# Analysis of economic impact among modalities of renal replacement therapy

Análise do impacto econômico entre as modalidades de terapia renal substitutiva

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#### **A**BSTRACT

Introduction: Chronic kidney disease (CKD) is a major health problem, determining the reduction in life expectancy and an increased risk of cardiovascular disease. Method: An observational, cohort, retrospective, based on patient's medical records data with CKD under hemodialysis, peritoneal dialysis and kidney transplantation in the city of Curitiba, in the period from January to June 2014, evacuativo the financial impact on the Unified Health System (SUS) and the supplementary health. Results: The lowest cost of a kidney transplant in the first year was R\$ 40,743.03 when cyclosporine was used and the highest was R\$ 48,388.17 with the use of tacrolimus. In the second year post-transplant, hemodialysis and peritoneal dialysis have a higher cost compared to kidney transplant. Transplantation with deceased donor, treated with tacrolimus: R\$ 67,023.39; Hemodialysis R\$ 71,717.51 and automated peritoneal dialysis automatic R\$ 69,527.03. Conclusions: After the first two years of renal replacement therapy, transplantation demonstrates lower costs to the system when compared to other modalities evaluated. Based on that, this therapy justifies improvements in government policies in this sector.

**Keywords:** kidney transplantation; renal dialysis; renal insufficiency, chronic; suppplementary health; Unified Health System.

#### **R**ESUMO

Introdução: A doença renal crônica (DRC) é um grande problema de saúde, determina redução na expectativa de vida e aumento dos riscos de doenças cardiovasculares. Método: Estudo observacional, de coorte, retrospectivo, baseado em dados de prontuários de pacientes com DRC em hemodiálise, diálise peritoneal e transplante renal na cidade de Curitiba, no período de janeiro a junho de 2014, avaliando o impacto financeiro no Sistema Único de Saúde (SUS) e na saúde suplementar. Resultados: O menor custo de um transplante renal no primeiro ano foi de R\$ 40.743,03, quando utilizada a ciclosporina, e o maior de R\$ 48.388,17, com a utilização do tacrolimo. Já no segundo ano pós-transplante, a hemodiálise e a diálise peritoneal têm valor superior ao transplante renal. Transplante com doador falecido, com tacrolimo: R\$ 67.023,39; hemodiálise R\$ 71.717,51 e diálise peritoneal automática R\$ 69.527,03. Conclusões: Após os dois primeiros anos da terapia renal substitutiva, o transplante demonstra menores custos ao sistema, quando comparado às outras modalidades avaliadas. Baseado nisso, esta terapia justifica melhorias nas políticas governamentais nesse setor.

Palavras-chave: diálise renal; insuficiência renal crônica; saúde suplementar; Sistema Único de Saúde; transplante de rim.

### Introduction

Chronic kidney disease (CKD) is a serious public health problem worldwide, and it is considered an "epidemic" of alarming growth. Recent data indicate that 10% of the adult population have some degree of renal dysfunction, and about 70% are unaware of having such disorder.<sup>1,2</sup>

The main risk factors for CKD are diabetes mellitus, hypertension, aging and family history. Obesity, dyslipidemia and smoking accelerate the disease progression, culminating with the need for Renal Replacement Therapy (RRT).<sup>3</sup> More than 20% of Brazilian adults have systemic arterial hypertension, 8%

have diabetes mellitus, 18% are smokers and 50% are overweight. The most alarming CKD outcomes are early mortality from cardiovascular disease and progression to chronic end-stage renal disease, and a need for RRT.<sup>4</sup>

RRT consists of hemodialysis, peritoneal dialysis and transplantation. In Brazil, current data indicate that more than 100,000 people are dependent on this therapy, and 30% are over 65 years of age. Still, according to data from the Brazilian Society of Nephrology (SBN), around 90% of the patients are on hemodialysis, with 85% of this treatment being financed by the Public Healthcare System (SUS), with an estimated annual expenditure of R\$ 2.2 billion.<sup>5</sup>

Despite treatment, the mortality of these patients is around 15% per year, being higher upon treatment onset due to the late diagnosis. Thus, early diagnosis is essential.<sup>6</sup>

The SBN has been working with the Ministry of Health to add CKD to healthcare programs, with the goals of stimulating and supporting the adoption of effective measures regarding surveillance, prevention, treatment and control for this disease.<sup>5</sup>

Thus, the present study aims to evaluate the costs of renal transplantation in the Public Healthcare System (SUS) and in the supplementary healthcare system (SHC) through the analysis of hospital bills, comparing costs associated with RRT modalities, in order to justify renal transplantation as a cost-effective treatment modality, and to stress its importance as a RRT, as well as to characterize the precariousness of an effective system of reimbursement of renal transplant procedures in the SHC, creating a basis for proposing negotiations with the HMOs.

