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RESEARCH ARTICLE

The per-patient costs of HIV services in South Africa: Systematic review and application in the South African HIV Investment Case

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Abstract

Background

In economic analyses of HIV interventions, South Africa is often used as a case in point, due to the availability of good epidemiological and programme data and the global relevance of its epidemic. Few analyses however use locally relevant cost data. We reviewed available cost data as part of the South African HIV Investment Case, a modelling exercise to inform the optimal use of financial resources for the country's HIV programme.

Methods

We systematically reviewed publication databases for published cost data covering a large range of HIV interventions and summarised relevant unit costs (cost per person receiving a service) for each. Where no data was found in the literature, we constructed unit costs either based on available information regarding ingredients and relevant public-sector prices, or based on expenditure records.

Results

Only 42 (5%) of 1,047 records included in our full-text review reported primary cost data on HIV interventions in South Africa, with 71% of included papers covering ART. Other papers detailed the costs of HCT, MMC, palliative and inpatient care; no papers were found on the costs of PrEP, social and behaviour change communication, and PMTCT. The results informed unit costs for 5 of 11 intervention categories included in the Investment Case, with the remainder costed based on ingredients (35%) and expenditure data (10%).

Conclusions

A large number of modelled economic analyses of HIV interventions in South Africa use as inputs the same, often outdated, cost analyses, without reference to additional literature review. More primary cost analyses of non-ART interventions are needed.



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Introduction

In economic analyses of HIV interventions, South Africa is often used as a case in point, due to both the size of its HIV-positive population and the number of people in need of and receiving services as well as the availability of good data on the course of its HIV epidemic and the outcomes of interventions. The fact that there is also often good local cost data available has however received less attention. As an example, a large number of economic evaluations of antiretroviral treatment (ART) options for low- and middle-income countries have used South Africa as a case study for decisions facing international donors supporting HIV programmes in sub-Saharan Africa more generally [1–7]. Very few of these analyses however use cost inputs from South Africa, despite the fact that the South African ART programme has been subjected to more cost analyses than any other ART programme outside the United States [8–25].

We present a summary of current unit costs for a large number of HIV services in South Africa that were generated during the analysis for the recent South African HIV and TB Investment Case [26]. The Investment Case used a novel optimisation methodology to inform local programme planners and both local and international funders about the most cost effective mix of interventions against both HIV and TB in South Africa over the next twenty years, using as the main outcome measure cost per life-year gained. It started with a long list of interventions proposed during a stakeholder workshop which was then subjected to a rigorous review of the evidence regarding each intervention's effectiveness. Because the analysis optimised interventions and intervention coverage based on cost effectiveness, the cost of services was a central input into the analysis; at the same time, the analytical framework mandated that even though interventions could be excluded if there was no evidence as to their effectiveness, they could not be excluded if data on their cost was missing, leaving us with the task of establishing the cost of each of the interventions included in the Investment Case.

For this, we first conducted a systematic review of available published data on the costs of the selected interventions, based on primary cost analyses published between 2000 and 2016 that described a mode of delivery of the intervention that was relevant to South Africa, with or without a comparator population or intervention, and as part of any type of economic analysis, including cost and cost-effectiveness or cost-utility analyses. We then included into the Investment Case analysis those unit costs that were either, an update on such literature, or, if no literature was available, on data from recent expenditure analyses of relevant providers. Only in the absence of any such information did we use ingredient costing to establish the unit cost of a service, based on published data on the type and number of resources used in the intervention (such as staff, consumables, equipment, drugs and laboratory tests), and input costs (prices, salaries, etc) from a variety of sources. Across all types of unit cost, we used the same public-sector input costs to establish a common frame of reference.

Methods

Types of services included

The South African Investment Case followed the Investment Case framework used by UNAIDS to a certain extent [27] but additionally introduced the category of technical efficiency factor. The Investment Case framework includes biomedical interventions most often implemented by the healthcare sector alongside structural enablers (activities that have the potential to improve the efficiency of more than one intervention) and development synergies



(investments into sectors other than health that have a positive effect on HIV outcomes amongst a broader range of impacts across different health and other development sectors).

This paper reviews available information on the costs for two types of services:

- a) Interventions: These were all biomedical or behavioural interventions for which available evidence showed a direct impact on HIV risk, transmission, morbidity and mortality;
- b) Technical efficiency (TE) factors: These were defined as activities that improve the technical efficiency of existing programmes, often by increasing their quality, uptake or coverage, but only affect a single intervention- in contrast to the enablers and synergies that have the potential to affect a number of interventions, possibly across different programme areas. (For a definition of "technical efficiency" and other economics terms used in this paper, please see Box 1.)

Systematic review of cost data

We searched Pubmed for publications on the cost of each of the interventions, using a combination of MeSH and manually set search terms describing the intervention as well combinations of "economics", "cost", "cost analysis", "costing", "financial", "budget" or "resource use", at the level of title or abstract, as well as "South Africa" at the level of the text. We included

Box 1. Definitions of economic terms.

Technical efficiency in the context of the South African Investment Case refers to the maximisation of output (for example, HIV tests done) given a set level of inputs (for example, healthcare staff).

Financial costs only include accounting (or monetary) costs, whereas **economic costs** include both accounting costs and opportunity (or non-monetary) costs, for example volunteers' time or donated goods.

Micro-costing is a method of cost estimation that enumerates and costs every input needed during a health intervention. Micro-costing may be used in either top-down or bottom-up costing, depending on the level of detail and precision required.

Top-down costing starts with the total expenditure on the intervention, and uses a metric (such as allocation factors based on patient volumes per service or similar) to assign total costs to individual services.

Bottom-up costing starts by identifying actual resource use (number of staff minutes, drugs etc. per patient year) of a sample of service recipients during the intervention, and multiplying the resource use by the cost of each resource from the same period as the resources were used. These resource costs are then summed to calculate the unit cost of the intervention.

Ingredient-based costing is used to cost interventions for which there is no data from economic study. It is similar to bottom-up costing but estimates resource use during the intervention based on assumptions, expert opinion, or other literature, rather than on a sample of service recipients, and applies costs taken from literature or other sources to reach a unit cost.



Table 1. Results of literature review by intervention category.

Intervention	Papers	Papers	Full	Duplicates	Total			Pape	rs excluded				Included	% of all
category	identified: Search	identified: Other	text not found		assessed	Irrelevant intervention	Irrelevant setting	No cost data provided	No primary cost data	Study protocol only	Not peer reviewed	Total excluded		included papers
Pre-ART care	10	0	1	0	9	1	1	4	2	0	0	8	1	2%
ART	469	3	8	54	410	35	23	255	62	3	2	380	30	71%
HCT	319	1	5	119	196	25	10	138	17	2	0	192	4	10%
PMTCT	4	0	0	4	0	0	0	0	0	0	0	0	0	-
MMC	49	1	0	6	44	0	5	16	18	1	1	41	3	7%
Condom distribution	65	0	1	9	55	2	5	46	2	0	0	55	0	-
PrEP	47	1	1	7	40	0	1	28	11	0	0	40	0	-
SBCC	27	0	0	5	22	1	5	16	0	0	0	22	0	-
PEP	6	0	1	0	5	0	0	5	0	0	0	5	0	-
Palliative care	8	0	0	1	7	0	0	6	0	0	0	6	1	2%
Inpatient care	43	1	1	14	29	3	6	16	1	0	0	26	3	7%
Total	1,047	7	18	219	817	67	56	530	113	6	3	775	42	100%

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papers published in any language between 01/01/2000 and 31/12/2016 (see S1 Table for the full list of search terms). In our full-text review we included papers containing primary cost data on at least one intervention with a mode of delivery of the intervention that was relevant to South Africa, and regardless of whether or not a comparator population or intervention had been included.

We identified 1,047 papers through the search and another 7 references either mentioned in these papers or suggested by experts (Table 1). Of these 1,054 papers, 219 (21%) were duplicates, and 18 (2%) could not be found, leaving us with 817 papers for full-text screening (see Fig 1 for PRISMA diagram). Of these, 775 (95%) papers were excluded after full-text review, 530 (68% of those excluded) because no cost data was provided, 113 (15%) because no primary cost data was reported, ie, the cost data used in the paper came from another study, 67 (9%) because the intervention was not relevant to our search, 56 (7%) because the setting was not South Africa, and 6 (1%) because the paper described a study protocol only, not the results. Results of only 42 papers (5%) were included in the review.

Data from these 42 papers were abstracted using an Excel sheet which was piloted on the first 10 papers, and updated thereafter. Data entries from one researcher were checked by another researcher working independently, and discrepancies resolved by reference to the original paper. The principal summary measures that we collected were the mean or median cost estimate with ranges, cost year, currency, and where relevant, exchange rate between the reported currency and USD in the cost year. Additional abstracted data included the year of publication and first author, the country or countries where cost data had been collected, the type of economic evaluation and its objective or objectives, whether the evaluation was part of a clinical trial, the target population and any population subgroups, a description of the intervention, intervention setting and level of care, the cost analysis method (economic vs. financial, and full vs. incremental cost), the cost data collection method and time period, whether costs were discounted and whether costs and any outcome indicators came from the same setting, whether a sensitivity analysis was done, and if so, of which parameters. Finally, we collected which cost categories had been reported as having been included in the cost analysis (costs of antiretroviral and any other drugs; laboratory, radiology, staff, consumable, overhead,



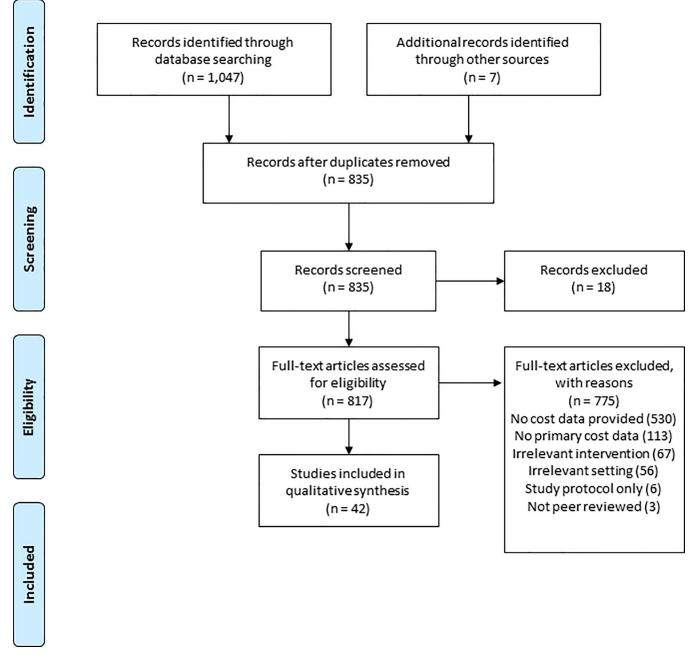


Fig 1. PRISMA diagram for systematic review.

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capital, and transportation costs). <u>S2 Table</u> summarises the definition of each parameter collected from the included papers.

