

Ensaio

Prescription Patterns of Pharmaceuticals

Ana Sofia Oliveira Gonçalves¹

1. Introduction

The majority of the population gets in touch with the health care sector in several occasions: as patients, policy makers, providers, taxpayers and as citizens.² In what concerns the European Union member-states, a Eurobarometer survey showed that healthcare is seen as the 5th vital matter, being considered more essential than both housing and education.³ According to the World Health Report of 2010, four out of ten causes that lead to inefficiencies in the health care sector are associated with pharmaceutical products: prices, quality, use and waste.

In May 2011, the Economic Adjustment Program for Portugal was settled, followed by the signing of the Memorandum of Understanding and the Loan Agreement.⁴ In line with the Memorandum of Understanding on Specific Economic Policy (MoU), Portugal needs to ensure a reduction in the public spending on pharmaceuticals of about 1% of GDP in 2013.⁵ Furthermore, Portugal has to develop its monitoring system of medical drugs. This includes a tougher assessment of each individual physician's prescriptions in what concerns volume and total value, regarding the prescription guidelines that were set. Therefore, feedback must be given to medical physicians on a regular basis, especially information concerning the most used and the costliest medical drugs. From August 2012 on, physicians working in Centro Hospitalar Central de Lisboa, the provider of this database, are given feedback on their expenditure in pharmaceuticals and the expenditure incurred by their colleagues. The unit price of each pharmaceutical product prescribed is not included. This feedback appears in each physician's intranet personal page, which requires a password. Physicians play an important role since they are the agents that decide which pharmaceutical product to prescribe, influencing the hospital's total expenditure.

The importance of understanding medical physicians' prescribing patterns relies on the fact that the pharmaceutical drugs expenditures can substantially change depending on the way doctors prescribe. According to the Law of Large Numbers, each physician should treat similar patients, on average, in terms of health status and related expenditure. Hence, it should be expected that prescription patterns of pharmaceuticals would be similar across hospitals and across physicians with the same medical specialty.

Therefore, the aim of this research work is to analyse the possibility of hospital cost reductions in pharmaceutical drugs, by better practice from physicians in the way they prescribe.^{6 7} This analysis considers pharmaceuticals administered in a hospital setting, only. Furthermore, it will focus on particular diseases: HIV/AIDS, oncology and rheumatism. The criteria used for the selection of medical conditions were its relative weight and growth tendency in hospital costs. Every year, Portugal incurs in very high expenditure levels related to HIV/AIDS treatment (an average cost of treatment of €13.625 per patient-year). Antiretroviral treatment represents the main cost driver (€8.767€ per patient-year).^{8 9} In this database, which only includes pharmaceutical products' costs, the six hospitals together spent €60.238.005,52 during the six

¹Corresponding author. Nova School of Business and Economics. Office for Strategy and Studies (GEE), Portuguese Ministry of Economy and Employment. Email: sofia172@gmail.com

² Gartner (2009)

³ Eurobarometer

⁴ European Commission (2011)

⁵ IMF (2013)

⁶ A good practice, according to the WHO (1987), is one which "requires that patient receive medication appropriate to their clinical needs, in doses that meet their own individual requirement for an adequate period of time and at the lowest cost to them and their community"

⁷ Hospital pharmaceutical drugs will be defined as all pharmaceutical drugs administered in a hospital setting, for both inpatients and outpatients

⁸ According to the WHO (2010) antiretroviral treatment "consists of the combination of at least three antiretroviral (ARV) drugs"

⁹ Oliveira et al. (2011)

quarters of available data. The direct costs of cancer treatment in Portugal, represent 3,91% of the total expenditure in health care, which means €53 per capita. In what concerns the total expenditure on pharmaceutical drugs, oncology drugs are responsible for 5,6% of this amount.¹⁰ Ageing is an important determinant in what concerns rheumatism.¹¹ Since the Portuguese population has been ageing, an increase in the costs associated with this disease is expected.¹²