#### **M**ETHODS

This study adopts the cost-procedure analysis to evaluate renal replacement therapy interventions for CKD (hemodialysis, peritoneal dialysis and renal transplantation), as well as its impact on the SUS and SHC. It is an observational, cohort, retrospective study based on data from patients' records of referral services in renal transplantation, hemodialysis (in the hospital setting) and peritoneal dialysis, in the city of Curitiba, PR, from January to June 2014. The values were extrapolated to annuals, since there is no price raise policy based on a given period or inflation.

Healthcare cost assessments were not based on the values of the procedures, because they did not capture all the variables relevant to CKD. We also considered

the hospitalizations related to the underlying disease and the RRT mode in question, as well as the use of medications needed for treatment maintenance.

The analysis of this study adopts the perspectives of the SUS and the SHC. Thus, only direct costs are included in this model, inferring hospitalization data and medication. The comparative analysis of RRT modalities is descriptive statistical data.

The costs associated with renal transplantation were divided into two categories. The first concerns the cost of the transplant itself, which consists of the surgical procedure for both the donor and the recipient. For this, we used the SUS procedures cost spreadsheet (Table 1). Payment is made by procedure, varying whether the donor is alive or deceased. The number of medical visits, examinations, hospitalizations and prescribed medications - especially immunosuppressants, were factored in. In this study, no costs were recorded for the donation of deceased donor organ and/or for the nephroureterectomy of the living donor. However, we added 30% increase applied to isolated kidney transplantation services, using data from results from every two years.

TABLE 1	Cost of renal transplantation from
	LIVING AND DECEASED DONORS IN THE SUS

Kidney transplant (organ from deceased donor)

Values
Hospital Care: R\$ 19,333.11
Medical Care: R\$ 8,289.56
Total Hospital Cost: R\$ 27,622.67

Values
Hospital Care:
R\$ 14.865.05

Medical Care: R\$ 6,373.77

Total Hospital Cost: R\$ 21,238.82

Ordinance No. 436, dated November 14, 2000, provides for the medical procedures paid by the SUS regarding the monitoring of patients over time. In the first year of transplantation, the Public Health Care system allows monthly payments for post-transplant patient follow-up visits, through the Authorization for High Cost/Complexity Procedures (APAC). In the following years, this billing happens every two months.

The costs associated with hemodialysis were calculated assuming that each patient is submitted to three hemodialysis sessions per week and one medical consultation per month. The cost of introducing an intravenous catheter was also included (about 70% of patients are unaware of the disease and have the option of placing a temporary catheter, called double or triple lumen catheter, to start hemodialysis) and an arteriovenous fistula - the first route in patients who already have knowledge of the disease and its evolution, since it is the best option of access for hemodialysis. The arteriovenous fistula requires, after its preparation, approximately 40 days of maturation for use.

The costs related to peritoneal dialysis were differentiated as being continuous or automatic. We considered a monthly medical consultation per patient and the cost of intraperitoneal catheter implantation.

The costs of medications and hospitalizations were also estimated. Obtaining prices for hemodialysis and peritoneal dialysis followed the same routine as that used for transplants.

#### RESULTS

A hemodialysis unit is paid for the sessions performed on end-stage CKD patients. These patients usually attend clinics three times a week, at pre-scheduled periods, being universal on Mondays, Wednesdays and Fridays or on Tuesdays, Thursdays and Saturdays.

This is the pattern followed by virtually the entire industry. Eventually, patients miss sessions or perform fewer weekly sessions depending on their residual kidney function. They can also be submitted to more sessions for shorter periods, to adjust the treatment, regardless of whether they belong to the SUS or SHC. In this way, the average number of sessions of a dialysis treatment unit is 11 to 14 monthly sessions.

Peritoneal dialysis clinics are paid a monthly income per patient, regardless of the number of peritoneal changes, varying only in the modality: continuous outpatient peritoneal dialysis (CPD) and automatic peritoneal dialysis (APD).

All proposed treatments have direct and indirect costs in their maintenance phase, regarding hospitalizations, directly related to the underlying disease, current treatment and morbidities associated with the use of high cost medications inherent to each treatment. It is evident that all the values attributed to the treatment of RRT are considered as a source of revenue for clinics, and represent expenditures for the SUS or SHC.

Initially, we present data collected from a hemodialysis clinic inside a hospital. Updated metrics

indicate a mortality rate of 10.85%; hospitalization rate of 3.3 patients/month - 78.78% for infectious reasons and 12.12% related to cardiovascular disease; the rate of patients leaving the program is 18%, because of: 46% deaths, 25% transplants, 28% migration to the peritoneal modality.

