nutrition, and Population Series, World Bank, Washington D.C.

Sangani P, Rutherford G, Wilkinson D. 2004 Population-based interventions for reducing sexually transmitted infections, including HIV infection. *Cochrane Database, Systematic Reviews Reviews*, 2004;(2):CD001220.

Savedoff, W. D., and T. P. Schultz. (2000). *Wealth from Health*. Washington, D.C.: Inter-American Development Bank.

Schultz P. 2002, Wage Gains Associated with Height as a Form of Human Capital, *American Economic Review*, Papers and Proceedings, 92: 349-353.

Shastry G.K. and D.N. Weil. 2002. How Much of Cross-Country Income Variation is Explained by Health? Journal *of the European Economic Association*, 1: 387-396.

Stabinski Lara, Karen Pelley, Shevin T Jacob, Jason M Long, and Jennifer Leaning, 2003, Reframing HIV and AIDS, British Medical Journal, 327: 1101 – 1103.

Teixeira PR, Vitoria MA, Barcarolo J. 2004 Antiretroviral treatment in resource-poor settings: the Brazilian experience. *AIDS*, 18, Supplement 3:S5-7.

Vidanapathirana J, Abramson MJ, Forbes A, Fairley C, 2005, Mass media interventions for promoting HIV testing, *The Cochrane Database of Systematic Reviews* 2005, Issue 4.

Walker D. 2003 Cost and cost-effectiveness of HIV/AIDS prevention strategies in developing countries: is there an evidence base? *Health Policy Plan*, 18(1):4-17.

Wawer MJ et al. 2005, Rates of HIV-1 transmission per coital act by stage of HIV-1 infection, in Rakai, Uganda. *Journal of Infectious Diseases*, 191: 1403 – 1409.

Williams Alan, 1999, Calculating the global burden of disease: time for a strategic reappraisal? *Health Economics*, 8(1): 1-8.

Wojcicki JM. 2005, Socioeconomic status as a risk factor for HIV infection in women in East, Central and Southern Africa: a systematic review, *Journal of Biosocial Science*, 37(1):1-36.

World Bank, 2005, World Development Indicators 2005, Washington, World Bank.

World Health Organization (2003). *World Health Report 2003*, World Health Organization, Geneva.

World Health Organization. 2004, World Health Report 2004, World Health Organization, Geneva.

World Health Organization. 2005, *Progress on global access to HIV antiretroviral therapy: an update on "3 by 5"*, World Health Organization, Geneva.

Young, Alwyn, 2004, The Gift of the Dying: The Tragedy of AIDS and the Welfare of Future African Generations, National Bureau of Economic Research, Working Paper w10991.

		HIV Infections	Adult Prevalence	Deaths Due to		
			Rate ^a	AIDS		
			Percentage			
Sub-Saharan A	Africa	25,800,000	7.2	2,400,000		
National	Swaziland	220,000	38.8	17,000		
data for end	Botswana	350,000	37.3	33,000		
2003.	Lesotho	320,000	28.9	29,000		
	Zimbabwe	1,800,000	24.6	170,000		
	South Africa	5,300,000	21.5	370,000		
	Namibia	210,000	21.3	16,000		
	Nigeria	3,600,000	5.4	310,000		
Caribbean		300,000	1.6	24,000		
East Europe and Central		1,600,000	0.9	62,000		
Asia						
South and South East Asia		7,400,000	0.7	480,000		
North America		1,200,000	0.7	18,000		
Latin America		1,800,000	0.6	66,000		
Oceania		74,000	0.5	3,600		
Western and Central Europe		720,000	0.3	12,000		
_						
North Africa and Middle		510,000	0.2	58,000		
East						
East Asia		870,000	0.1	41,000		
World Total		40,300,000	1.1	3,100,000		

Table 1HIV and AIDS Estimates 2005

World Total40,300,0001.13,100,000Source: Joint United Nations Program on HIV/AIDS (2005) for regional data. National data are
for end 2003 are from the UNAIDS/ World Health Organization Global HIV/AIDS
Online Database, http://www.who.int/globalatlas/default.asp0

^a The adult prevalence rate is the proportion of those 15 to 49 years of age living with HIV.