The patterns of drug prescriptions by physicians in six public hospitals in Lisbon were modelled using a relative distribution model and a stochastic dominance analysis. The relative distribution analysis represents a way to compare expenditure levels between physicians. Therefore it is possible to identify outlier physicians (physicians incurring into higher or lower expenditure levels than the norm, on average, in comparison to their colleagues).¹³ The stochastic dominance analysis allows an intertemporal comparison of expenditure levels for each physician. A regression using a probit model was carried out to determine which variables are significant in explaining outlier physicians above and below the norm. My findings show that there is great divergence of prescribing patterns across physicians, the hospitals where they prescribe and their medical specialty. Outlier physicians, incurring into higher or lower expenditure levels than the norm, tended to maintain their prescribing patterns in the observation period. Even after the introduction of individual feedback their prescribing behaviour did not show major changes. Concerning the evolution of total pharmaceutical product costs, a decrease in product prices does not seem to have had a permanent impact in expenditure levels.

2- Literature Review

Pharmaceutical treatment choices can be influenced by both demand-side and supply-side factors.¹⁴ Demand-side factors include prices, personal preferences and income. The database of this work only refers to pharmaceutical products that are given for free to every patient, in public hospitals, making the treatment choice independent of the patients' decisions. Supply-side factors include the technology available to perform the different treatment choices. Carone et al. (2012) identified several ways to influence and try to improve the way medical physicians prescribe as prescription monitoring, prescription guidelines, targets for prescription costs, prescription quotas, financial incentives and educational training.

Hogerzeil (1995) classified strategies to promote a rational prescribing behaviour into three different categories: educational, managerial and regulatory. In what concerns their possible effectiveness, the author concludes that educational strategies, such as printed materials of drug lists and treatment guidelines, alone are unlikely to influence prescription patterns unless they are followed by introductory campaigns and these physicians take part on the process. Furthermore, he highlighted the importance of achieving consensus among physicians when introducing treatment protocols and drug lists. Moreover, Schroeder et al. (1984) found that given feedback individually for each physician may actually be so costly that will offset the potential gains.

Allan et al. (2007) found that physicians lack knowledge in what concerns the costs of pharmaceutical drugs, often underestimating the price of expensive drugs. Nonetheless, Hart (1997) concluded, that family physicians tend to prescribe less expensive drugs even before having prior information about their costs.

¹⁰ Araújo et al. (2009)

¹¹ Lucas (2012)

¹² Pordata

¹³ According to Barnett and Lewis (1994) an outlier is: "An observation (or subset of observations) which appears to be inconsistent with the remainder of that set of data."

¹⁴ Culver et al. (2000)

3- Data and Methodology

3.1- Data

The data source for this project comes from different hospitals located in Lisbon. They belong to Centro Hospitalar de Lisboa Central. The hospitals are: Hospital Curry e Cabral (HCC), Hospital de São José (HSJ), Hospital Santo António dos Capuchos (HSAC), Hospital de Santa Marta (HSM), Hospital Dona Estefânia (HDE) and the Maternity Hospital Alfredo da Costa (MAC). The data refers to the period from the 1st quarter of 2012 to the 2nd quarter of 2013.

In 2012, HSJ, HSAC and HDE were already using a common database. Only in 2013, HCC, HSM and MAC started to be included in this database. Therefore, for any analysis of the current database, one needs to take into account that before 2013 HCC, HSM and MAC were registering episode numbers and patient numbers independently from the other hospitals already with a common database. Thus, different patients from different hospitals may, in 2012, have the same patient number.

Furthermore, it may happen that the same patient has different hospital numbers. Likewise, it may occur that the same episode number in different hospitals in 2012 represented different and not related episodes. The latter problem was overcome by analysing not the episode number but a code aggregating both, the episode and the patient number, so repetitions are avoided. In what concerns patients, a code aggregating both the patient number and the hospital name was created. In 2013 this problem does not arise since the database already includes all hospitals in a consistent coding procedure. A given episode contains information regarding patient's health problem currently being treated and the description of the provided service according to different medical specialties as well as the drugs that were provided and their cost. There is also information concerning which one was the physician responsible for each episode. This record focuses on the characteristics of the drugs prescribed to the patients in the hospital. It includes the beginning and the end date of the treatment of the given pharmaceutical, the dose and the frequency, the quantity, the unit cost and the total cost of the prescribed pharmaceutical.¹⁵

The information gathered allows not only an intra-hospital comparison, but also the possibility to monitor pharmaceutical costs between institutions.