The patients were divided into SUS and SHC, and the SUS were further divided into HIV seropositive and seronegative patients. In HIV-negative patients, the annual sessions totaled R\$ 25,780.32. These values, as well as the others described, are in agreement with the basic pay, without increment, referenced by the SIGTAP table (Table of Procedures Management System, Medications and OPM of SUS) (Table 2).

## TABLE 2 COST PER HEMODIALYSIS SESSION IN THE SUS

Hemodialysis (maximum of 3 sessions per week)

High Complexity: FAEC fund

Maximum 14/month

Outpatient Care:

R\$ 179.03

Hemodialysis in HIV patients (maximum of 3 sessions per week)

Outpatient Care:

R\$ 265.41

Laboratory costs totaled R\$ 539.76 per year. Annual values with medications (Table 3) are around R\$ 7,829.89 (this value makes up a standard basic package of medications common to end-stage chronic renal disease in hemodialysis, considering the use of iron saccharate, erythropoietin, calcium carbonate, Calcitriol and sevelamer). The total value could be much higher if the patient were to use paracalcitol as an alternative for bone renal disease and it would be a little lower if the use of any of the mentioned drugs was disregarded.

Additionally, venous access (considered a central puncture with placement of a double lumen catheter and, subsequently, the creation of an arteriovenous fistula) was considered. The total mean values were R \$ 817.57 per procedure. However, this value does not predict failures both in the placement of temporary (double lumen catheter) or permanent (fistula) access.

The gross annual costs of consultations with nephrologists were R\$ 120.00 (considering the amount of R\$ 10.00 per month paid by the SUS for a standard consultation with a specialist). An estimated average annual cost of hospitalizations was also assessed and added up, totaling R\$ 1,180.00 per patient.

TABLE 3	Commonly used medications in hemodialysis								
Hemodialysis		Mean dose	% patient in use	Individual cost	Year total				
Iron saccharate 100 mg		2 vials/month	month 91.20% 3.72/vial		R\$ 89.28				
Erythropoietin 1,000 UI		14,000 UI/week	96.70%	4.17/vial	R\$ 3,035.76				
Calcium carbonate 500 mg		9 pills/day	93.40%	0.08/pill	R\$ 262.80				
Calcitriol 0.25 mcg/pill		1 pill/day	pill/day 43.00% 0.29/pill		R\$ 105.85				
Sevelamer 800 mg		6 pills/day 72% 1.98/pill		1.98/pill	R\$ 4,336.20				
Paracalcito	ol 5 mcg/ml	3 vials/week	1.3%	48.62/vial	R\$ 7,584.72				

Thus, the annual total expenses with a non-HIV hemodialysis patient was R\$ 36,267.54 on average. The HIV-positive patient presented a cost difference of R\$ 12,270.68 in sessions, with the cost per session being of R\$ 265.41; the laboratory having a cost of R\$ 372.12, of which the monthly mean value is of R\$ 31.01. The other costs: medications, accesses, consultations and hospitalizations were considered similar to those from HIV-negative patients. Therefore, the total expenses with one HIV patient, in hemodialysis, in the first year were R\$ 48,538.22. In both types of patients, HIV-positive or HIV-negative, the venous access was deducted as of the second year of treatment (Tables 4 and 5).

The dialysis patients from the SHC belong to the Unimed Curitiba and Unimed Intercâmbio HMOs, had mean costs of R\$ 47,877.12 in one year. The estimated annual cost in tests was R\$ 1,158.00. The drugs used were the same as those described in Tables 3 and 4, totaling R\$ 7,829.89/year. The cost of a double lumen catheter implant and, subsequently, the making of an arteriovenous fistula, was R\$ 897.00. Nephrologist consultations and fees, per session, were R\$ 60.00 (totaling R\$ 8,640.00 in the year). The average cost of hospitalization was approximately R\$ 3,865.41. Adding up the entire process, the first year in hemodialysis of a patient with the standard healthcare plan totaled R\$ 70,267.42. From the second year, venous access was subtracted, totaling R\$ 69,370.42.

The peritoneal dialysis modality, represented by a unit with a total of 179 patients, presented the following updated indicators: mortality rate of 11.4%; rate of exit of the program of 18%, being 46% death, 25% transplantation and 28% migration to modality hemodialysis.

The annual values paid by SUS, on average, for each patient in CPD was R\$ 21,498.72. To this value, R\$ 549.75 was added, referring to the catheter implant. The annual cost of the tests was R\$ 607.76

and for medications it was R\$ 4,642.30 (a calculation consisting of a standard basic package of the usual medications prescribed for end-stage chronic renal disease in dialysis - the use of Iron, erythropoietin, calcium carbonate, calcitriol and sevelamer at standard doses was considered in most patients, similar to hemodialysis patients) (Table 6).

