A transition from using multi-step procedures to a fully integrated system for performing extracorporeal photopheresis: A comparison of costs and efficiencies

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Abstract

Introduction: The Pitié Salpêtrière Hospital Hemobiotheraphy Department, Paris, France, has been providing extracorporeal photopheresis (ECP) since November 2011, and started using the Therakos® CELLEX® fully integrated system in 2012. This report summarizes our single-center experience of transitioning from the use of multi-step ECP procedures to the fully integrated ECP system, considering the capacity and cost implications.

Materials and Methods: The total number of ECP procedures performed 2011–2015 was derived from department records. The time taken to complete a single ECP treatment using a multi-step technique and the fully integrated system at our department was assessed. Resource costs (2014€) were obtained for materials and calculated for personnel time required. Time-driven activity-based costing methods were applied to provide a cost comparison.

Results: The number of ECP treatments per year increased from 225 (2012) to 727 (2015). The single multi-step procedure took 270 min compared to 120 min for the fully integrated system. The total calculated per-session cost of performing ECP using the multi-step procedure was greater than with the CELLEX® system (€1,429.37 and €1,264.70 per treatment, respectively).

Conclusions: For hospitals considering a transition from multi-step procedures to fully integrated methods for ECP where cost may be a barrier, time-driven activity-based costing should be utilized to gain a more comprehensive understanding the full benefit that such a transition offers. The example from our department confirmed that there were not just cost and time savings, but that the time efficiencies gained with CELLEX® allow for more patient treatments per year.

KEYWORDS
cost comparison, photopheresis, time-driven activity-based costing

1 INTRODUCTION

Extracorporeal photopheresis (ECP) is a therapeutic procedure recommended for the treatment of patients with conditions associated with cutaneous T-cell lymphoma (CTCL) as well as acute and chronic graft-versus-host disease (GvHD). ECP has also been reported to benefit patients with solid organ transplant rejection and autoimmune diseases such as scleroderma and Crohn’s disease.1–3

ECP combines leukapheresis and photodynamic therapy, and can be performed using a multi-step procedure or using a fully integrated system.4 The mechanism of photopheresis...
is not fully understood, however, it is theorized that T-lymphocyte cells damaged by ultraviolet A (UVA) during the procedure activate the patient’s immune system.1–5

Multi-step ECP procedures require separate components for leukocyte collection (including cell separation using an instrument such as the COBE® Spectra [Terumo BCT]), addition of the photosensitizing agent (methoxsalen/8-methoxypsoralen, such as UVADEX® [Therakos®] or 8-MOP [MacoPharma®] solutions), UVA radiation (performed using a system such as Theraflex [MacoPharma®]), and re-infusion of treated cells.1,6 UVADEX® is the only formulation of methoxsalen with widespread regulatory approval for the treatment of CTCL as part of ECP.7 8-MOP solution, recommended for use with the Theraflex system, is only approved as a related therapeutic product in France.8 The European Guidelines for minimal cell manipulation state that multi-step procedures should be performed in a class A laminar airflow cabinet located in a class D laboratory.9

Fully integrated ECP systems combine all the required processes, with the patient remaining connected to the system throughout the treatment.1 This ensures sterility, alleviates the need to cross match re-infused materials, and reduces the risk of improper infusion, contamination, or infection.1 The duration of the ECP procedure with fully integrated systems is shorter than procedures requiring multiple steps.1,10–14 In Europe, the only approved fully integrated instruments specifically designed for ECP are the Therakos® UVAR-XTS and CELLEX® systems,1 which utilize UVADEX® photosensitizing solution.7 Fully integrated systems are associated with higher set up costs compared to the multi-step procedures, and some reports suggest that the fully integrated systems may be less suitable for patients with restricted venous access.15,16 Current estimates suggest that around 75% of apheresis departments in France perform ECP using integrated systems.17