3.2- Methodology

3.2.1- Relative Distribution

To compare expenditure levels between physicians, a relative distribution analysis is used. Therefore it is possible to identify outlier physicians, physicians incurring into higher or lower expenditure levels than the norm, in comparison to their colleagues.¹⁶ Hence, this method detects physicians which incurred into significant deviations from a given (average) prescription pattern.

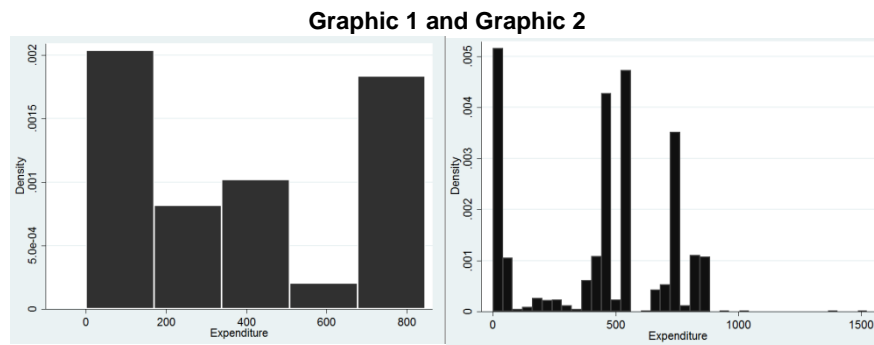
Relative distributional methods consist of non-parametric statistical processes. Nonparametric density estimates allow for the comparison between groups and to a group with a benchmark density (e.g. the normal).¹⁷ It does not require the adoption of assumptions in what concerns the mathematical form of the probabilities distribution of a variable.¹⁸ Hence, non-parametric models prevent violations of the hypothesis of models which would lead to misleading answers. The two histograms below, showing the expenditure level of two specific medical physicians, highlight how hard it would be to find a benchmark density suitable for every medical physician regarding his/her number of prescriptions over prices/value per prescription:

¹⁵ Further information can be found in the Extra Annexes, section A.1.

¹⁶ According to Barnett and Lewis (1994) an outlier is: "An observation (or subset of observations) which appears to be inconsistent with the remainder of that set of data."

¹⁷ Cameron et al. (2005)

¹⁸ Being a distribution the depiction of the relative numbers of times each potential outcome will occur in a number of trials



Moreover, it contains a graphical component, which simplifies the analysis.

This method establishes an association among distributions, allowing the possibility of making direct comparisons between groups or points in time, with respect to a continuous outcome variable. A reference group must be set, as well as a comparison group. Comparisons are done by getting comparison group's points and understand where they fall in the reference distribution (i.e. in which quantile). Thus, it converts two distributions into a single one allowing for an analysis that is scale independent (it shows the set of percentage positions that the observations of one distribution would have were they located in another distribution). If both distributions are equal, then the "relative data" will be uniformly distributed.

As stated by Handcock e Morris (1999), relative distribution methods include particularities such as not being affected by the scale choice or to monotonic transformations, basing its study in the population and not the individual and it computes proportions of individuals and calculates their ranking position.

Let Y_0 be the outcome variable in the reference group, and Y be the outcome variable in the comparison group. It is usually the measurement for a different group or it can be the identical group in another time period. Being F_0 the distribution function of the reference group we have that:

$$R = F_0(Y)$$

The Cumulative Distribution Function (CDF) of the relative data R is given by:

$$G(r) = F[F_0^{-1}(r)] = Q_0(r) \quad 0 \leq r \leq 1 \quad \text{where } Q_0(r) \text{ is the quantile function of the reference distribution}$$

Taking the first order derivative of $G(r)$ the density function is obtained:

$$g(r) = f\{[F_0^{-1}(r)]\} / f_0\{[F_0^{-1}(r)]\} = f[Q_0(r)] / f_0[Q_0(r)] \quad 0 \leq r \leq 1 \quad \text{where } f \text{ and } f_0 \text{ are density functions}$$

The relative distribution allows to directly compare one distribution with another.