The annual cost in consultations with a nephrologist totaled R\$ 660.00. In relation to hospitalization, the average value was R\$ 464.86. Finally, the annual cost of a patient in the CPD modality, for the SUS, was R\$ 28,423.39. From the second year (disregarding the catheter implant already performed the first year), the value was R\$ 27,873.64.

The patient in APD had an annual cost of R\$ 28,113.72. The mean costs of tests, medications, peritoneal access, consultations and hospitalizations were similar to those from patients in CPD. Thus, the annual cost of a SUS patient in APD was R\$ 35,038.39 and R\$ 34,488.64, in the first and second years, respectively. This difference was due to the catheter implantation required in the first year.

The annual amount paid by the reference HMO plan was R\$ 45,796.92 for CPD and R\$ 56,417.16 for APD. In both modalities, the annual cost of tests totaled R\$ 1,382.00 and R\$ 4,642.30 in medicines. The peritoneal catheter implant had a cost of R\$ 793.00. Annual costs in consultations totaled R\$ 5,184.00. The average cost of hospitalization was R\$ 2,324.31. Accordingly, the annual cost, for the reference HMO healthcare plan, for the CPD modality was R\$ 60,123.33 and for APD it was R\$ 70,743.57. From the second year, the costs related to the peritoneal catheter implantation were subtracted, totaling R\$ 59,329.53 for CPD and R\$ 69,949.77 for APD (Tables 7 and 8).

Finally, in relation to renal transplantation, the data was collected on the mirror accounts of patients transplanted in the same period (which includes all hotel expenses, materials and medications - including

TABLE 4 HEMODIALYSIS TREATMENT COST IN THE 1ST YEAR **HD SUS** HD SS HIV -HIV + R\$ 25,780.32 Sessions R\$ 38,219.04 R\$ 47,877.12 Tests R\$ 539.76 R\$ 372.12 R\$ 1,158.00 Medication R\$ 7,829.89 R\$ 7,829.89 R\$ 7,829.89 Vascular access R\$ 817.57 R\$ 817.17 R\$ 897.00 Medical visits R\$ 120.00 R\$ 120.00 R\$ 8,640.00 Hospitalization R\$ 1,180.00 R\$ 1,180.00 R\$ 3,865.41 R\$ 36,267.54 R\$ 48,538.22 Total R\$ 70,267.42

HD SUS: hemodialysis patients treated in the Public Healthcare System (SUS); HD SHC: hemodialysis patients treated in the supplementary healthcare system.

TABLE 5	Hemodialysis treatment cost in the	emodialysis treatment cost in the 2nd year							
		HD SHC							
	HIV -	HIV +	UD SUC						
Sessions	R\$ 25,780.32	R\$ 38,219.04	R\$ 47,877.12						
Tests	R\$ 539.76	R\$ 372.12	R\$ 1,158.00						
Medication	s R\$ 7,829.89	R\$ 7,829.89	R\$ 7,829.89						
Vascular ad	cess								
Medical vis	its R\$ 120.00	R\$ 120.00	R\$ 8,640.00						
Hospitaliza	ção R\$ 1,180.00	R\$ 1,180.00	R\$ 3,865.41						
Total	R\$ 35,449.97	R\$ 47,721.05	R\$ 69,370.42						

HD SUS: pacientes em hemodiálise tratados no Sistema Único de Saúde; HD SS: pacientes em hemodiálise tratados na Saúde Suplementar.

Table 6 Commonly used N	BLE 6 COMMONLY USED MEDICATION IN PERITONEAL DIALYSIS								
Peritoneal dialysis	Mean dose	% pat. In use	Individual cost	Year total					
Iron saccharate 100 mg	1 vial/month	84%	3.72/vial	R\$ 44.64					
Erythropoietin 1,000 UI	4,000 UI/week	81%	4.17/vial	R\$ 867.36					
Calcitriol 0.25 mcg/pill	1 pill/day	42%	0.08/pill	R\$ 204.40					
Calcium carbonate 500 mg	6 pills/day	95.40%	0.29/pill	R\$ 635.10					
Sevelamer 800 mg	mg 4 pills/day 80.20% 1.98/pill		1.98/pill	R\$ 2,890.80					
Paracalcitol 5 mcg/day	3 vials/week	2.20%	48.62/pill	R\$ 7,584.72					

high-cost induction drugs and procedures performed), distributed as follows: total number of transplants belonging to the SUS was 21 - 12 from deceased donors and 9 from live donors. In addition, five transplants from living donors belonging to the SHC were also reported.