Ensuring effective and efficient use of healthcare resources is a system-wide priority. When seeking to optimize hospital services, particularly where there is an option to incur substantial acquisition costs related to new technologies, time-driven, activity-based costing (TDABC) methods can be useful as these go beyond methods which utilize traditional hospital cost accounting systems. TDABC provides a more comprehensive understanding of resource usage and its associated costs, while measuring processes and encouraging quality improvement.18,19 A number of studies have investigated differences in procedure times and costs associated with the Therakos® CELLEX® fully integrated ECP system and other main alternatives, and have suggested advantages for the integrated system.12,20,21

The Hemotherapy Department at Pitié Salpêtrière Hospital in Paris, France, has focused on the mobilization and collection of hematopoietic stem cells (HSC) for autologous and autologous HSC transplantation (HSCT) since 1989. The department has been performing ECP procedures for patients presenting with GvHD after HSCT and patients with CTCL since November 2011, and for lung transplant patients since June 2013. At the time of the study, the department’s resources included three hospital beds, two Spectra Optia® cell separators, two CELLEX® fully integrated ECP systems, two full-time nurses, and one full-time physician. Given its size and resource availability, the department sought to assess and address the increasing demand for ECP procedures while also maintaining the same number of necessary healthcare providers within the unit.

The aim of this report is to summarize a single-center experience of transitioning from the use of multi-step ECP procedures to a fully integrated ECP system, considering the capacity and cost implications through the use of TDABC.

2 | MATERIALS AND METHODS

2.1 | Data acquisition

Total ECP procedures performed by the department were derived from a review of the department records from 2011 to 2015. A detailed review of procedures carried out in 2015 was also undertaken to investigate the reliability of the integrated ECP procedures.

To investigate the relative resource requirements of each ECP method, one patient undergoing ECP using the fully integrated system and one patient undergoing ECP using multi-step procedures at Pitié Salpêtrière Hospital were observed during February 2014. Specifically, the fully integrated Therakos CELLEX® system was used, and the multi-step procedures included cell separation using Spectra Optia®, and UVA irradiation using MacoPharma® Theraflex MB-Plasma. Informed, written consent was obtained from both patients prior to the procedures.

Details of each activity’s execution were recorded, including time spent per activity performed by hospital personnel. The time for pre- and post-treatment observations (eg, blood pressure), cannulation time, treatment time, addition of methoxsalen, UVA irradiation, biological sampling, and the reinfusion of cells were measured. Total duration of patient retention time using both systems was compared. Results were reviewed by experienced system users to confirm that the observations were typical of each procedure method.

2.2 | Time-driven, activity-based costing comparisons

TDABC methods were applied to provide a meaningful cost comparison between administration of ECP using multi-step procedures and fully integrated systems.17,18

Costs reviewed included unit costs of hardware and calculated costs of personnel based on timeline comparison results.
The costs of individual activities used in both techniques were obtained from the purchasing department of the Pitié Salpêtrière Hospital. The costs of every activity and consumable associated with performing ECP using both the multi-step procedure and fully integrated system were considered. It was assumed that the associated typical costs are independent of patients’ disease type and other patient characteristics.

Bed retention times were calculated by taking the daily overhead cost incurred by the hospital while providing a bed. This cost includes all associated overheads but does not include personnel costs. Daily costs were prorated for each system based on the number of hours required to complete one treatment.

Personnel costs were calculated by taking the hourly rate of salary for each involved healthcare provider, plus all associated additional employment costs incurred by the hospital (pension, tax, insurance, etc). The number of hours and minutes that a provider (nurse and/or physician) needed to be present with a patient undergoing treatment was then calculated. The hourly rate was then multiplied by the time that the provider was in attendance with the patient to create an assigned personnel cost. In the case of the multi-step procedures, the additional costs of porters (for transport of cells) and for laboratory technicians (for the offline treatment of cells) were also included in the calculation. Costs were recorded in 2014 euros, and not adjusted for inflation.

### 3 RESULTS

Initially, the hospital utilized multi-step procedures for all ECP treatments. In the first full year of practice (2012), 225 treatments were carried out. A transition from the multi-step procedures to the Therakos® CELLEX® fully integrated system began at the end of 2012. The following year, 397 ECP procedures (75.3% of all procedures) were carried out using the CELLEX® fully integrated system.