It is expressed as a random variable which was attained by converting a variable from the comparison group by the CDF of that variable for a reference group. This conversion will lead to a set of observations (relative data), which characterise the rank of the initial comparison value in terms of the CDF of the reference group. The Probability Density Function (PDF) may be interpreted as a density ratio between two different distributions i.e., "the ratio of the fraction of respondents in the comparison group to the fraction in the reference group at a given level of the outcome attribute". To use the relative distribution model one only needs to include those that have prescribed at least more than four times.

To complement the visual component, the Kolmogorov-Smirnov test is used to test the hypothesis whether two samples follow the same distribution or not.¹⁹ Specifically, it will be compared the distribution of the expenditure level of each physician with the expenditure level of the other physicians excluding himself/herself. The null hypothesis assumes that the samples are drawn from the same distribution. This test requires continuous distributions, as in this case, and it is a non-parametric method. Therefore, such analytical components are appropriate to use in this analysis.

¹⁹ Further information can be found in the Extra Annexes, section B.1.

3.2.1- Stochastic Dominance

To compare the evolution of each physician's expenditure across time the stochastic dominance analysis is applied. Stochastic dominance approaches have been traditionally applied in economics in relation to assets with monetary payoffs and in relation to poverty and income distribution analysis.²⁰

The several types of pharmaceutical drugs used in each type of medical treatment entail an expenditure distribution. In this case, the First Stochastic Dominance (FSD) will be useful to compare expenditure levels, between two periods, for each doctor. Therefore, applying a stochastic dominance technique offers the opportunity to identify whether expenditure levels for a given physician have been rising or falling across time. However, one has to consider that a physician's expenditure level in pharmaceuticals is also affected by price effects. A specific case is a modification in law that may contribute to changes in pharmaceutical expenditures across time. Indeed, since the 1st of March of 2013, the reference prices for pharmaceutical products changed.²¹ The assumption is that both, the physicians' and patients' characteristics, remain constant across time.

3.2.1.1- First-degree stochastic dominance (FSD)

Suppose that there are two possible distributions F and G, with cumulative density functions (CDF) of $F(x)=P_r(X\leq x)$ and $G(x)=P_r(X\leq x)$.

The CDF of F(x) dominates G(x) by first degree stochastic dominance if and only if:

$$F(x)\leq G(x), \text{ for all } x$$

3.2.3- Ordered probit model

An ordered probit model is implemented to assess which characteristics associated with the outlier physicians are significant in explaining their expenditure levels' deviation to the norm. This model was not run for rheumatism since there are too few observations. The dependent variable, being an outlier (y) is a limited dependent variable which takes the following form: $y=1$ if the physician is an outlier with an expenditure level above the norm and $y=-1$ if the physician is an outlier with an expenditure level below the norm and $y=0$ his expenditure is according to the norm.

$$Y = \begin{cases} -1 & \text{if the physician is an outlier with an expenditure level below the norm} \\ 0 & \text{his expenditure is according to the norm} \\ 1 & \text{if the physician is an outlier with an expenditure level above the norm} \end{cases}$$

Being $y_i^* = x_i'\beta + u_i$ a latent index model where $y_i = (-1,0,1)$ For a three alternative ordered probit model:

$$y_i = -1 \text{ if } y_i^* \leq \alpha_1$$

$$y_i = 0 \text{ if } \alpha_1 < y_i^* \leq \alpha_2$$

$$y_i = 1 \text{ if } y_i^* \geq \alpha_2$$

Thresholds separate the ordering of alternatives. In this case we have:

$$\Pr(y_i = -1) = \Pr(y_i \leq \alpha_1) = \Pr(x_i \beta + \varepsilon_i \leq \alpha_1) = \Pr(\varepsilon_i \leq \alpha_1 - x_i \beta) = \Phi[\alpha_1 - x_i \beta] = 1 - \Phi[x_i \beta - \alpha_1]$$