As for the SUS transplants from live donors, the mean hospitalization time was 10 days. The total amount of revenues was R\$ 251,914.64, with an average of R\$ 27,914.96 per transplant. The mean number of hospitalizations from procedures using cadaveric donors was also 10 days, with a total cost of R\$ 357,035.43 (average of R\$ 29,752.95 per transplant). In the SHC, the total cost was R\$ 77,178.17 (average of R\$ 15,435.63 per transplant),

even with hospitalization time higher than that in the SUS (mean of 13 days) (Table 9).

Concerning SUS patients, the average annual cost with tests was R\$ 523.00. Regarding the medication, the main cost is related to immunosuppressants, and most renal transplantation centers use triple schemes, with prednisone and sodium mycophenolate (usually of choice) and the third drug varying between tacrolimus or cyclosporine. Alternative schemes may contain sirolimus, everolimus, or azathioprine (Table 10).

Considering three-drug regimens, the annual cost was R\$ 6,179.45 (when cyclosporine is added); R\$ 11,986.60 (when tacrolimus is added); R\$ 10,025.82 (when sirolimus or everolimus is added), and R\$ 8,146, 80 (when using tacrolimus and azathioprine).

TABLE 7 PERITONEAL DIALYSIS TREATMENT COST IN THE 1ST YEAR DP SUS DP SHC CPD APD CPD APD Treatment R\$ 21,498.72 R\$ 28,113.72 R\$ 45,796.92 R\$ 56,417.16 Tests R\$ 607.76 R\$ 607.76 R\$ 1,382.00 R\$ 1,382.00 Medication R\$ 4,642.30 R\$ 4,642.30 R\$ 4,642.30 R\$ 4,642.30 Peritoneal access R\$ 549.75 R\$ 549.75 R\$ 793.80 R\$ 793.80 Medical visits R\$ 660.00 R\$ 660.00 R\$ 5,184.00 R\$ 5,184.00 Hospitalization R\$ 464.86 R\$ 464.86 R\$ 2,324.31 R\$ 2,324.31 R\$ 35,038,39 R\$ 70,743.57 Total R\$ 28,423.39 R\$ 60,123.33

PD SUS: patients in peritoneal dialysis treated in the public healthcare system; PD SHC: patients in peritoneal dialysis treated in the Supplementary Healthcare System.

Table 8   Peritone	PERITONEAL DIALYSIS TREATMENT COSTS AS OF THE 2ND YEAR								
	PD S	SUS	PD S	SHC					
	CPD	APD	CPD	APD					
Treatment	R\$ 21,498.72	R\$ 28,113.72	R\$ 45,796.92	R\$ 56,417.16					
Tests	R\$ 607.76	R\$ 607.76	R\$ 1,382.00	R\$ 1,382.00					
Medication	R\$ 4,642.30	R\$ 4,642.30	R\$ 4,642.30	R\$ 4,642.30					
Peritoneal access									
Medical visits	R\$ 660.00	R\$ 660.00	R\$ 5,184.00	R\$ 5,184.00					
Hospitalization	R\$ 464.86	R\$ 464.86	R\$ 2,324.31	R\$ 2,324.31					
Total	R\$ 27,873.64	R\$ 34,488.64	R\$ 59,329.53	R\$ 69,949.77					

PD SUS: patients in peritoneal dialysis treated in the public healthcare system; PD SHC: patients in peritoneal dialysis treated in the Supplementary Healthcare System.

Table 9	Cost per patient in transplant									
		DD SUS	LD SUS	LD SHC						
Total number		12	9							
Mean number of days hospitalized		10	10	13						
Total		R\$ 357,035.43	R\$ 251,234.64	R\$ 77,178.17						
Mean		R\$ 29,752.95	R\$ 27,914.96	R\$ 5,435.43						

DD: deceased donor; LD: live donor; SUS: Public Healthcare System; SHC: Supplementary Healthcare.

In the first year, consultations with nephrologists are paid monthly, with most centers passing on a value of R\$ 90.00 (this is part of the post-transplant follow-up APAC), totaling R\$ 1,080.00 in the year. When hospitalized for post-transplant complications, the total amount paid per day of hospitalization was R\$ 134.00 (R\$ 34.00 being paid to the nephrology team). The main causes of hospitalizations are listed on Table 11. The mean cost of hospitalization was R\$ 5,045.62.

After accounting for all the annual costs, the SUS transplants from a live donor totaled R\$ 40,743.03 (when cyclosporine was used as immunosuppressant); R \$ 46,550.18 (when tacrolimus was used); R\$ 44,589.40 (when mTOR-sirolimus or everolimus inhibitor was used) and R\$ 42,710.38 (when using tacrolimus and azathioprine).

The SUS transplants from deceased donors totaled, in the first year: R\$ 42,581.02 (with cyclosporine use); R\$ 48,427.39 (with tacrolimus); R\$ 46,427.39 (with mTOR) and R\$ 44,548.37 (with tacrolimus and azathioprine).