Calculation of unit costs

For the purposes of this paper we define a unit cost as the cost per unit of output delivered by an intervention or technical efficiency factor (summarised as "services" in the following). The unit of output can be a person, test, visit, patient year, or a whole programme, but most often it



will be a person reached by the particular intervention. We established the unit cost for each intervention using a number of methods:

First, we used data from papers identified in our review without further adjustment if the cost year was recent and the service described was represented implementation of the service under the most recent guidelines, for the target population in question and at the relevant level of care.

Second, wherever necessary we updated the published unit cost estimates to represent more recent input costs relevant to South Africa, such as staff salaries and drug costs, while maintaining information on the types and quantities of inputs required for the intervention. All cost data taken from published estimates was updated to 2016 South African Rand (ZAR) using the South African consumer-price index published by StatsSA [28] and relevant exchange rates if the estimate was not given in ZAR.

Third, where no unit cost could be found in the literature, cost was established using ingredient costing based on published data on the type and number of resources used in the intervention, and input costs from government commodity tenders, public servant remuneration documents, retail advertisements, government programme costings and budgets from South Africa's current and past portfolio of grants with the Global Fund to Fight Aids, Tuberculosis and Malaria (GFATM). Finally, for selected interventions for which no published data on either unit cost or ingredients was available, we used information from budgets or expenditure records. This method was used for the cost of male medical circumcision and for the three social behaviour change communication campaigns included in the analysis.

The results of both the literature review and the cost calculation exercise are presented in both ZAR and US dollars (USD), using the 2017/18 period average exchange rate of 1 USD = 13.32 South African Rand.

Structure of unit cost model

The analysis used a unit cost model that calculates the cost of each ingredient used in producing 1 output of an intervention and then aggregates the ingredient costs to arrive at a total unit cost per output of the intervention. For some ingredients assumptions regarding staff productivity are required to estimate the ingredient's cost per output (for instance clients counselled per day by a counsellor). A useful benefit of building unit costs in this manner is that one is able to view the proportional contribution of each ingredient to the total unit cost.

For those interventions costed based on ingredients (rather than on literature or expenditure data), the cost per ingredient was the same across all interventions for which we used this ingredient- eg, for every intervention that required a primary healthcare nurse's time, a minute of that nurse's time costs the same. All ingredient costs were based on the most recently available public-sector data, either from sources in the public domain (such as the Department for Public Service Administration's salary scales for salaries, the Essential Drug and ARV tender price lists list for drug prices and the price lists of the National Health Laboratory Service) or from budgets from past and current South African grants with the GFATM. Unit costs based on ingredients are denoted as "From ingredients" in Table 2; unit costs based on literature updated by more recent input prices are denoted as "From ingredients, based on Author (year)" in Table 2.

Interventions for which the unit cost came directly from our literature review are denoted with the reference to the paper from which the cost was sourced in <u>Table 2</u>. For most of these estimates we updated input costs to more recent prices, using the same input costs as for those interventions costed based on ingredients. For those estimates that did not include enough details regarding ingredients and quantities, we forward-adjusted the estimate for inflation to



Table 2. Summary of results, methods and sources used in calculating unit costs. PC: Personal communication; "From ingredients": For more details regarding the quantity and prices of ingredients, please see <u>S4 Table</u>.

Service	Unit cost [2017/18 ZAR]	Unit cost [2017/18 USD]	Cost value	Costing method/ source	Notes
1. Interventions	Laik,	CoDj		I	<u> </u>
ART (Adults)	R 3,318.62	\$ 249.15	per patient year	National ART Cost Model (NACM) 201/18 [50]	Only relevant for 2017/18. Please contact the corresponding author for
ART (Paediatric)	R 3,784.19	\$ 284.10	per patient year	National ART Cost Model (NACM) 2017/18 [50]	updates if required.
Male medical	R	\$ 132.90	per	[<u>51</u>]	
circumcision	1,770.29		circumcision		
Early infant male circumcision	R 885.14	\$ 66.45	per circumcision	Based on above	Assumed to be 50% of adult MMC cost (D. Taljaard, PC)
Condom use	R 0.74	\$ 0.06	per condom distributed	From ingredients	Weighted average of male and female condoms, including distribution costs
Male and female condom education	R 66.06	\$ 4.96	per person trained	From ingredients, based on [52]	
PMTCT (mother not on any ART)	R 316.65	\$ 23.77	per mother- baby pair	National ART Cost Model (NACM) [50]	
PMTCT B (mother not on lifelong ART)	R 2,125.74	\$ 159.59	per mother- baby pair		
Infant testing at birth	R 442.23	\$ 33.20	per test	From ingredients,	
Infant testing at 6 weeks	R 416.59	\$ 31.28	per test	based on [54]	
General population HCT (negative result)	R 48.24	\$ 3.62	per test		
General population HCT (positive result)	R 74.83	\$ 5.62	per test		
Testing of pregnant women (negative result)	R 104.43	\$ 7.84	per test		
Testing of pregnant women (positive result)	R 111.64	\$ 8.38	per test		
Testing of adolescents (negative result)	R 12.1	\$ 0.91	per test		
Testing of adolescents (positive result)	R 6.19	\$ 0.46	per test		
SBCC mass media campaign 1	R 3.39	\$ 0.25	per person reached	Expenditure records from implementing	Message: testing, multiple partners
SBCC mass media campaign 2	R 1,093.2	\$ 82.07	per person reached	agencies	Message: condom usage and self-efficacy
SBCC mass media campaign 3			per person reached		Message: testing, condom usage and self-efficacy, MMC
Post-Exposure Prophylaxis (PEP)	R 1,918	\$ 143.97	per patient	From ingredients	
Pre-Exposure Prophylaxis (PrEP)	R 1,647	\$ 123.65	per patient year		
Young women, first year	R 1,900	\$ 142.65			
Young women, every year thereafter	R 1,631	\$ 122.48			

(Continued)



Table 2. (Continued)

	I			T	I
Service	Unit cost [2017/18 ZAR]	Unit cost [2017/18 USD]	Cost value	Costing method/ source	Notes
Young men, first year	R 1,915	\$ 143.77			
Young men, every year thereafter	R 1,647	\$ 123.65			
Female adolescents, first year	R 1,900	\$ 142.64			
Female adolescents, every year thereafter	R 1,631	\$ 122.42			
Male adolescents, first year	R 1,939	\$ 145.55			
Male adolescents, every year thereafter	R 1,637	\$ 122.91			
Female sex workers, first year	R 1,890	\$ 141.86			
Female sex workers, every year thereafter	R 1,621	\$ 121.68			
Men who have sex with men, first year	R 948.48	\$ 71.21			
Men who have sex with men, every year thereafter					
Palliative care	R 1,694.49	\$ 127.21	per patient	A. Lolliot, HPCA (PC)	
Inpatient care	R 989.47	\$ 74.28			
pre-ART, <200 cells/ microl	R 808.77	\$ 60.72	per patient year	[53], adjusted to different CD4 strata	
pre-ART, 200–349 cells/microl	R 393.23	\$ 29.52			
pre-ART, 350–500 cells/microl	R 2079.97	\$ 156.15			
pre-ART, >500 cells/ microl	R 1660.27	\$ 124.64			
ART, <200 cells/ microl	R 852.19	\$ 63.98			
ART, 200–349 cells/ microl	R 756.87	\$ 56.82			
ART, 350–500 cells/ microl	R 3,318.62	\$ 249.15			
ART, >500 cells/ microl	R 3,784.19	\$ 284.10			
2. Technical efficiency	factors				
Provider initiated counselling and testing (negative result)	R 56.07	\$ 4.21	per test	Updated based on [54]	
Provider initiated counselling and testing (positive result)	R 82.66	\$ 6.21	per test	Updated based on [54]	
Mobile HCT (negative result)	R 74.48	\$ 5.59	per test	[34]	
Mobile HCT (positive result)	R 87.92	\$ 6.60	per test		

(Continued)



Table 2. (Continued)

Service	Unit cost [2017/18 ZAR]	Unit cost [2017/18 USD]	Cost value	Costing method/ source	Notes
Home based HCT (negative result)	R 69.73	\$ 5.23	per test	[34]	
Home based HCT (positive result)	R 77.42	\$ 5.81	per test		
Workplace HCT (negative result)	R 63.14	\$ 4.74	per test	From ingredients	
HCT invitations to partners of pregnant women (negative result)	R 55.50	\$ 4.17	per test	From ingredients	
HCT invitations to partners of pregnant women (positive result)	R 50.79	\$ 3.81	per test	From ingredients	

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2016 costs as above. Estimates that were made available to us ahead of being published are denoted as "personal communication", or "PC", in Table 2.

Lastly, for those interventions for which neither literature on costs nor detailed ingredients nor expenditure data were available, most often because they were either so new that cost analyses had not yet been undertaken, we used expenditure data instead (denoted as "Expenditure records from implementing agencies" in Table 2).

It should be borne in mind that the purpose of this paper is to summrise potential sources of cost data rather than every sub-aspect of our own or other analysts' cost analyses. For more information, we suggest to contact either the authors of the papers we reference or, if more information on the ingredients cost summarised in S4 Table is required, the corresponding author of this paper. Alternatively, we suggest searching the unit cost study repository of the Global Health Cost Consortium (https://ghcosting.org/pages/data/ucsr/app/index) which, at the time of writing, contained data from four of the papers included in our review).

Results

Literature review

Of the 42 papers identified through the literature review (5% of all papers included in full-text review), 30 (71%) reported on the cost of antiretroviral treatment in different models of care, 4 (10%) reported on HIV counselling and testing (HCT), 3 (7%) each on medical male circumcision and inpatient care, and 1 each (2%) on palliative care and pre-ART care (cotrimoxazole preventive therapy only) (Table 2). We did not find any cost papers reporting primary cost data relevant for South Africa on PMTCT, pre-exposure prophylaxis (PrEP), condom use, social and behaviour change communication (SBCC), or post-exposure prophylaxis (PEP). Papers found spanned the 12 years between 2005 and 2016, with slightly more papers (60%) published in the second six years than in the first.

Of the 42 papers, 34 (81%) reported resource use from the perspective of the healthcare provider (with the exception of one paper [29], the provider was the public sector), and 6 papers (14%) reported resource use from the patient perspective (with 5 and 1 papers, respectively, reporting on the costs associated with accessing ART and VMMC). Two papers (4%) reported on both provider and patient perspectives [30,31]. One paper (2%) on the cost of NIMART purported to have used a societal perspective by summing costs from the healthcare provider



and patient perspectives [12]. Thirty-two papers (76%) presented full cost, 8 papers (19%) incremental costs, one paper (2%) reported both [32], and one paper (2%) did not give any information [33]. Thirty-two papers (76%) reported financial costs only, 5 papers (12%) economic costs, and another 5 papers (12%) both financial and economic costs. 55% of papers used a bottom-up approach to analysing resource use, 43% an ingredients or top-down approach, and one paper (2%) used both [34]. The majority of papers (62%) reported results as a point estimate only, without giving a range.