$$\Pr(y_i = 0) = \Pr(\alpha_1 < y_i^* \leq \alpha_2) = \Pr(y_i^* \leq \alpha_2) - \Pr(y_i^* \leq \alpha_1) = \Pr(x_i \beta + \varepsilon_i \leq \alpha_2) - \Pr(x_i \beta + \varepsilon_i \leq \alpha_1) = \Pr(\varepsilon_i \leq \alpha_2 - x_i \beta) - \Pr(\varepsilon_i \leq \alpha_1 - x_i \beta) = \Phi[\alpha_2 - x_i \beta] - \Phi[\alpha_1 - x_i \beta] = 1 - \Phi[x_i \beta - \alpha_2] - 1 + \Phi[x_i \beta - \alpha_1] = \Phi[x_i \beta - \alpha_1] - \Phi[x_i \beta - \alpha_2]$$

$$\Pr(y_i = 1) = \Pr(y_i^* > \alpha_2) = \Pr(x_i \beta + \varepsilon_i > \alpha_2) = \Pr(\varepsilon_i > \alpha_2 - x_i \beta) = 1 - \Phi[\alpha_2 - x_i \beta] = \Phi[x_i \beta - \alpha_2]$$

The parameters in an ordered response model can be estimated by maximum likelihood:

$$\log(L(\beta, \alpha_1, \alpha_2, \alpha_3)) = \sum_{i=1}^n \sum_{j=1}^m y_{ij} \log(p_{ij}) = \sum_{i=1}^n \log(p_i y_i)$$

20 Madden (2009)

21 Ordinance n.º 91/2013

It would be desirable to include other characteristics of the physicians such as sex, age and education level. Unfortunately such information was not available. Unordered multinomial models could also be used, but to take into account the ordering makes this model more parsimonious. P is the probability of the outcome.

To ensure that $0 \leq p \leq 1$ it is natural to specify $F(\cdot)$ to be a cumulative distribution function.²² In the case of the probit, this cumulative distribution function will be the standard normal one. The explanatory variables are the current hospital where the physician works, the number of prescriptions prescribed by him/her and his/her medical specialty. The aim is to understand which factors influence physicians' prescribing patterns. Given the small number of observations in what concerns rheumatism, the model will only focus on HIV and oncology pharmaceutical products' prescriptions. Since the ordered probit is constructed based on the results attained by the relative distribution, all physicians prescribing less than four times were excluded.

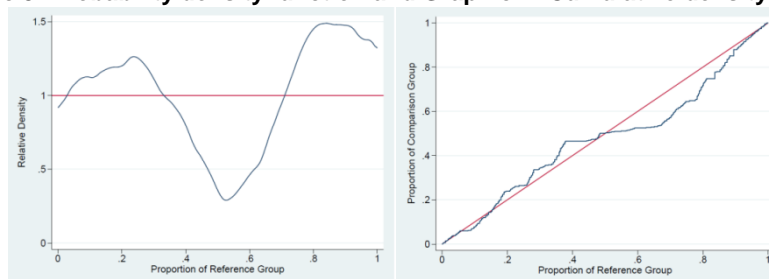
4- Results

4.1- HIV

4.1.1- Comparisons among physicians

This database includes 176 physicians prescribing HIV-related pharmaceutical products. However, only those prescribing more than four times were included in this analysis. Each physician's expenditure was compared with his colleagues' expenditure. Afterwards each graph was analysed as to conclude about the physician's expenditure: physicians incurring into higher or lower expenditure levels than the norm, in comparison to their colleagues. HSJ, HSAC and HCC are the hospitals whose specialty is HIV cases and therefore, register a higher number of cases and a higher overall expenditure. Analysing a specific physician as an example:

Graphic 3- Probability density function and Graphic 4- Cumulative density function



By looking at the probability density function, one sees that this physician shows a big discrepancy compared to his colleagues from around the 38th to the 70th percentile, showing a higher expenditure, on average, than his colleagues. However, from the 0th to the 37th and from the 70th percentile on, it shows, on average, a lower expenditure than his colleagues. This means that this physician in particular prescribes a higher quantity of less expensive pharmaceutical drugs than his colleagues, but also a higher quantity of more expensive pharmaceutical products.