The major difference in transplant costs from the second year onwards was the withdrawal of the cost of the surgical procedure and the reduction in office visits. Thus, in the second year, the average cost of living donor and cadaver transplantation was R\$ 12,828.07 (when cyclosporine regimens were used); R\$ 18,635.22 (with tacrolimus); R\$ 16,674.44 (with mTOR) and R\$ 14,795.42 (with azathioprine and tacrolimus), respectively.

In relation to live donor transplants performed by the reference HMO healthcare plan, office visits

Table 10	$m{M}$ EDICATIONS OF FREQUENT USE IN TF	RANSPLANT		
	Mean dose	% pat. In use	Individual cost	Yearly Total
Prednisone	5 mg/day	99.40%	5 mg = 0.48	R\$ 175.20
Tacrolimus	6 mg/day	69.90%	5 mg = 17.95/cap. e 1 mg = 3.61/cap.	R\$ 7,869.40
Cyclosporine	100 mg/day	12.30%	100 mg = 5.65/ capsule	R\$ 2,062.25
Sodium mycophenolate	1,080 mg/day	83.20%	360 mg = 3.60/ capsule	R\$ 3,942.00
Sirolimus	1.2 mg/day	29.40%	1  mg = 13.49	R\$ 5,908.62
Everolimus	1.5 mg/day	12.10%	0.75  mg = 11.05	R\$ 8,066.50
Azathioprine	100 mg/day	5.20%	50  mg = 0.14	R\$ 102.20

TABLE 11 CAUSES OF HOSPITA	Causes of hospitalizations								
Causes of hospitalizations	Hemodialysis	Peritoneal dialysis	Kidney transplant						
Infectious	78.78%	54.40%	62.40%						
Cardiovascular	12.12%	19%	24.50%						
Other	9.10%	26.60%	13.10%						

cost R\$ 1,296.00 in the first year and R\$ 390.00 in the second year (considering R\$ 65.00 per visit). Medication costs were the same as that in the SUS, since these patients get their immunosuppressive medications from public pharmacies.

The costs of tests averaged R\$ 4,523.00/year. The hospitalization average cost was R\$ 2,295.00. Considering the immunosuppressive regimens, the total transplantation cost in the reference healthcare plan in the first year was R\$ 19,728.88 with cyclosporine; R\$ 25,536.03 with tacrolimus; R\$ 23,575.25 with mTOR and R\$ 21,696.23 with azathioprine and tacrolimus (Table 12). In the second year, the cost was R\$ 13,387.45 with cyclosporine; R\$ 19,194.60 with tacrolimus; R\$ 17,233.82 with mTOR and R\$ 15,354.80 with azathioprine and tacrolimus (Tables 12 and 13).

#### DISCUSSION

According to the Ministry of Health, in 2008, Brazil spent 3.6% of its GDP (Gross Domestic Product) on public healthcare - this amount is equivalent to almost R\$ 109 billion. Factoring in the private healthcare industry (healthcare plans and private expenditures); total healthcare expenditures in Brazil reached 8.4% of the GDP, much lower than the amounts invested by other South American countries.<sup>7</sup>

According to the Brazilian Society of Nephrology, in 2013, 100,397 patients were on dialysis in Brazil. Most dialysis centers have a contract with the SUS

(98.2%). It is estimated that 47% of patients on dialysis are waiting in the kidney transplant queue and 25-50% of dialysis patients are diabetic. Data from the Brazilian Association of Transplants and Organs (ABTO) estimate that 92% of kidney transplants are reimbursed by SUS. In addition, most of the medicines needed to maintain the different modalities of RRT are paid for by the public healthcare system.<sup>8</sup>

The complexity of renal replacement therapy makes the costs of hemodialysis higher than the costs of peritoneal dialysis, whether manual or automated (USRDS Annual Data Report, 2013).9

Table 14 compares the various RRT modalities with the total costs at 15, 18 and 24 months. The transplant was represented in its most expensive modality: using tacrolimus.

In patients treated at the SUS, transplantation appears as the highest-cost alternative during the first and part of the second year. However, at the end of the second year, transplantation is consolidated as a less costly alternative for the treatment of CKD. In patients on RRT paid for by the standard healthcare plan, live donor transplantation presents a lower cost as of the first year. Still, at the end of 24 months, it does not even cover the cost of dialysis for the entire first year.