At the start of 2014, the Hemobiotherapy Department was faced with capacity constraints. A complete transition to the fully integrated system was completed in 2014, with a total of 686 ECP procedures performed via the CELLEX® system that year. In order to optimize the delivery of ECP procedures with the new system, a ProcEx Solutions Limited work-flow assessment was undertaken. The aim was to improve ways of managing the increasing demand while maintaining the same numbers of personnel necessary to operate the unit. The combination of improved efficiencies in work-flow and the CELLEX® fully integrated system enabled a patient capacity increase of 223% from 2012 (225 procedures) to 2015 (727 procedures).

In 2015, all ECP procedures were completed with the CELLEX® system, with approximately 5% experiencing transient problems (resolved through a system stop followed by resuming the procedure) and two cases of failed procedures due to mechanical errors. There were no recorded cases of failed procedures due to patient complications. The CELLEX® systems required an average of three machine stops a year for maintenance.

#### 3.1 Timeline comparison

The patient treated using the multi-step procedure was retained for more than double the time of the patient treated with the fully integrated system (270 min vs. 120 min, respectively) (Figure 1). Given the assumption that there are only 450 min (7.5 h) in a working day, these findings translate to the ability to treat only one patient per bed per day using multi-step procedures compared to three patients per bed per day using the fully integrated system.

#### 3.2 Time-driven activity-based costing comparison

Following the timeline analysis and comparison, time-driven costs were obtained for each of the steps required under both the multi-step procedure and CELLEX® fully integrated system (Table 1). The cost of performing ECP using the multi-step procedures was greater than with the CELLEX® system (€1,429.37 and €1,264.70 per treatment, respectively, Table 1). While the CELLEX® system had substantially higher costs upfront, the additional steps and personnel costs required to perform ECP using multi-step procedures had a higher total cost.

A total cost comparison for the number of treatments carried out in 2015 (based on 727 treatments) revealed cost savings of €119,715 associated with performing all ECP treatments on the CELLEX® system.

### 4 DISCUSSION

Previous studies within our department reported that the use of plerixafor to aid peripheral blood stem cell mobilization in patients undergoing autologous stem cell transplantation
TABLE 1  Time-driven activity-based cost comparison of the multi-step procedures and the fully integrated CELLEX® ECP system

| Phases in treatment cycle                        | Multi-step | CELLEX® |
|-------------------------------------------------|------------|---------|
| Collection of leukocyte concentrate (PTC)       | €173.75    | €1,009.20 |
| Biological analysis (patient)                   | €59.00     | €59.00  |
| Biological analysis (cells collection)          | €31.05     | €0.00   |
| Transportation of cells to cell treatment area   | €7.76      | €0.00   |
| Handling of cell product by cell manipulation facility | €616.55 | €0.00   |
| Biological analysis (irradiated cells)          | €76.00     | €0.00   |
| Transportation of cells to ward                 | €7.76      | €0.00   |
| Injection (triple access)                       | €12.00     | €0.00   |
| Personnel costs                                 | €108.00    | €40.50  |
| Bed retention cost/treatment (per hours used)   | €337.50    | €156.00 |
| **Total Cost**                                  | €1,429.37  | €1,264.70 |

*The total cost is per one treatment cycle, values are in 2014 EUR.  

In terms of procedural time, the CELLEX® fully integrated system benefits from a double-needle mode, providing more rapid treatment times (110 min vs. 135 min using the single-needle mode). However, some patients, particularly those with sclerotic skin and reduced joint movement around the elbows, have been reported to find the double-needle mode to be uncomfortable and impractical. Some centers consider the single-needle mode, which is still more rapid than performing ECP using a multi-step procedure, more suitable for these patients. However, our protocol of anxiolytic medication, heating covers, and a local anesthetic applied 15 min prior to the ECP procedure ensures a good fit even in patients with poor venous access, and we have not found the single-needle mode necessary in practice.

5 | CONCLUSIONS

For hospitals considering a transition from multi-step procedures to fully integrated methods for ECP where cost may be a barrier, time-driven activity-based costing should be applied to gain a more comprehensive understanding of the full costs of such a transition. The example from Pitié Salpêtrière Hospital confirmed that time efficiencies gained with CELLEX® allow for more patient treatments per year.

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