Cost categories included varied between intervention areas and perspectives. Cost items included in papers that only reported the provider perspective were staff costs (in 81% of these papers), consumables (69%), ARVs (in 61% of papers), non-ARV drugs (69%), laboratory tests (58%), equipment (58%), radiology (36%), inpatient cost (19%), transport costs (17%). Other specified cost items not attributable to these categories comprised nutritional support or supplements [35,36], communication costs [37], the costs of end of life care [34], field materials [38] and start-up costs [39], bank and interest charges [40] and insurance costs [41].

See S3 Table for the detailed cost results by paper.

Unit costs

Table 2 gives an overview of the methods used in arriving at the unit cost of each intervention and TE factor included in the South African HIV Investment Case, the sources of all data, and the resulting unit cost. S4 Table furthermore gives details on the type, cost and quantities of ingredients included in each unit cost where applicable.

We were able to find estimates of the cost of 55% of the services in the literature, almost all of which we further updated to represent the most recent South African input prices. 35% of services were costed based on ingredients; the remaining 10% of services were costed based on expenditure data.

Discussion

We reviewed the available literature on primary cost data for HIV services in South Africa. Despite HIV interventions making up a large proportion of the health interventions in South Africa for which cost estimates are available [42], and despite a large number of reviews showing that South Africa is home to more cost estimates for HIV interventions than any other low- and middle-income country [43–46], our search yielded limited results.

Only 5% of the papers on HIV interventions that used the term "cost" or "economics" as well as "South Africa" in the text reported the results of a primary cost analysis. Instead, a large number of modelled economic analyses of HIV interventions in South Africa which made reference to the same, often outdated, cost analyses, with no reference to any additional review of other literature. For example, a commonly cited cost source is one of the very early costs of ART in an NGO-run programme years before the public-sector roll-out [9]. This cost data was collected in 2003 and published in 2006. While the importance of the paper as one of very few sources of ART cost data in the early years of the public-sector ART programme cannot be overstated, it was still used as a cost source for papers published as recently as 2016 [47–49]- at which stage the data used in the models was over 10 years old.

We also found that the actual age of the input costs used in models was often lost through chains of referencing, where a modelled analysis used input costs that were the outputs of an older modelled analysis based on an even older primary analysis. In an effort to correct for outdated costs, those modelled analyses tended to inflate primary cost data over many years using general inflation indices such as the consumer price index, instead of sourcing and using more recent input prices for salaries, drug costs, etc, which skewed resulting estimates considerably,



often upwards. Medical costs in South Africa have not increased with general inflation over the last 15 years. Labour costs, which are often the largest contributor to the cost of an intervention, have generally increased at a higher rate than inflation- for example, the introduction of Occupational Specific Dispensation in 2007 marked a structural shift in health worker remuneration. Similarly, drug costs, also a significant contributor to total costs, do not increase each year with inflation as the costs are negotiated through a tender process and may be fixed for a number of years, and in the past the prices tended to decrease as a result of tender negotiations. For these reasons we would advise caution in applying straight-line inflation adjustments to the costs of health interventions in South Africa over a number of years.

In line with the impact ART has on the total cost of the HIV programme in South Africa, more than two thirds of the identified 42 papers detailed the cost of ART. However, in our review we were able to find publications on the costs of a number of other interventions such as HCT, MMC, palliative and inpatient care. The majority of the reviewed publications included all cost categories relevant to the intervention under study, and more than half had collected data through bottom-up resource use analyses, often using large patient samples. However, only 62% of papers reported some estimate of uncertainty, increasing the risk of potential reporting bias.

Our review has a number of other limitations. Firstly, as mentioned above, we used inflation adjustment for the results of the literature review, in order to be able to present mean and median cost in a single cost year to aid comparison (see \$\frac{S3 Table}{2}\$). For the results of our own ingredient cost analysis, however, we added the details of the exact ingredients, quantities and unit prices included for each intervention and technical efficiency factor in \$\frac{S4 Table}{2}\$ in order to allow other analysts to easily update our estimates to their settings and future prices. Secondly, our search strategy might have not retrieved all relevant papers, especially those reporting on results from smaller settings or using smaller sample sizes. Since we do not attempt to synthesize the evidence further, for example by creating a mean cost and cost range of a single intervention from several such reported estimates, we do not think that our analysis is subject to reporting bias.

In our review we were able to find literature detailing the results of primary cost analyses for five out of the 11 intervention categories included in the Investment Case, including for most of the HCT technical efficiency factors. The remaining interventions had to be costed using an ingredients-based approach, which is more prone to over- or underestimation than the detailed analysis of actual resources used in a bottom-up cost analysis. Only a small part of HIV services had to be costed based on expenditure. In this method, the relationship between inputs and outputs (in this case, services rendered) is even harder to establish, as invoices might have not been paid on time and the charges paid could have been higher than the actual cost of inputs. The quality of some of the results of this analysis is therefore limited; however, in each of these cases an estimate based on inferior methodology was deemed better than no estimate at all.

Our analysis yielded some insight into those interventions that need further cost analysis. In the area of HIV, more primary cost analyses are currently needed on the costs of PrEP, especially once routine delivery has started, social and behaviour change communication, and PMTCT. More work should also be directed to generate additional estimates of the cost of technical efficiency factors, including adherence interventions and different modalities of delivering HIV counselling and testing services, especially given the heightened demand for these factors.



Supporting information

S1 Table. Search strategies by intervention/ technical efficiency factor. (PDF)

S2 Table. Definitions of variables collected in systematic review.

(PDF)

S3 Table. Results of literature review.

(PDF)

S4 Table. Details of ingredient-based unit costs [2017/18 ZAR].

(PDF)

S1 File. PRISMA checklist.

(PDF)

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References

 Granich RM, Gilks CF, Dye C, De Cock KM, Williams BG (2009) Universal voluntary HIV testing with immediate antiretroviral therapy as a strategy for elimination of HIV transmission: a mathematical model. Lancet 373: 48–57. https://doi.org/10.1016/S0140-6736(08)61697-9 PMID: 19038438



- Hontelez JAC, de Vlas SJ, Tanser F, Bakker R, Bärnighausen T, Newell M-L, et al (2011) The Impact of the New WHO Antiretroviral Treatment Guidelines on HIV Epidemic Dynamics and Cost in South Africa. PLoS ONE 6:e21919 https://doi.org/10.1371/journal.pone.0021919 PMID: 21799755
- Walensky RP, Wood R, Ciaranello AL, Paltiel AD, Lorenzana SB, Anglaret X, et al (2010) Scaling Up the 2010 World Health Organization HIV Treatment Guidelines in Resource-Limited Settings: A Model-Based Analysis. PLoS Med 7(12): e1000382. https://doi.org/10.1371/journal.pmed.1000382 PMID: 21209794
- Bendavid E, Grant P, Talbot A, Owens DK, Zolopa A (2011) Cost-effectiveness of antiretroviral regimens in the World Health Organization's treatment guidelines: a South African analysis. AIDS 25: 211–220. https://doi.org/10.1097/QAD.0b013e328340fdf8 PMID: 21124202
- Ciaranello AL, Lockman S, Freedberg KA, Hughes M, Chu J, Currier J, et al (2011) First-line antiretroviral therapy after single-dose nevirapine exposure in South Africa: a cost-effectiveness analysis of the OCTANE trial. AIDS 25:479–492. https://doi.org/10.1097/QAD.0b013e3283428cbe PMID: 21293199
- Bachmann MO (2006) Effectiveness and cost effectiveness of early and late prevention of HIV/AIDS progression with antiretrovirals or antibiotics in Southern African adults. AIDS Care 18(2): 109–120. https://doi.org/10.1080/09540120500159334 PMID: 16338768
- Vijayaraghavan A, Efrusy MB, Mazonson PD, Ebrahim O, Sanne IM, Santas CC, et al (2007) Cost effectiveness of alternative strategies for initiating and monitoring highly active antiretroviral therapy in the developing world. J Acquir Immune Defic Syndr 46(1): 91–100. https://doi.org/10.1097/QAI. 0b013e3181342564 PMID: 17621241
- Bacaer N, Pretorius C, Auvert B (2010) An age-structured model for the potential impact of generalized access to antiretrovirals on the South African HIV epidemic. Bull Math Biol 72(8): 2180–98. https://doi.org/10.1007/s11538-010-9535-2 PMID: 20349152
- Cleary SM, McIntyre D, Boulle AM. The cost-effectiveness of antiretroviral treatment in Khayelitsha, South Africa—A primary data analysis. Cost Eff Resour Alloc. 2006; 4:1–14. https://doi.org/10.1186/ 1478-7547-4-1
- Smith de Cherif TK, Schoeman JH, Cleary S, Meintjes GA, Rebe K, Maartens G (2009) Early severe morbidity and resource utilization in South African adults on antiretroviral therapy. BMC Infect Dis 9: 205. https://doi.org/10.1186/1471-2334-9-205 PMID: 20003472
- Deghaye N., Pawinski RA, and Desmond C, Financial and economic costs of scaling up the provision of HAART to HIV-infected health care workers in KwaZulu-Natal. S Afr Med J, 2006. 96: p. 140–143. PMID: 16532083
- Foster N. and McIntyre D, Economic evaluation of task-shifting approaches to the dispensing of anti-retroviral therapy. Human Resources for Health, 2012. 10: p. 32. https://doi.org/10.1186/1478-4491-10-32 PMID: 22974373
- Granich RM, Gilks CF, Dye C, De Cock KM, Williams BG (2009) Universal voluntary HIV testing with immediate antiretroviral therapy as a strategy for elimination of HIV transmission: a mathematical model. Lancet 373(9657): 48–57. https://doi.org/10.1016/S0140-6736(08)61697-9 PMID: 19038438
- 14. Harling G, Wood R (2007) The Evolving Cost of HIV in South Africa: Changes in Health Care Cost With Duration on Antiretroviral Therapy for Public Sector Patients. J Acquir Immune Defic Syndr 45:348–354. https://doi.org/10.1097/QAI.0b013e3180691115 PMID: 17496562
- 15. Hontelez JAC, de Vlas SJ, Tanser F, Bakker R, Bärnighausen T, Newell M-L, et al. (2011) The Impact of the New WHO Antiretroviral Treatment Guidelines on HIV Epidemic Dynamics and Cost in South Africa. PLoS ONE 6(7): e21919. https://doi.org/10.1371/journal.pone.0021919 PMID: 21799755
- Kevany S, Meintjes G, Rebe K, Maartens G, Cleary S (2009) Clinical and financial burdens of secondary level care in a public sector antiretroviral roll-out setting (G F Jooste Hospital). S Afr Med J 99:320– 325. PMID: 19588792
- Leisegang R, Cleary S, Hislop M, Davidse A, Regensberg L, Little F, et al. (2009) Early and Late Direct Costs in a Southern African Antiretroviral Treatment Programme: A Retrospective Cohort Analysis. PLoS Med 6(12): e1000189. https://doi.org/10.1371/journal.pmed.1000189 PMID: 19956658
- Long LC, Rosen SB, Brennan A, Moyo F, Sauls C, Evans D, et al. Treatment outcomes and costs of providing antiretroviral therapy at a primary health clinic versus a hospital-based hiv clinic in south africa. PLoS One [Internet]. 2016; 11(12):1–12. Available from: http://dx.doi.org/10.1371/journal.pone.0168118
- Long LC, Fox MP, Sauls C, Evans D, Sanne I, Rosen SB. The high cost of HIV-positive inpatient care at an urban hospital in Johannesburg, South Africa. PLoS One [Internet]. 2016; 11(2):1–12. Available from: http://dx.doi.org/10.1371/journal.pone.0148546.
- Martinson N, Mohapi L, Bakos D, Gray GE, McIntyre JA, Holmes CB. Costs of Providing Care for HIV-Infected Adults in an Urban HIV Clinic in Soweto, South Africa. JAIDS J Acquir Immune Defic Syndr. 2009; 50(3):327–30. https://doi.org/10.1097/QAI.0b013e3181958546 PMID: 19194308