In the Public Healthcare System (SUS), the cost of hemodialysis patients is higher in relation to peritoneal dialysis, whether in patients with HIV or not. One of the highest costs is the need to use a disposable extracorporeal system. Other justifications for the TABLE 12 Transplant cost in reference services in the 1st year Tx SHC Tx SUS LD DD LD Treatment R\$ 27,914.96 R\$ 29,752.95 R\$ 5,435.43 Tests R\$ 523.00 R\$ 523.00 R\$ 4,523.00 Immune with Csa R\$ 6,179.45 R\$ 6,179.45 R\$ 6,179.45 Immune with Fk R\$ 11.986.60 R\$ 11.986.60 R\$ 11.986.60 Immune with Mtor R\$ 10,025.82 R\$ 10,025.82 R\$ 10,025.82 Immunes with Fk + Aza R\$ 8.146.80 R\$ 8.146.80 R\$ 8.146.80 Medical visits R\$ 1,080.00 R\$ 1,080.00 R\$ 1,296.00 Hospitalizations R\$ 5.045.62 R\$ 5.045.62 R\$ 2,295.00 Total tx w/csa R\$ 42,581.02 R\$ 40,743.03 R\$ 19,728.88 Total tx w/fk R\$ 46.550.18 R\$ 48.388.17 R\$ 25,536.03 Total tx w/Mtor R\$ 44,589.40 R\$ 46,427.39 R\$ 23,575.25 Total tx w/FK + Aza R\$ 42,710.38 R\$ 44,548.37 R\$ 21,696.23

Tx: transplant; SUS: Public Healthcare System; SHC: Supplementary Healthcare; Csa: Cyclosporine; Fk: tacrolimus; mTOR: sirolimus or everolimus; Aza: azathioprine.

TABLE 13 TRANSPLANT COS	T IN REFERENCE CLINICS IN THE	2nd year			
	Tx S	SUS	Tx SHC		
	LD DD				
Treatment					
Tests	R\$ 523.00	R\$ 523.00	R\$ 523.00		
Immune with Csa	R\$ 6,179.45	R\$ 6,179.45	R\$ 6,179.45		
Immune with Fk	R\$ 11,986.60	R\$ 11,986.60	R\$ 11,986.60		
Immune with Mtor	R\$ 10,025.82	R\$ 10,025.82	R\$ 10,025.82		
Immune with Fk + Aza	R\$ 8,146.80	R\$ 8,146.80	R\$ 8,146.80		
Medical visits	R\$ 540.00	R\$ 540.00	R\$ 540.00		
Hospitalization	R\$ 5,045.62	R\$ 5,045.62	R\$ 5,045.62		
Total tx w/csa	R\$ 12,288.07	R\$ 12,288.07	R\$ 12,288.07		
Total tx w/fk	R\$ 18,095.22	R\$ 18,095.22	R\$ 18,095.22		
Total tx w/Mtor	R\$ 16,134.44	R\$ 16,134.44	R\$ 16,134.44		
Total tx w/FK + Aza	R\$ 14,255.42	R\$ 14,255.42	R\$ 14,255.42		

Tx: transplant; SUS: Public Healthcare System; SHC: Supplementary Healthcare; Csa: Cyclosporine; Fk: tacrolimus; mTOR: sirolimus or everolimus; Aza: azathioprine.

costs of hemodialysis being higher than those for peritoneal dialysis are the costs of supplies, facilities, specialized labor, as well as the impact of expenses with vascular access, medication and hospitalization.

In SHC, the costs of hemodialysis patients are also higher in relation to those in manual peritoneal dialysis, but equivalent to automated peritoneal dialysis.

To maintain the different replacement therapies, numerous medicines are necessary, which increases the financial impact. Many of them are considered strategic or exceptional, being provided by pharmacies of the public healthcare network.

Reliable data on the amounts invested in dialysis drugs in Brazil are not available. However, in 2011, the US healthcare system 'Medicare' reported spending \$1.69 billion on drugs supplementing dialysis, while for kidney transplantation \$315 million were invested in the same period.<sup>5</sup>

One of the largest CKD databases, the United States Renal Data System (USRDS), shows that renal transplantation has a lower rate of hospitalization, with infectious causes accounting for the majority.<sup>5</sup> Similarly, the data found in this study is compatible with that in the literature.

TABLE 14	Comparing total expenditures in rrt modalities in up to 24 months								
	HD		PD	SUS	PD	SHC	Tx SU	S w/Fk	Tx SHC w/Fk
	SUS	SHC	CPD	APD	CPD	APD	LD	DD	LD
1st year	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
	36,267.54	70,267.42	28,423.39	35,038.39	60,123.33	70,743.57	46,550.18	48,388.17	25,536.03
2nd year	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
	35,449.97	69,370.42	27,873.64	34,488.64	59,329.53	69,949.77	18,635.22	18,635.22	20,100.60
Monthly	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
2nd year	2,954.16	5,780.87	2,322.80	2,874.05	4,944.13	5,829.15	1,552.94	1,552.94	1,675.05
15	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
months	45,130.03	87,610.03	35,391.80	43,660.55	74,955.71	8,231.01	51,208.99	53,046.98	30,561.18
18	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
months	53,992.53	104,952.63	42,360.21	52,282.71	89,788.10	105,718.46	55,867.79	57,705.78	35,586.33
24	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$	R\$
months	71,717.51	139,637.84	56,297.03	69,527.03	119,452.86	140,693.34	65,185.40	67,023.39	45,636.63

HD: hemodialysis; PD: peritoneal dialysis; CPD: Continuous peritoneal dialysis; APD: automatic peritoneal dialysis; LD: live donor; DD: deceased donor; SUS: Public Healthcare System; SHC: Supplementary Healthcare; Tx: transplant; Fk: tacrolimus.