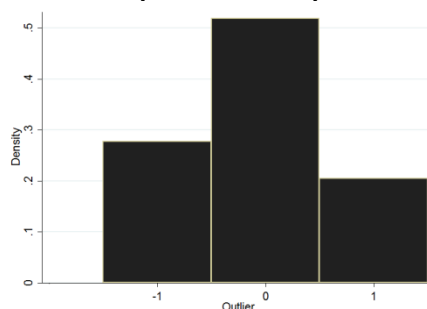
By looking at all the computed graphs, it was notorious that there are many different patterns of prescription thus, it is not possible to establish a typology. Following the visual analysis of the graphs obtained in the relative distribution and the Kolmogorov-Smirnov test, physicians were separated in three different groups. Let -1 represent physicians incurring into lower expenditure levels than the norm, in comparison to their colleagues, 0 represent those prescribing values according to the norm, and 1 represent those incurring into higher expenditure levels.

²² Cameron et al. (2005)

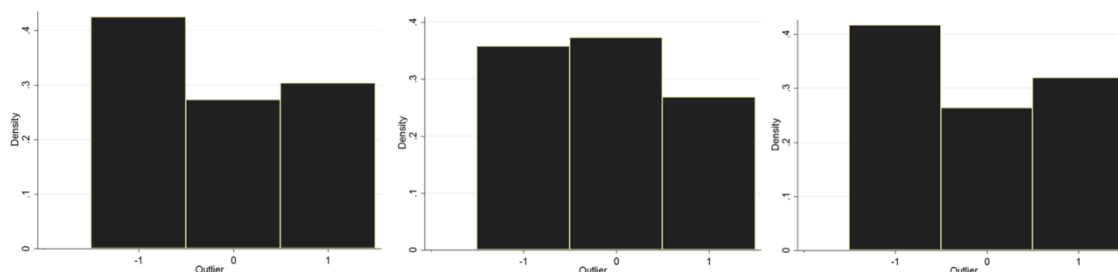
Table 1 – Relative distribution results after the Kolmogorov-Smirnov test

Nº of physicians	Overall period	1st and 2nd Quarter 2012	3rd and 4th Quarter 2012	1st and 2nd Quarter 2013
Above the norm outlier	28	20	18	23
Below the norm outlier	36	27	24	30
According to the norm	28	18	25	19
Total	135	65	67	72

Considering this, histograms were computed, taking into account that data is discrete and that the each class width is one, and therefore the density equals the relative frequency (since the density is the quotient of the relative distribution with the class width). Seeing the overall period of analysis (six quarters), the histogram below shows the density of the physicians' expenditure levels. One can conclude that the class with the highest density was the one that includes physicians incurring into expenditures according to the norm (0,5), followed by those incurring into lower expenditure levels than the norm (approximately 0,3) and lastly, by the ones incurring into higher expenditure levels (approximately 0,2).

Graphic 5- Overall period

Analysing semester by semester, one can see that the densities among each class in the first semester of 2012 and in the first semester of 2013 look a-like, although in the latter one there is a higher density of physicians with an above the norm expenditure level.

Graphic 6- 2012 1st and 2nd Quarters, Graphic 7- 2012 3rd and 4th Quarters and Graphic 8- 2013 1st and 2nd Quarters

To analyse prescription patterns per hospital where each physician was prescribing, in the overall period, histograms were computed. However, one should note that in each quarter, the average, which is the reference point, changes. In MAC and the density of physicians that are outliers prescribing below the average's norm, in comparison to their colleagues is the greatest. Those prescribing in both HSAC and HSJ, have the same quantity of physicians prescribing on the norm and above it. One should also highlight that those prescribing in HSJ, HCC and HSAC; HSJ, HDE and in HSJ, HSM were never outliers, when considering the overall period).²³ It should be highlighted that in the specific case of HSJ, physicians dealing with HIV cases gather and make prescribing decisions together.²⁴ Across the three semesters, this hospital shows great density of physicians incurring into expenditure levels below the norm, in comparison to their colleagues (0,6; 0,4 and 0,6 respectively). Thus, there is a great variation in prescribing patterns across hospitals.