- Meyer-Rath G, Brennan AT, Fox MP, Modisenyane T, Tshabangu N, Mohapi L, et al. Rates and cost of hospitalization before and after initiation of antiretroviral therapy in urban and rural settings in South Africa. J Acquir Immune Defic Syndr. 2013; 62(3)
- 22. Meyer-Rath G, Brennan A, Long L, Ndibongo B, Technau K, Moultrie H, et al. Cost and outcomes of paediatric antiretroviral treatment in South Africa. AIDS. 2013; 27(2)
- Meyer-Rath G, Miners A, Santos AC, Variava E, Venter WD. Cost and resource use of patients on antiretroviral therapy in the urban and semiurban public sectors of South Africa. J Acquir Immune Defic Syndr. 2012; 61(3).
- Rosen S, Ketlhapile M, Sanne I, DeSilva MB (2007) Cost to patients of obtaining treatment for HIV/ AIDS in South Africa. S Afr Med J 97:524–529. PMID: 17805455
- 25. Thomas LS, Manning A, Holmes CB, Naidoo S, van der Linde F, Gray GE, et al (2007) Comparative Costs of Inpatient Care for HIV-Infected and Uninfected Children and Adults in Soweto, South Africa. J Acquir Immune Defic Syndr 46(4):410–416. https://doi.org/10.1097/QAI.0b013e318156ec90 PMID: 17786130
- Department of Health, South Africa, and South African National AIDS Council: South African HIV and TB Investment Case—Reference Report Phase 1. March 2016.
- Schwartländer B, Stover J, Hallett T, Atun R, Avila C, Gousw E, et al (2011) Towards an improved investment approach for an effective response to HIV/AIDS. Lancet.; 377(9782):2031–41. https://doi.org/10.1016/S0140-6736(11)60702-2 PMID: 21641026
- 28. Statistics South Africa: Consumer Price Index: Index numbers and year-on-year rates. http://www.statssa.gov.za/?page_id=1854&PPN=P0141 (last accessed 19 June 2018)
- 29. Leisegang R, Cleary S, Hislop M, Davidse A, Regensberg L, Little F, et al. (2009) Early and Late Direct Costs in a Southern African Antiretroviral Treatment Programme: A Retrospective Cohort Analysis. PLoS Med 6(12): e1000189. https://doi.org/10.1371/journal.pmed.1000189 PMID: 19956658
- Sinha A, Kim S, Ginsberg G, Franklin H, Kohberger R, Strutton D, et al. Economic burden of acute lower respiratory tract infection in South African children. Paediatr Int Child Health. 2012; 32(2):65–73. 0 https://doi.org/10.1179/2046905512Y.0000000010 PMID: 22595212
- 31. Bango F, Ashmore J, Wilkinson L, van Cutsem G, Cleary S. Adherence clubs for long-term provision of antiretroviral therapy: cost-effectiveness and access analysis from Khayelitsha, South Africa. Trop Med Int Heal. 2016; 21(9):1115–23.
- Deghaye N., Pawinski R.A., and Desmond C., Financial and economic costs of scaling up the provision of HAART to HIV-infected health care workers in KwaZulu-Natal. S Afr Med J, 2006. 96: p. 140–143. PMID: 16532083
- Chimbindi N, Bor J, Newell ML, Tanser F, Baltussen R, Hontelez J, et al. Time and Money: The True Costs of Health Care Utilization for Patients Receiving "Free" HIV/Tuberculosis Care and Treatment in Rural KwaZulu-Natal. J Acquir Immune Defic Syndr. 2015; 70(2):e52–60. https://doi.org/10.1097/QAL. 0000000000000728 PMID: 26371611
- Smith JA, Sharma M, Levin C, Baeten JM, van Rooyen H, Celum C, et al. (2015). Cost-effectiveness of community-based strategies to strengthen the continuum of HIV care in rural South Africa: a health economic modelling analysis. The Lancet HIV 2(4): e159–e168. https://doi.org/10.1016/S2352-3018(15) 00016-8 PMID: 25844394
- **35.** Cobb G, Bland RM. Nutritional supplementation: The additional costs of managing children infected with HIV in resource-constrained settings. Trop Med Int Heal. 2013; 18(1):45–52.
- **36.** Tagar E, Sundaram M, Condliffe K, Matatiyo B, Chimbwira F, Chilima B, et al. Multi-Country analysis of treatment costs for HIV/AIDS (match): Facility-level art unit cost analysis in Ethiopia, Malawi, Rwanda, South Africa and Zambia. PLoS One. 2014; 9(11).
- 37. de Tolly K, Skinner D, Nembaware V, Benjamin P. Investigation into the Use of Short Message Services to Expand Uptake of Human Immunodeficiency Virus Testing, and Whether Content and Dosage Have Impact. Telemed e-Health. 2012; 18(1):18–23.
- 38. Tabana H, Nkonki L, Hongoro C, Doherty T, Ekström AM, Naik R, et al. A cost-effectiveness analysis of a home-based HIV counselling and testing intervention versus the standard (facility based) HIV testing strategy in rural South Africa. PLoS One. 2015; 10(8):1–13. http://dx.doi.org/10.1371/journal.pone.0135048
- Barton GR, Fairall L, Bachmann MO, Uebel K, Timmerman V, Lombard C, et al. Cost-effectiveness of nurse-led versus doctor-led antiretroviral treatment in South Africa: Pragmatic cluster randomised trial. Trop Med Int Heal. 2013; 18(6):769–77.
- 40. Hongoro C, Dinat N. A cost analysis of a hospital-based palliative care outreach program: Implications for expanding public sector palliative care in South Africa. J Pain Symptom Manage. 2011; 41(6):1015–24. https://doi.org/10.1016/j.jpainsymman.2010.08.014 PMID: 21330096



- Larson B Schnippel K, Ndibongo B, Long L, Fox MP, Rosen S (2012). How to estimate the cost of point-of-care CD4 testing in program settings: an example using the Alere Pima Analyzer in South Africa.
 PLoS One 7(4): e35444. https://doi.org/10.1371/journal.pone.0035444 PMID: 22532854
- 42. Gavaza P, Rascati KL, Oladapo AO, Khoza S: The State of Health Economic Research in South Africa —A Systematic Review. Pharmacoeconomics 2012; 30 (10): 925–940 https://doi.org/10.2165/ 11589450-000000000-00000 PMID: 22809450
- Beck JE, Harling G, Gerbase S, DeLay P: The cost of treatment and care for people living with HIV infection: implications of published studies, 1999–2008. Curr Op HIV AIDS 2010; 5: 215–24
- 44. Beck JE, Avila C, Gerbase S, Harling G, DeLay P: Counting the cost of not costing HIV health facilities accurately—Pay now, or pay more later. Pharmacoeconomics 2012; 30 (10): 887–902 https://doi.org/10.2165/11596500-000000000-00000 PMID: 22830633
- 45. Galarraga O, Wirtz VJ, Figueroa-Lara A, Santa-Ana-Tellez Y, Coulibaly I, Viisainen K, et al: Unit Costs for Delivery of Antiretroviral Treatment and Prevention of Mother-to-Child Transmission of HIV. A Systematic Review for Low- and Middle-Income Countries. Pharmacoeconomics 2011; 29 (7): 579–599 https://doi.org/10.2165/11586120-000000000-00000 PMID: 21671687
- 46. Siapka M, Remme M, Obure D, Maier CB, Dehne L, Vassall A. Is there scope for cost savings and efficiency gains in HIV services? A systematic review of the evidence from low- and middle-income countries. Bull Heal Organ. 2014; 4(April):499–511.
- Akudibillah G, Pandey A, Medlock J: Maximizing the benefits of ART and PrEP in resource-limited settings (2017) Epidemiol Infect. 145(5): 942–956. https://doi.org/10.1017/S0950268816002958 PMID: 28031062
- 48. Walensky RP, Jacobsen MM, Bekker L-G, Parker RA, Wood R, Resch SC, et al: Potential Clinical and Economic Value of Long-Acting Preexposure Prophylaxis for South African Women at High-Risk for HIV Infection (2016) J Inf Dis 213: 1523–31
- **49.** Francke JA, Penazzato M, Hou T, Abrams EJ, MacLean RL, Myer L, et al: Clinical Impact and Cost-effectiveness of Diagnosing HIV Infection During Early Infancy in South Africa: Test Timing and Frequency (2016) J Inf Dis 214:1319–28
- 50. Meyer-Rath G, Johnson LF, Pillay Y, Blecher M, Brennan AT, Long L, ET AL: Changing the South African national antiretroviral treatment guidelines: The role of cost modelling. PLoS One 12 (10), e0186557. (2017) https://doi.org/10.1371/journal.pone.0186557 PMID: 29084275
- **51.** Tchuenche M, Haté V, McPherson D, Palmer E, Thambinayagam A, Loykissoonlal D, et al. Estimating client out-of-pocket costs for accessing voluntary medical male circumcision in South Africa. PLoS One. 2016; 11(10):1–10.
- Callegari LC, Harper C, van der Straten A, Kamba M, Chipato T and Padian NS (2008). Consistent condom use in married Zimbabwean women after a condom intervention. Sex Transm Dis 35(6): 624–630 https://doi.org/10.1097/OLQ.0b013e31816b3208 PMID: 18545141
- 53. Meyer-Rath G., Brennan AT, Fox MP, Modisenyane T, Tshabangu N, Mohapi L, et al (2013). Rates and cost of hospitalization before and after initiation of antiretroviral therapy in urban and rural settings in South Africa. J Acquir Immune Defic Syndr 62(3): 322–328. https://doi.org/10.1097/QAI. 0b013e31827e8785 PMID: 23187948
- 54. Bassett IV, Govindasamy D, Erlwanger AS, Hyle EP, Kranzer K, van Schaik N, ET AL (2014). Mobile HIV screening in Cape Town, South Africa: clinical impact, cost and cost-effectiveness. PLoS One 9 (1): e85197. https://doi.org/10.1371/journal.pone.0085197 PMID: 24465503