In this study, the cost of hospitalizations on hemodialysis was higher than that for peritoneal dialysis in the SUS, although it was lower in the SHC, although this difference may be influenced by the lower number of SHC patients compared to the SUS. However, although the referred hemodialysis service is a hospital unit, hospitalization rates in this modality and renal transplantation were similar (even with a shorter hospital stay on hemodialysis).

Such a bias may be due to the lack of exclusive beds for the hemodialysis sector, having the clinic take on responsibility for day-care for patients, which reduces the average cost of hospitalization, with a financial impact lower than expected.

In general, the index of patient hospitalizations, either in hemodialysis or peritoneal dialysis, is equivalent: 11.7 days/patient/year in 2011;<sup>5</sup> while in kidney transplantation it would be in the order of 5.7 days/patient/year. If compared to the general population, chronic renal patients have a 16% higher re-hospitalization rate.

Considering epidemiological data, the number of new patients diagnosed with CKD and the number of patients undergoing treatment (starting or even under maintenance) increased almost 11-fold between 1980 and 2011.<sup>5</sup>

Among the options of RRT, transplantation has the lowest mortality rate and better quality of life, surfacing as the modality of choice for treatment. In addition, the cost of immunosuppressive medication is lower in relation to the medications used in hemodialysis and peritoneal dialysis. In the first year, the cost of hemodialysis and automated peritoneal dialysis was lower than the average cost of renal transplantation, regardless of the immunosuppressive regimen used, except for hemodialysis in HIV patients. In the second year of treatment, hemodialysis and peritoneal dialysis have a higher value than renal transplantation, confirming data presented by other studies. <sup>10-12</sup>

In relation to SHC, living donor transplant costs are already lower as of the first year, and at the end of 24 months, the cost of dialysis during the first year has not yet being reached. Another situation to be considered is the lack of stimulation for renal transplantation, since the medical compensation for patients on hemodialysis or peritoneal dialysis is higher than the fees charged for transplant procedures.

Data on the few transplants made in the reference HMO, compared to patients on RRT in the hemodialysis and peritoneal dialysis modalities, showed that regardless of any immunosuppressive agent used, the cost of kidney transplantation is lower than hemodialysis or peritoneal dialysis as of the first year of treatment, when it is equivalent to 32-38% of the cost of the previous ones. From the second year onwards, the cost of a kidney transplant can be 18% of the amount spent on hemodialysis or automated peritoneal dialysis, or 21% of that of manual peritoneal dialysis.

In the patients treated by the SUS, transplantation appeared as a costlier alternative during the first year and part of the second year, and at the end of the second year it was established as the best alternative for RRT.

Some limitations of the study could not be avoided. First, the data does not represent values of extreme accuracy; however, these were obtained through the instruments that govern the financial and billing system of hospital costs related to the SUS and the SHC. Second, the population represents the national average of a continental-size country, with considerable interregional differences. Third: the automatic extrapolation of treatment costs in absolute terms, although there were no adjustments in the SUS or SHC during the period analyzed. Fourth, the costs related to the different RRT modalities and their aggregate items vary between different healthcare services and hospitals, as well as the budget incentive values between the different therapeutic modalities and the participation of the third sector in face of the numerous social difficulties, and the availability of vacancies for hospitalization.

#### CONCLUSION

Regardless of structural factors and socioeconomic conditions, kidney transplantation costs are lower as of the second year, strengthening this modality as the main RRT option, since in addition to the financial factor, transplanted patients have higher survival, better quality of life, lower hospitalization index, lower mortality rate and reduction in direct and indirect costs with their comorbidities. In addition, many return to full working condition, contributing with taxes and growth for the country, with less burdening of the system.

Even with some costs underestimated due to the absence of reliable indicators regarding the treatment of the RRT population in Brazil, the information generated is useful for cost impact analyses, in the SUS and SHC, serving as a reference for future studies, epidemiological estimates, budget planning and negotiations with the HMOs.

Prevention policies should be planned, postponing or avoiding the need for RRT. However, when this therapy is necessary, the option should be directed to the most cost-effective modality, that is, renal transplantation.

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