Table 2

HIV/AIDS Expenditures

			HIVAIDS	
		Adult HIV	spending, per	As Percentage of
Country	Year	prevalence rate %	capita, \$US	GDP
Kenya	2002	15.0	7.61	2.23
Burkino Faso	2003	6.5	2.04	0.79
Ghana	2003	3.0	1.33	0.47
Guyana	2002	2.7	2.71	0.28
Belize	2003	2.0	6.93	0.19
Honduras	2001	1.6	3.97	0.43
Panama	2003	1.5	4.46	0.11
Thailand	2003	1.3	1.10	0.04
El Salvador	2003	1.0	5.04	0.23
Guatemala	2000	1.0	1.18	0.07
Argentina	2002	1.0	4.53	0.07
Brazil	2000	0.7	3.61	0.10
Costa Rica	2002	0.6	2.44	0.47
Venezuela	2002	0.5	3.05	0.07
Columbia	2002	0.4	1.01	0.05
Uruguay	2002	0.3	3.54	0.07
Mexico	2002	0.3	2.26	0.04
Chile	2002	0.3	1.39	0.03
Nicaragua	2002	0.2	1.61	0.21
Bolivia	2002	0.1	0.45	0.04

Authors Calculations. Source for HIV/AIDS prevalence rates, expenditures and population, Joint United Nations Program on HIV/AIDS (2004). GDP per capita from World Bank (2005).

 Table 3

 Cost Effectiveness of Prevention and Treatment of HIV/AIDS in Africa

 (U.S. dollars per loss of disability-adjusted life year averted)

	Marseille, Hofmann and Kahn (2002)	Creese, Floyd, Alban and Guinness (2002)	Masaki, Green, Greig, Walsh, and Potts (2003)	Hogan, Baltussen, Hayashi, Lauer, and Salomon (2006) (taken alone)	Hogan, Baltussen, Hayashi, Lauer, and Salomon (2006) (combined in optimal sequence)
Mass Media Campaigns				3	3
Peer education for sex workers		4-7	4		
Peer education and treatment of sexually transmitted infections for sex workers	1			3	4
Condom distribution to commercial sex workers	1	12			
Short-course tuberculosis treatment		2-21			
Blood transfusion safety	8	1-43	3		
Prevention of mother to child Transmission through single	11-19	1-81	11	34	46

dose					
Diagnosis and treatment of sexually transmitted infections	13	12	22	19-32	68-290
Voluntary counseling and testing	12-78	18-22	22	82	220
Condom distribution to high and medium risk women		48-99			
Preventive therapy for the HIV positive against tuberculosis			169-288		
Prevention of mother to child Transmission through advocating limited breast feeding or formula milk provision.		131-731			
School based HIV/AIDS education				376-530	631
antiretrovial therapy - first line drugs only	350-2000	1100-1800	1317-2029	556	547
antiretrovial therapy - first and second line drugs				1977	5175

Note: Costs are in U.S. dollars for different years. Also, costs and effectiveness both depend on assumptions on the rate of discount to be applied to future costs and the stream of future disability-adjusted life-years gained. I have not tried to rebase to a common year to allow for inflation, nor to recalculated all studies with a common discount rate.

Outcome	Cost (\$ millions)	DALY loss	HIV
Stratogy		averted (millions)	Infections Prevented
Strategy		(minons)	(IIIIIIOIIS)
Prevention	859	30.9	3.53
Antiretroviral therapy first line drugs only	1,350	2.4	0.04
Antiretroviral therapy first and second line drugs	6,434	3.2	0.04

Table 4The Cost and Benefits of Prevention and Treatmentin Twenty High Mortality Countries in Africa

Source: Adapted from data in Hogan, Baltussen, Hayashi, Lauer, and Salomon (2006). The effect of each treatment and prevention strategy is relative to no intervention.

Note: These are 20 African countries with high child and very high adult mortality rates defined by the World Health Organization (2003). It includes all of the countries in Africa with HIV prevalence rates over 20 percent.