Appendix S3: Results of literature review

Appendix for the paper "The per-patient costs of HIV services in South Africa: Systematic review and application in the South African HIV Investment Case" by Gesine Meyer-Rath, Craig van Rensburg, Calvin Chiu, Rahma Leuner and Steve Cohen

No.	Author, year 1. Cotrimox	Objective	Target population, setting, level of care	Cost perspective	Costing method	Costing period	Mean cost (2016 USD)	Range (2016 USD)	ARVs	Other drugs	Labs	Radiology	Staff/Consultation	Consumables	Overhead ¹	Equipment	Iransport Inpatient	Other
1	Hausler (2006)	Cost and effectiveness of a package of TB and HIV interventions	General population in Cape Town; CHC, PHC and STI clinic	Provider	Full economic cost, ingredients approach	2001 - 2002	Screening for cotrimoxazole preventative therapy \$3.13 in community health centre \$1.57 primary health clinic Cost of screening per person started on cotrimoxale preventative therapy \$7.84 in community health centres \$2.97 in primary health clinic Cost of 6 months of cotrimoxazole preventative therapy \$12.54 community health centre \$9.40 primary health clinic	N/S		x			x	x	x	x		

¹ includes other capital costs in some instances

3	Chimbindi (2015)	Cost associated with accessing public sector care over different stages of illness (TB, pre-ART, ART)	General population in rural KwaZulu- Natal, PHC	Patient	Economic and financial costs, bottom up approach	2009-2010	Financial costs per month \$18.61 pre-ART \$17.84 on ART Monetized time costs \$6.45 pre-ART \$9.49 on ART	All SD \$14.61-22.56 \$15.37-20.38 \$5.07-7.84 \$4.11-14.88		x		x			x	Opportunity costs
6	(2012)	Explore barriers to access of ART at different treatment sites	Adult population (>18) in rural and urban treatment sites	Patient	Full financial costs, bottom up	N/S	Expenditure past month \$6.43- to reach ART facility (rural) \$18.34- all healthcare (rural) \$1.41- reach ART facility (Urban) \$5.77- all healthcare (Urban)	N/S		x					х	Patient health expenses
7	Cobb (2013)	Cost of applying WHO guidelines for the nutritional management of HIV-infected children		Provider	Incremental financial costs, ingredients	2010	\$56.25 /child nutrition, 26 weeks, requiring NCP-B only \$178.33 / child nutrition, 26 weeks requiring NCP-C and NCP-B \$923.63 average cost of supplying ART	N/S	x							Nutritional supplement
9	Deghaye (2006)	Provide comprehensi ve costing of HAART to health care workers in different hospitals	Health Care Workers	Provider	Full Economic and incremental financial costs, bottom up	2004	Total financial cost per year \$1379.18 urban \$896.74 peri-urban Total economic cost per year \$1399.80 urban \$1066.89 peri-urban	N/S	х	x	x	х	х	x		

11	Harling (2007a)	Cost of a dedicated ART clinic over pre-ART and 2 years of ART	General population	Provider	Full financial costs, top down	March 2004- Feb 2006	Cost per patient month \$40.81 in 2004/05 \$36.94 in 2005/06 Cost per visit \$55.49 in 2004/05 \$42.16 in 2005/06	N/S	х	x				x	x	х	
16	Leisegang (2009)	Describe the direct costs and establish cost drivers over time in HIV managed care programme	19+ yrs old, ART naïve	Provider	Incremental financial costs, ingredients using AfA data base	N/S	Monthly cost \$96.96 before ART \$484.79 peri-ART \$193.92 on ART	N/S	x	x							
18	Long (2010)	Estimate the costs and outcomes of second line ART	Adult patients on second line treatment	Provider	Full financial costs, bottom up	N/S	Cost 12 months after initiating \$1045.39 all outcomes \$1278.26 care and responding \$1122.01 t in care and not responding \$482.88 no longer in care	N/S	х	x	X		х	x	x		
19	Martinson (2009)	Estimate the cost of initiating and maintaining patients on ART	Adults	Provider	Full financial costs, bottom up	Octobe r 2004- March 2005	Monthly costs \$96.48 overall per patient \$61.77 per patient not on ART \$117. per patient on ART \$105.24 for month prior to ART \$206.25 month of ART initiation \$127.69 after ART initiation		х	x	x	х	х			х	
21	Meyer-Rath (2013)	Annalise ART cost	Children (initiated ART, 13 yrs old)	Provider	Full financial costs, bottom up	2005-2009	Per 24 month outcome \$723.63 all in care \$652.11 all in sample \$304.88 not in care \$758.45 in care not responding \$707.63 in care responding		х	х	х	х	х	х	х		

22	Moshabela (2012)	Investigate factors associated with patterns of plural healthcare utilization	Adults	Patient	Incremental financial costs, bottom up	April 2008- March 2009	Report expenditure 4 weeks prior to ART visit \$5.65 private chemist \$8.47 self-care \$17.88 doctor \$6.59 primary health clinic	\$2.82-12.23 \$2.82-23.52 17.88-28.23 0.94-2.82 IQR		x		x					
25	Rosen (2007)	Estimate the costs patient occur in obtaining ART	Adults	Patient	Full financial and economic costs, bottom up	July 2005- June 2006	Total cost per visiit \$15.83 urban hospital \$10.20 informal settlement clinic \$18.88 rural clinic \$14.62 all					х				х	Opportunity cost
26	Rosen (2008)	Estimate outpatient cost per patient in care and responding to treatment	General population	Provider	Full financial costs, bottom up	N/S	Annual cost per patient initiated: \$747.59 referral hospital \$886.04 private GP \$921.64 HIV clinic \$1113.48 NGO clinic \$917.68 all sites Cost to produce patient in care and responding: \$115.46 referral hospital \$1703.84 private GP \$1463.54 HIV clinic \$1465.52 NGO clinic \$1422.01 all sites	N/A	х	x	х	х	X	X	x		
27	Tagar (2014)	Describe facility-level cost of ART in a random sample of facilities	General population	Provider	Full financial costs, top down	2011	Average cost of treatment per year \$521.46 weighted mean \$597.71 simple mean Average 1st line cost \$160 cost for adults		х	x	х	x	х	x	х		Nutritional

28	Vella	Identify	General	Provider	Full financial	April	6 month cycle cost:	6 month	Х	х	х	х	х	х	х	х		
	(2011)	which	population		costs, bottom	2005-	,	cycle (95%)										
		delivery			up	March	\$471.58 Part time MD part time nurse,	\$447.96-										
		profiles of			'	2006	<200 new patients per MD per yr	495.21										
		ART sites					\$484.38 Part time MD part time nurse,											
		influenced					>= 200 new patients per MD per yr	\$476.51-										
		retention and					\$509.98 full time MD + full time nurse,	492.26										
		at what cost					<200 new patients per MD per yr											
		patients were					\$531.64 full time MD + full time nurse,	\$502.10-										
		retained					>= 200 new patients per MD per yr	517.86										
							Incremental cost per patient on	\$506.04-										
							treatment after 10 years:	557.24										
							\$8864.60 Part time MD part time nurse,											
							<200 new patients per MD per yr											
							\$12081.02 full time MD + full time nurse,	ICERs (95%)										
							<200 new patients per MD per yr											
								\$8,369-9,367										
								\$10,430.97-										
								13,709										
29	Wouters	Investigate	General	Patient	Full financial	2004-		SD									х	
	(2010)	characteristic	population		costs, bottom	2006	\$1.22 mean transport cost	\$0-2.31										
		s of patients			up													
		enrolled in																
		public sector																
		ART																
		programme																
		in Free State																
		nd outpatient co				T			ı	ı	ı	ı	1					
5	Cleary	Cost and cost		Provider	Full economic	N/S	Cost per clinic visit		Х	х	Х	Х	Х	Х	х	Х		х
	(2006)	effectiveness	population in		cost,		\$22.11 ART											
		of HIV	Khayelitsha,		ingredients		\$21.64 no-ART											
		services, cost	Cape Town,		and step													
		per LY and	HIV clinic for		down		Incremental cost effectiveness											
		QALY gained	ART, tertiary		approach		\$1,187.25 per LY gained	\$984-1,187										
			and .					per LY gained										
			secondary facility for TB				\$1,329.62 per QALY	\$1,102-1,330 per QALY										
			Tacility IOI 1B					per QALI										

12	Harling (2007b)	Analyse ART costs over 3 treatment periods	General population	Provider	Full financial costs, top down	March 2004- Feb 2006	Pre ART costs \$409.19 intention to treat \$380.45 on program Treatment costs per patient year 1 \$2,533.71 intention to treat \$2045.73 on program Treatment costs per patient year 2 \$1,389.83 intention to treat \$1.392.74 on program	N/S	x	x	x			x	x	x	х
15	Kevany (2009)	Determine the costs of inpatient and outpatient at a dedicated ARV referral unit	Older than 13 yrs	Provider	Incremental economic costs, bottom up	Apr & May 2005	Outpatient costs \$194.87 cost per patient \$148.47 cost per visit Inpatient costs \$148 cost per inpatient \$488.49 cost per day	N/S	x	x	х	x	x				x
20	Meyer-Rath (2012)	Compare resource use and cost between 2 settings	General population, CD4 < 200	Provider	Full financial costs, bottom up	April 2006- Dec 2008	Inpatient costs per year \$304.99 urban \$366.34 semi-urban Outpatient costs per year \$606.48 urban \$610.86 semi-urban Total costs per year \$911.47 urban \$977.20 semi-urban	95% CI \$127.08- 482.03 \$70.99- 662.57 \$552.14- 660.81 \$589.82- 631.01 \$701.13- 1121.80 \$680.09- 1273.42	x	x	х	x	x	х	x	x	x

23	Nachega	Determine	Adults	Provider	Incremental	N/S	Mean monthly cost	SD	Х	Х	х	Х	х		х	
	(2010)	the effect of		(private)	financial		\$356.45 all patients	\$0-996.45								
		ART			costs, top		\$362.14 adherence 4-45%	\$214.52-								
		adherence on			down		\$402.15 adherence 40-75%	938.80								
		the direct					\$358.86 adherence 75-91%	\$400.61-								
		health care					\$302.26 adherence 92-100%	1,204.90								
		costs among						\$134.31-								
		adults					Median monthly cost	852.02								
							\$203.44 all patients	\$0- 935.90								
							\$202.47 adherence 4-45%									-
							\$170.66 adherence 40-75%	IQR								Other
							\$208.26 adherence 75-91%	\$147.52-								0
							\$215.97 adherence 92-100%	303.71								
								\$135.95-								
								323.96								
								\$102.20-								
								319.13								
								\$140.77-								
								345.17								
								\$151.37-								
								327.81								
24	Naidoo	Measure the	General	Provider	Incremental	2005-	Total median variable cost per patient	IQR	х	х	х	Х	Х	х	х	
	(2015)	effectiveness	population		financial	2010	month									
		of initiating			costs, bottom											
		ART at			up		\$100.57 with ART within 4 weeks	\$93.38-								
		different					\$96.96 ART after intensive phase of	142.62								
		points in TB					treatment	\$89.99-								
		treatment					\$86.59 ART after completing treatment	126.49								
								\$78.10-								
								113.76								
		1			1	<u> </u>				<u> </u>	<u> </u>	1				Ш_
	3. Inpatie	nt cost														