²³ More details are available in the extra annexes, in the C.1.2. section

²⁴ According to the Centro Hospitalar de Lisboa Central, physicians prescribing in HSJ have meetings where they decide together which pharmaceutical products to prescribe to each patient

Histograms across medical specialties were also computed. In what concerns the medical specialty of the physicians prescribing HIV related pharmaceutical products, the assumption is that the “type” of patients is not related to the specialty of the physician. Considering the “extreme” cases²⁵, specialists in *Dermatovenereology* were only outliers below the norm, *Endocrinology* always above and *Anesthesiology*, *Internal Physician*, *Clinical Pathology* and *Urology* were always prescribing according to the norm.²⁶

4.1.2- Outlier physicians across semesters

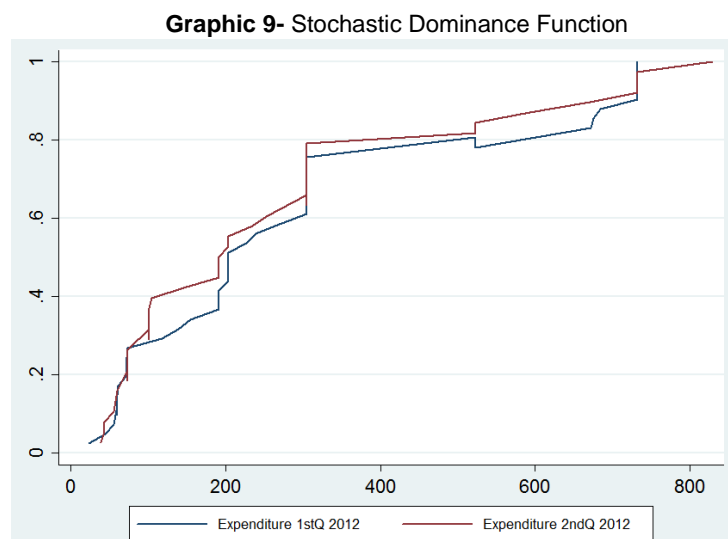
It is important to assess whether physicians are always showing the same outlier prescribing patterns in the period of analysis or if there is any change. For this purpose, the histograms below were computed for physicians with according to the norm, above the norm and below the norm expenditure levels.

In what concerns physicians identified as outliers because they incurred into higher expenditure levels, half of them were outliers only during one period. The other half was showing this behaviour throughout three semesters (27%) and two semesters (23%). Thus, approximately half of the physicians considered outliers incurring into higher expenditure levels than the norm tend to maintain their prescribing patterns across the period of observation.

Regarding physicians considered outliers with lower expenditure levels than the norm, the majority was showing this behaviour during two semesters, followed by those displaying this behaviour one semester.

4.1.3- Expenditure levels across semesters

A stochastic dominance approach was used to compare each physician’s expenditure level between two different periods. Consecutive quarters were analysed and compared, throughout a graphical analysis. The graph below illustrates this approach:



This figure represents a particular medical physician and his/her expenditure levels in the 1st quarter of 2012 and on the 2nd quarter of 2012. One can see that the distribution of the 2nd quarter of 2012 is everywhere above the distribution of the 1st quarter of 2012. Therefore, the distribution of the 1st quarter of 2012 first order stochastically dominates the one of the 2nd quarter of 2012. Assuming that the benefits are the same, lower expenditure levels are considered better. Therefore, in this specific case, expenditure with pharmaceutical products increased from one quarter to the other and the distribution developed in a undesired way.