8	Smith de Cerif (2009)	Determine whether reasons for hospitalizatio n were different for HAART vs non-HAART	Adult population (>18), secondary hospital, Cape Town	Provider	Full financial costs, ingredients (some costs from Cleary 2006)	July 2003- March 2004	Total hospitalization cost \$1513.91 in HAART group \$1401.09 in NON-HAART group	IQR \$777.91- 2855.9 \$873.53- 2231.64	x	x	x	х	x	x	x	x	
42	Meyer-Rath (2013)	Compare hospitilisatio n rates and costs in cohort of HIV+ patients before and after ART initiation	Adult population (over 18 years)	Provider	Full financial costs, ingredients approach	2009	Cost per patient per year, pre ART \$142.21 at 100 cells/mm³ \$74.34 at 101 – 200 cells/mm³ \$56.56 at 201 – 350 cells/mm³ \$35.55 at >350 cells/mm³ Cost per patient per year, on ART \$330.49 at 100 cells/mm³ \$74.34 at 101 – 200 cells/mm³ \$56.56 at 201 – 350 cells/mm³ \$35.55 at >350 cells/mm³	95% CI \$76.76- 234.33 \$39.59- 122.82 \$56.56-83.23 \$20.20-54.14 \$120.40- 694.91 \$75.96- 327.25 \$58.18- 139.79 \$18.58-64.64									

43	Sinha	Determine	HIV+ and HIV	Provider	Full financial	2000 –	HIV+ costs, cost per ALRTI episode	95% CI	Χ	Х	х	х		х	Х	
	(2012)	the economic	- Children (>6	and patient	costs, top	2001	\$969.61, direct paediatric ward costs,	\$4874.53-								
		burden of	months)		down for bed		(retrospective period)	1,064.44								
		acute lower			days, bottom		\$1,117.10 direct paediatric ward cost	\$892.08-								
		respiratory			up for other		(prospective period)	1342.11								
		tract			resource use		\$198.28 short stay ward costs									
		infection					(prospective period)	\$184.32-								
		(ALRTI) in					\$11.88 patient costs	193.90								
		South Africa														ty
							HIV- costs, cost per ALRTI episode	\$8.78-16.76								Onportunity cost
							\$804.31, direct paediatric ward costs,									Ė
							(retrospective period)									Ę
							\$809.10.10 direct paediatric ward cost	\$711.75-								٥
							(prospective period)	896.07								Č
							\$193.90 short stay ward costs									
							(prospective period)	\$646.32-								
							\$10.37 patient costs	971.88								
								\$184.32-								
								212.25								
								212.25								
								\$8.78-16.79								
48	Long (2016)	Report the	Adult (>18)	Provider	Full financial	2012-	Cost per HIV+ admission	N/S	х	х	х	х	Х	х	Х	
		primary			costs, bottom	2013	\$3,518 renal disease									
		reasons for			up		\$2,298 drug adverse event									
		HIV-related					\$2,205 endocrine and metabolic disease									
		admissions at					\$2,172 HIV and related OI									
		a regional,					\$2,135 gastro-intestinal disease									
		urban					\$2,096 unknown reason for admission									
		hospital					\$1,818 hematologic disorder									
							\$1.719 parasitic infection									
							\$1,714 TB and other mycobacteria									
							\$1,677 musculo-skeletal									
							\$1,250 other									
							\$1,590 all									

30	Barton (2013)	Estimate cost effectiveness of nurse-led ART vs doctor lead ART	population (16+)	Provider	Full financial costs, bottom up	2008-2010	Cost at 12 months post enrollment, patients CD4 <=350 and not yet on ART \$383.05 nurse lead ART \$285.70 doctor lead ART \$23,444 per death averted Costs at 12 months post enrollment, Patients on ART at enrollment 460.68 nurse lead ART 402.91 doctor lead ART 12,042 per undetectable viral load	95% CI \$17,529- 19,625	х	x	x	x	x			x	Set up costs
31	Foster (2012)	Evaluate indirectly supervised pharmacists' assistant (ISPA) and nurse based care against full time pharmacist	General population	Societal	Incremental financial and economic costs, bottom up	2009	Average annual costs: \$109.67 per patient with full time pharmacist (baseline) \$10.68 incremental cost for patient through ISPA \$27.11 incremental cost per patient through nurse Average societal cost per visit: \$14.10 with full time pharmacist (baseline) -\$2.89 incremental cost with ISPA -\$0.11 incremental cost with nurse Average cost to patient per visit \$8.54 with full time pharmacist (baseline) -\$2.21 incremental cost with ISPA -\$3.19 incremental cost with nurse Average cost to patient per visit \$5.57 with full time pharmacist (baseline) -\$0.69 incremental cost with ISPA \$3.07 incremental cost with nurse	N/S			x			×	X		Indirect patient costs

32	Long (2011)	Evaluate cost effectiveness		Provider	Full financial costs, bottom	N/S	Cost per patient:	Cost per	х	х	х	:	(x	х	х	
		of down	patients on ART > 11		up		514.55 DR/Hospital	patient (SD) \$379.95-								
		referring	months		ир		314.33 Div/103pital	649.15								
		from doctor					463.95 Down referral	0.5.25								
		managed to						\$70.40-								
		nurse						557.51								
		managed ART					Cost per patient in care and responding:									
		in primary					526.01 DR/Hospital									
		health care					469.98 Down referral	Cost in care								
								(SD)								
								\$403.81-								
								648.20								
								\$385.67-								
		_						553.69					_			 _
45	Fatti	Compare	ART naïve	Provider	Incremental	Jan	Mean staff cost per visit	N/S				×				
	(2016)	•	adults, 16 yrs		financial	2013-	\$130 through ISPA									
		al care	and older		costs, bottom		1.82 through nurse managed									
		quality, clinical			up	2013	Staff cost per item dispensed									
		outcomes					\$0.41 through ISPA									
		and provider					\$0.81 through nurse managed									
		staff costs					70.01 through harse managed									
		between task														
		shifting														
		pharmaceutic														
		al care														
		models														

53	Long (2016)	Compare outcomes and costs of NIMART between primary health clinics and hospital based HIV clinics	Adults >18yr	Provider	Full financial costs, bottom up	2014	Average cost for all patients: 323.74 HIV outpatient 201.63 PHC Average cost per patient alive & in care 405.15 HIV outpatient 225.29 PHC Average cost per dead/LTFU 115.49 HIV outpatient 94.66 PHC	All patients (95%) 308.60-338.89 193.11-210.15 Alive & in care 399.47-427.87 220.56-230.97 Dead/LTFU 100.34-129.69	x	x	х	x	x	x	x		
	4. SMS not	ification						82.36-106.97							<u> </u>		
			T	T	T	1	1	1 4			1	1		-		т г	
33	De Tolly (2012)	Investigate effectiveness of SMS to encourage HCT	General population	Provider	Full financial costs, bottom up	N/S (?2011)	\$0.02 per sms \$1.76 per additional tester	N/S									Cost per sms
	5. Adherer	nce clubs															

34	Bango (2016)	Assess cost effectiveness of lay health worker led group adherence clubs and describe and evaluate the associated patient costs	Adult, stable patients	Provider and patient costs	Full financial costs, top down	2011	Total cost per visit: \$12.97 adherence club \$14.16 standard of care Patient transport costs \$0.71 adherence club \$1.15 standard of care Total cost per patient yr \$235.45 adherence club \$293.53 standard of care	N/S	x		x	x	x	x	x	x	Patient opportunity costs
36	Basset (2007)	Evaluate the yield of routine VCT vs provider referred VCT	General population	Provider	Full financial costs, top down	Jan – Mar 2005	Cost per test \$9 referral \$8.56 routine VCT Cost per HIV positive person identified \$13.48 referral \$25.81 routine VCT	N/S				x	x	x			
1	Hausler (2006)	Cost and effectiveness of a package of TB and HIV interventions	General population in Cape Town; CHC, PHC and STI clinic	Provider	Full economic cost, ingredients approach	2001 - 2002	Cost per VCT completed \$14.10 at community care center \$17.24 at primary health clinic \$10.97 at STI clinic Cost per HIV infection averted \$145.74 at community health center \$175.51 at primary health clinic \$104.99 at STI clinic	N/S		х		x	x	x	x		

37	McConnel (2005)	Determine the cost of a client completing VCT through rapid test technology	General population	Provider	Full economic and financial costs, top down	June 2002 – May 2003	Cost per counselling and test \$116.45 economic cost \$68.85 financial cost Cost per completed VCT sequence \$121.89 economic cost \$72.07 financial cost	N/S				x x	×	x			
	7. Home H	IV counselling a	nd testing		1	1	I			1				<u>l</u>	ļ		
39	Smith (2015)	Evaluate impact and cost effectiveness of a community based home HTC package	Adult population (over 18 years)	Provider	Full financial costs, bottom up and ingredients	N/S	Cost per HIV negative test \$20.52 research model \$8.67 operation model Cost per HIV positive test \$35.29 research model \$23.28 operational model Incremental cost effectiveness \$21,920 per HIV infection averted \$1,335 per DALY averted	90% of model variability \$11,300-\$75,900 \$1080-1760	Х		x	×	x	x	x	x	End of life care
40	Tabana (2015)	Examine the cost effectiveness of home based testing in rural SA	General population	Provider	Full financial costs, ingredients approach	Jan – Dec 2010	\$23.08 cost per home based client \$30.24 cost per clinic client	N/S				x	×	x	x		Field materials

41	Hongoro	Establish cost	General	Provider	Full financial	2007/	Cost per home visit	N/S	Х	Х	Х	Х		Х	\top	
	(2011)	and cost	population		costs, top	2008	\$77.51 not including capital costs	'								
		drivers of a			down		\$109.87 including capital costs									
		hospital														es
		palliative					Cost per home visit registered patient									arg
		outreach					\$724.00 not including capital costs								-	5
		service					\$1,026.29 including capital costs								'	rest
							Cost per hospital visit									inte
							\$87.45 not including capital costs									aug
							\$119.82 including capital costs								-	Bank and interest charges
							Cost per hospital registered patient								(מ
							\$156.88 not including capital costs									
							\$214.94 including capital cost									
	9. Point o	f care CD4 testin	g									•				
44	Larson	Estimate cost	N/A	Provider	Full financial	2010	\$19.42 cost per test	N/S		Х	х		х			┪
	(2012)	of CD4 point			costs, bottom											
		of care			up		\$20.02 cost per successfully completed									یب
		testing					test									SO
																Insurance cost
																rar
																nsn
															•	=
	10. Volun	tary medical ma	le circumcision	(VMMC)												

46	Tchuenche (2016a)	Derive the unit cost of delivering VMMC at the facility level and to identify the level of spending currently incurred for VMMC demand creation	Men and boys	Provider	Full financial costs, ingredients	Jan-Dec 2014	Per circumcision performed \$124.95 across all facilities \$131.11 in facilities with outreach services \$123.15 in facilities without outreach services \$110.75 urban facilities \$137.26 peri-urban facilities \$111.70 rural facilities \$145.78 in hospitals \$114.54 health care centre/clinic	\$116.24- 145.97 \$115.37- 130.94 \$136.87- 154.69 \$105.97-	x	***************************************	×	x	x	x	×		
47	Tchuenche (2016b)	To provide useful information on the scope of financial barriers that VMMC clients face	Follow up VMMC clients	Patient	Full economic costs, bottom up	-	Average transport costs \$8.26 all clients \$9.43 urban clients \$7.45 peri-urban clients \$9.70 rural clients	123.10 N/S	×	(х	Opportunity costs

IQR – Interquartile range

CI – Confidence interval

ISPA – Indirectly supervised pharmacist assistant

Appendix S4: Details of ingredient-based unit costs [2017/18 ZAR]

Appendix for the paper "The per-patient costs of HIV services in South Africa: Systematic review and application in the South African HIV Investment Case" by Gesine Meyer-Rath, Craig van Rensburg, Calvin Chiu, Rahma Leuner and Steve Cohen

NB: For updates or additional details regarding unit costs based on ingredients, please contact the corresponding author at gesine@bu.edu; for updates regarding unit costs based on literature, please contact the corresponding author of the original paper.

1. PMTCT

Intervention	PMTCT (m	other not or	n ART, presei	nting in lab	our)	
Costing method			are drug costs and postnatal		ned to be inc	remental to
Total cost				R 316.6	5 per mothe	r-baby pair
Ingredient	Unit cost	Cost unit	Source of unit cost	Quantity	Source of ingredient and/ or quantity	Total cost
Single dose NVP + AZT 3-hourly in labour; TDF+FTC stat	R 5.20	per child	2018 drug tender prices	1		R 5.20
NVP coverage during breastfeeding (12 months)	R 311.45	per child	2018 drug tender prices	1		R 311.45

Intervention:	PMTCT (mot	her not on li	felong ART)			
Costing method:	Ingredients (N standard ante			nly; assume	d to be inc	remental to
Total cost				R 2,125.74	per moth	er-baby pair
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost
NVP coverage during breastfeeding (6 weeks)	R 36.31	per child	2018 drug tender prices	1		R 36.31
Triple ART during pregnancy and breastfeeding	R 2,089.43	per mother	2018 drug tender prices	1		R 2,089.43

Intervention:	Infant testi	ng at birth				
Costing method:	Ingredients					
Total cost					R 442	2.23 per test
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost
Staff Nurse	R 2.56	per minute of staff time	2018 Government Salary Scale	20	ZoeLife	R 51.28
Counsellor	R 0.99	per minute of staff time		10	ZoeLife	R 9.92
HIV PCR	R 380.02	per test	NHLS prices March 2017	1	ZoeLife	R 380.02
Gloves	R 50.82	per pack of 50 pairs	Treasury contract RT76-2016	0.02	Own assump tion	R 1.02

Intervention:	Infant test	ing at six we	eks			
Costing method:	Ingredients					
Total cost					R 416.	59 per test
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total
						cost
Staff Nurse	R 2.56	per minute of staff time	2018 Government Salary Scale	10	ZoeLife	R 25.64
Counsellor	R 0.99	per minute of staff time		10	ZoeLife	R 9.92
HIV PCR	R 380.02	per test	NHLS prices March 2017	1	ZoeLife	R 380.02
Gloves	R 50.82	per pack of 50 pairs	Treasury contract RT76-2016	0.02	Own assumption	R 1.02

2. Social behaviour change communication (SBCC)

Intervention	SBCC campaign 1 (message: adolescent testing, multiple partners)				
Costing method	Expenditure (personal communication, implementing agencies)				
Total cost	R 12.10 per person reached				

Intervention	SBCC campaign 2 (message: condom usage)				
Costing method	Expenditure (personal communication, implementing				
	agencies)				
Total cost	R 6.19 per person reached				

Intervention	SBCC campaign 3 (message: testing, condom usage)				
Costing method	Expenditure (personal communication, implementing agencies)				
Total cost	R 3.39 per person reached				

3. Comprehensive condom programming

Intervention:	Condom o	listribution				
Costing method:	Ingredients	3				
Total cost				R 0.74	per condom	distributed
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total
						cost
Male condom +	R 0.63	per	2015-18	0.98	2015-18	R 0.62
distribution		condom	Condom		Condom	
			tenders		tenders	
Female condom +	R 6.16	per	2015-18	0.021	2015-18	R 0.13
distribution		condom	Condom		Condom	
			tenders		tenders	

¹ These represent the relative quantities of male and female condoms planned in the 2012 condom tender.

Intervention:	Male and female condom education						
Costing method:	Literature, updated from [1]						
Total cost				R 66.	06 per pers	on trained	
Ingredient	Unit	Cost unit	Source	Quantity	Source	Total	
	cost					cost	
Male condom	R 0.52	per condom	2015-18 Condom tenders	1	[1]	R 0.52	
Female condom	R 6.14	per condom	2015-18 Condom tenders	1	[1]	R 6.14	
Counsellor (first session)	R 0.99	per minute of	2018 Government Salary Scale	30	[1]	R 29.70	
Counsellor (follow-up session)	R 0.99	staff time		30	[1]	R 29.70	

4. Care and treatment

Intervention:	ART							
Costing method:	National ART Cost Model (NACM) 2017/18							
Total cost	R 3,318.62 per patient year (all adults)							
	R 2,760.87 per adult on first line treatment							
	R 2,771.13 per adult in first line treatment faile							
	R 8,705.15 per adult on second line							
	R 9,272.15 per adult in second line treatment failure							
	R 18,688.12 per adult on third line ART							
	R 3,784.19 per patient year (children)							
	R 3,720.73 per child on first line treatment							
	R 4,466.98 per child in first line treatment failure							
	R 4,763.82 per child on second line ART							

Intervention:	Palliative care
Costing method:	Personal communication, A. Lolliot
Total cost	R 948.48 per patient

Intervention:	Inpatient care						
Costing method:	Updated from literature [2], using different CD4 strata						
Total cost	R 1,694.94 per patient year (pre-ART, <200 cells/microl)						
	R 989.47 per patient year (pre-ART, 200-349 cells/microl)						
	R 808.77 per patient year (pre-ART, 350-500 cells/microl)						
	R 393.23 per patient year (pre-ART, >500 cells/microl)						
	R 2,079.97 per patient year (ART, <200 cells/microl)						
	R 989.47 per patient year (ART, 200-349 cells/microl)						
	R 808.77per patient year (ART, 350-500 cells/microl)						
	R 393.23 per patient year (ART, >500 cells/microl)						

5. Medical male circumcision

Intervention:	Medical male circumcision
Costing method:	Literature [3]
Total cost	R 1,770.29 per circumcision

Intervention:	Early infant male circumcision
Costing method:	Ingredients
Total cost	R 885.14 per circumcision (assumed to be 50% of adult circumcision)

6. HIV counselling and testing (HCT)

Intervention:	General po	pulation HC	T (negative re	sult)			
Costing method:							
Total cost		T	T	1		24 per test	
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost	
Rent and electricity	R 3,181.95	per month	Estimate	0.001	Assumed	R 3.18	
Security guard	R 0.55	per minute of staff time	2018 Government Salary Scale	0.5	[9]	R 0.27	
Counsellor	R 0.99	per minute of staff time	2018 Government Salary Scale	17	[9]	R 16.86	
Clerk	R 1.29	per minute of staff time	2018 Government Salary Scale	4	[9]	R 5.17	
Furniture and equipment	R 1 178	rental cost per month	Retail quote	0.001	Assumed. Equipment supports 1,000 patients seen per month	R 1.18	
Paper flip file	R 5.00	per material	Retail quote	1	Assumed	R 5.00	
Bond paper	R 58.80	per 500 pages	Retail quote	0.004	Assumed 2 sheets used	R 0.36	
Biosscrub	R 155.18	Per 5L	Treasury contract	0.0002	Assumed	R 0.03	
Gloves	R 50.82	per pack (100)	Treasury contract RT76-2016	0.02	RSA GF proposal for continued funding, 2013	R 1.03	
Cotton wool swabs	R 81.65	per pack (100)	Treasury contract	0.01	RSA GF proposal for continued funding, 2013	R 0.82	
First response test kit	R 8.17	per test	Treasury contract RT41-2017	1.01	RSA GF proposal for continued funding, 2013	R8.25	
Biohazard bag	R 0.09	per bag	Treasury contract	1	RSA GF proposal for	R 0.09	

					continued funding, 2013	
Result card	R 0.76	per card	Retail quote	1.01	Assumptio n	R 0.77
Male condom	R 0.52	per condom	Treasury contract	10.01	Assumptio n	R 5.23

Intervention:	General population HCT (positive result)							
Costing method:	Update fro	Update from literature [4]						
Total cost	R 74,83 per							
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost		
As above, plus:								
Advance test kit	R 6.50	per test	Treasury contract RT41-2017	1.01	RSA GF proposal for continued funding, 2013	R 6.57		
First response test kit	R 8.17	per test	Treasury contract RT41-2017	0.007	Assumed	R 0.06		
Advance test kit	R 6.50	per test	Treasury contract RT41-2017	0.007	Assumed	R 0.05		
ELISA	R 55.18	per test	NHLS 2018 price list	0.0002	Assumed	R 0.01		
Counsellor	R 0.99	per minute	April 2018 Government Salary Scale		Assumed	R 19.90		

Intervention:	Provider-initiated HCT
Costing method:	Updated from literature [4]
Total cost	R 56.07 per test (negative result)
	R 82.66 per test (positive result)

Intervention:	Mobile HCT
Costing method:	Updated from literature [5]
Total cost	R 74.48 per test (negative result)
	R 87.92 per test (positive result)

Intervention:	Home-based HCT
Costing method:	Updated from literature [5]
Total cost	R 69.73 per test (negative result)
	R 77.42 per test (positive result)

Intervention:	HCT invitations to male pregnancy partners (negative result)						
Costing method:	Ingredients						
Total cost		R 50.79 per t					
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total	
						cost	
HCT invitation card	R 1.82	per card	Retail quote	1	Assumption	R 1.82	
BCC material design	R 5,600.00	per day	RSA Global Fund	0.1	Assumption	R 0.56	

BCC material	R1,238.18	per day	Retail quote	0.07	Assumption	R 0.17
translation						
General population	R 48.24	per test	See above	1		R 48.24
HCT (negative)						

Intervention:	HCT invitat	ions to male	e pregnancy p	artners (po	sitive result)	
Costing method:	Ingredients				-	
Total cost					R 79	.87per test
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total
						cost
HCT invitation card	R 1.82	per card	Retail quote	1	Assumption	R 1.82
BCC material design	R 5,600.00	per day	RSA Global	0.1	Assumption	R 0.56
			Fund			
BCC material	R1,238.18	per day	Retail quote	0.07	Assumption	R 0.17
translation						
General population	R 74,83	per test	See above	1		R 77.32
HCT (positive)						

Intervention:	Workplace	HCT (negati	ve result)					
Costing method:	Ingredients							
Total cost	R 55.50 per test							
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total		
						cost		
Mobile van	R 2,108.26	per month		0.001	Assumption	R 2.11		
Senior nurse			2018	0.001	Assumption	R 26.36		
			Government					
	R 26 360	per month	Salary Scale					
Community health			Global fund	0.001	Assumption	R 10.53		
workers (*4)	R 2,631.75	per month	proposal					
Consent form		per 500	Retail quote	0,002	Assumption	R 0.22		
	R 108.00	forms						
Bioscrub			Treasury	0.00101	Assumption	R 0.16		
	R 155.18	per 5L	contract					
Gloves			Treasury	0.02	RSA GF	R 1.03		
			contract		proposal for			
			RT76-2016		continued			
		per pack			funding,			
	R 50.82	100			2013			
Cotton wool swabs			2015-17	0.01	RSA GF	R 0.82		
			NDOH		proposal for			
			bandages &		continued			
			dressings		funding,			
		per pack	tender		2013			
	R 81.25	100	contract					
First response test kit			Treasury	1.01	RSA GF	R 8.25		
			contract		proposal for			
			RT41-2017		continued			
					funding,			
	R 8.17	per test			2013			
Biohazard bag		per	Retail quote	1.01	Assumption	R 0.09		
	R 0.09	material						
Results card	R 0.76	per card	Retail quote	1.01	Assumption	R 0.77		
Ground travel	R 3.55	per km	SARS rate	2	Assumption	R 7.10		

Intervention:	Workplace HCT (positive result)							
Costing method:	Ingredients							
Total cost	R 63.14 per test							
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost		
As above, plus:						R 55.50		
Advance test kit	R 6.50	per test	Treasury contract RT41-2017	1.01	RSA GF proposal for continued funding, 2013	R 6.57		
First response test kit	R 8.17	per test	Treasury contract RT41-2017	0.06	Assumption	R 0.48		
Advance test kit	R 6.50	per test	Treasury contract RT41-2017	0.06	Assumption	R 0.38		
ELISA	R 55.18	per test	NHLS 2018 price list	0.004	Assumption	R 0.22		

Intervention:	Testing of a	dolescents (negative resu	lt)					
Costing method:	Ingredients								
Total cost	R 104.34 per te								
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost			
Mobile van	R 2,108,26	per month	Updated based on [6]	0.0003	Assumption	R 0.59			
Administrative assistant	R 19,451.74	per month	2018 Government Salary Scale	0.0003	Assumption	R 5.43			
Senior nurse	R 26,359.83	per month	2018 Government Salary Scale	0.001	Assumption	R 29.43			
Community health worker	R 2,631.75	per month	RSA GF proposal for continued funding, 2013	0.007	Assumption	R 17.63			
Bond paper	R 58.80	per 500 pages	Retail price	0.002	Assumption	R 0.12			
Data technician	R 16,319	per month	2018 Government Salary Scale	0.0001	Assumption	R 2.28			
Driver	R 17,995	per month	2018 Government Salary Scale	0,001	Assumption	R 20.09			
Cleaner	R 3,775	per month	2018 Government Salary Scale	0.0001	Assumption	R0.53			
Bioscrub	R 155.18	per 5L	Treasury contract	0.0002	Assumption	R 1.03			
First response test kit	R 8.17	per test	Treasury contract RT41-2017	1.01	Assumption	R 8.25			
Biohazard bag	R 0.09	per bag	Retail quote	1.01	Assumption	R 0.09			
Cotton wool swab	R 81.65	per 100	Treasury contract	0.01	Assumption	R 0.82			

Male condom	R 0.52	per .	Treasury	10	Assumption	R 5.18
		condom	contract			
Results card	R 0.76	per card	Retail quote	1	Assumption	R 0.76

Intervention:	Testing of adolescents (positive result)							
Costing method:	Ingredients							
Total cost					R 111.0	64 per test		
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total		
						cost		
As above, plus:								
Advance test kits	R 6,50	per test kit	Treasury contract RT41-2017	1.01	Assumption	R 6.57		
First response test kit	R 8,17	per test kit	Treasury contract RT41-2017	0.06	Assumption	R 0.48		
Advance test kits	R 6,50	per test kit	Treasury contract RT41-2017	0.006	Assumption	R 0.04		
Elisa	R 55,18	per test	NHLS 2018 price list	0.004	Assumption	R 0.22		

7. Other bio-medical prevention interventions

Intervention:	Pre-exposure prophylaxis (example for PrEP to young women)									
Costing method:	Ingredients									
Total cost	R 1,917.62 per patient year (first year)									
R 1,647.07 per patient year (every year thereafter)										
			First year Every year thereafter			ter				
Drugs			R 810.84		R 810	R 810.84				
Labs			R 468.63		R 226	R 226.51				
HTS			R 104.21		R 100	R 100.62				
STI			R 37.69		R 33.	R 33.70				
Social mobilisation			R 109.82		R 109	R 109.82				
Counselling/ dispen	sing		R 138.33		R 134	R 134.35				
Combination prever	ntion		R 33.33		R 32.	R 32.33				
Overheads			R 149.01		R 133	R 133.12				
Training			R 60.07			R 60.07				
M+E			R 5.69 R 5.69							
	Health system level cost	Screening & initiation	Follow- up clinical visit	Follow-up pharmacy visit	Re- initiation visit	Creatinine	Syphilis testing (follow- up)	Total cost		
Cost per visit (patient-level costs)										
		336.43	109.46	78.60	213.30	28.71	29.72			
Cost per visit (health-system level cost)										
	219.11	23.55	7.66	5.50	14.93					
Number of visits										
First year		1	3	8	1	3	0			

Every year						1	
thereafter	-	4	8	1	1	'	

Intervention:	Post-exposure prophylaxis								
Costing method:	Ingredients								
Total cost	R 1,093.20 per patient year								
Ingredient	Unit cost	Cost unit	Source	Quantity	Source	Total cost			
PEP regimen (TDF, 3TC and ATV/Rit)	R 418.04	per month	2018 NDOH contract prices	1	2008 SAHIVSOC guidelines	R 418.04			
Space	R 110.00	per square meter	Estimate	0.007	Assumed	R 0.73			
Electricity	R1.97	per square meter	Estimate	0.007	Assumed	R 0.01			
Social Worker	R 4.36	per minute of staff time	April 2018 Government Salary Scale		GF Budget 2013-2016	R 121.07			
Counsellor	R 0.99	per minute of staff time	April 2018 Government Salary Scale	277.76	GF Budget 2013- 2016RSA	R 275.46			
Driver NGO	R 0.84	per minute of staff time	Estimate	27.78	Estimate	R 23.41			
Monitoring & Evaluation officer NGO		per minute of staff time	RSA Global Fund Grant portfolio budgets 2018	13.89	Assumed	R 29.81			
Programme Manager	R 7.02	per minute of staff time	RSA Global Fund Grant portfolio budgets 2016	6.94	GF Budget 2013-2016	R 48.75			
IEC material (design, print, distribute)	R 2.47	per material	RSA GF proposal for continued funding, 2014, inflation adjusted	1	RSA GF Proposal for continued funding, 2013	R 2.47			
Patient forms - printing	R 0.10	per form	RSA GF proposal, inflation adjusted	1	RSA GF Proposal for continued funding, 2013	R 0.10			
HIV rapid test (average cost of all test kits)	R 6.95	per test	Treasury contract RT41-2017	1.31	Assumed (for every 100 people tested, there will be 30 retests (HIV+ve + discordant tests) and 1 kit wasted	R 9.10			
Comfort packs	R 147.90	per pack	Provincial DOH CG	1	NACOSA GF Budget 2013-2016	R 147.90			

			Business Plans			
Needle	R 13.11	per pack 100	Contract HM08- 2015SYR	1	RSA GF Proposal for continued	R 0.01
Gloves	R 50.82	per pack 100	RSA GF proposal	0.01	funding, 2013	R 1.02
Cotton wool swabs	R 81.65	per pack 100	Treasury tender contract	0.01		R 0.82
Rapid HIV test disposal	R 74.09	per 5I container	2013 RSA GF proposal, inflated	0.001		R 0.07
Phone	R 449.00	rental cost per month	Retail price	0.0064	Assumed	R 2.87
Ground travel	R 3.61	per km	SARS rate	3.2	Assumed	R 11.55

REFERENCES

- Callegari, L., C. C. Harper, A. van der Straten, M. Kamba, T. Chipato and N. S. Padian (2008). "Consistent condom use in married Zimbabwean women after a condom intervention." Sex Transm Dis 35(6): 624-630
- Meyer-Rath, G., A. T. Brennan, M. P. Fox, T. Modisenyane, N. Tshabangu, L. Mohapi, S. Rosen and N. Martinson (2013). "Rates and cost of hospitalization before and after initiation of antiretroviral therapy in urban and rural settings in South Africa." J Acquir Immune Defic Syndr 62(3): 322-328.
- 3. Tchuenche M, Haté V, McPherson D, Palmer E, Thambinayagam A, Loykissoonlal D, et al. Estimating client out-of-pocket costs for accessing voluntary medical male circumcision in South Africa. PLoS One. 2016;11(10):1–10.
- 4. Bassett, I. V., D. Govindasamy, A. S. Erlwanger, E. P. Hyle, K. Kranzer, N. van Schaik, F. Noubary, A. D. Paltiel, R. Wood, R. P. Walensky, E. Losina, L. G. Bekker and K. A. Freedberg (2014). "Mobile HIV screening in Cape Town, South Africa: clinical impact, cost and cost-effectiveness." PLoS One 9(1): e85197.
- 5. Smith, J. A., M. Sharma, C. Levin, J. M. Baeten, H. van Rooyen, C. Celum, T. B. Hallett and R. V. Barnabas (2015). "Cost-effectiveness of community-based strategies to strengthen the continuum of HIV care in rural South Africa: a health economic modelling analysis." Lancet HIV 2(4): e159-e168.
- Tabana H, Nkonki L, Hongoro C, Doherty T, Ekström AM, Naik R, et al. (2015) A Cost-Effectiveness Analysis of a Home-Based HIV Counselling and Testing Intervention versus the Standard (Facility Based) HIV Testing Strategy in Rural South Africa. PLoS ONE 10(8): e0135048. doi:10.1371/journal.pone.0135048