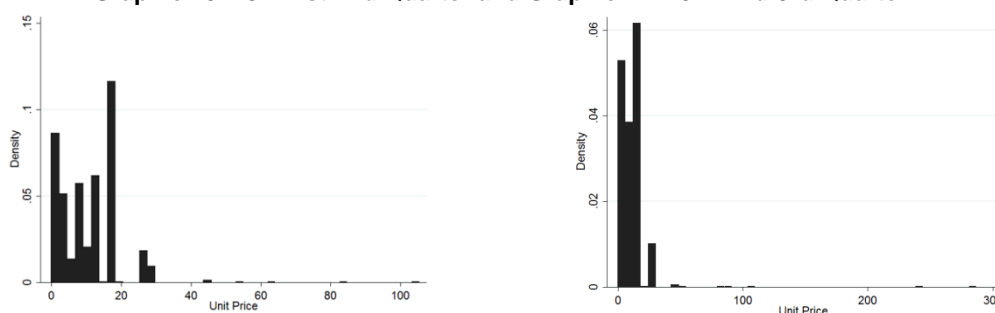
Being $D_{it} = 1$ if t stochastically dominates $t-1$ (the expenditure level in t is lower than in $t-1$) and $D_{it} = 0$ if $t-1$ stochastically dominates t (the expenditure level in $t-1$ is lower than in t).

²⁵ Hospitals which only registered outliers that incur into expenditure levels above or below the norm

²⁶ More details are available in the extra annexes, in the C.1.3. section

For the majority of the physicians prescribing HIV related pharmaceutical products in the first semester of 2012, the distribution of the 1st quarter of 2012 first order stochastically dominates the one of the 2nd quarter of 2012. The expenditure level of HIV related pharmaceutical products has, therefore, increased from the 1st to the 2nd quarter of 2012 for more than a half of medical physicians. From the 2nd to the 3rd quarter of 2012, approximately more than a half of medical physicians increased their expenditure levels in pharmaceuticals. The same pattern occurs from the 3rd to the 4th quarter of 2012 and well as for the 1st to the second quarter of 2013. The period from the 4th quarter of 2012 to the 1st quarter of 2013 was an exception, in the sense that there were more physicians incurring into lower expenditure levels in the latter period (the 1st quarter of 2013). Such event may be related to the fact that since August (3rd quarter of 2012) physicians started to receive feedback about their and the total expenditure level in pharmaceuticals. It may also have to do with the fact that, since the 1st of March of 2013, the reference prices for pharmaceutical products changed.²⁷ The newly selected countries were chosen based on their lower pharmaceutical products' price.²⁸ The 1st semester of 2013 is the one that shows a higher discrepancy between the two quarters, with the majority of physicians incurring into higher expenditure levels in the 2nd quarter. Thus, one can conclude that expenditure levels have been increasing over time for most physicians, except in one period. However, it is not possible to assess if this change was due to the number of treated patients since only after 2013 all the hospitals were using the same database. Thus, different patients from different hospitals may, in 2012, have the same patient number. Nevertheless, one can look at the quantity and unit price of the pharmaceuticals prescribed. In what concerns the unit price, there is an increase in the prescription of very costly pharmaceutical products since the first two quarters of 2012 (although they represent really small densities). In what concerns the quantity, it seems that there are no major changes across time.²⁹

Graphic 10- 2012 1st-2nd Quarter and Graphic 11- 2012 2nd-3rd Quarter



4.1.4- Significant Characteristics

The explanatory variables include the number of prescriptions made by each physician, the medical specialty (a dummy variable with 1 being Infectiology and General Medicine and 0 the other specialties, since these are the two specialties which physicians register a larger number of episodes) and the hospital where he/she prescribes (dummy variables, being the omitted variable HSM). As it was previously stated in the data description, it is not possible to follow a patient by his/her hospital number. Neither patients' nor physicians' characteristics (e.g. age, education and job) are available. It might be possible that some of these variables could be significant into explaining different expenditure levels.

²⁷ Ordinance n.º 91/2013

²⁸ According to the Ordinance n.º 91/2013: "Atendendo à necessidade de racionalização dos encargos públicos com medicamentos, o conjunto de países selecionados atende ao critério de países europeus com nível de preços de medicamento mais baixos"

²⁹ Further details can be found in Section C.1.4. of the Extra Annexes

Table 2 – Ordered probit results

Variables	All Period of Analysis		2012 1st-2nd Quarter		2012 3rd-4th Quarter		2013 1st-2nd Quarter	
	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values
N° Prescriptions	0,0000867	0,008*	7.69e-06	0,875	0,0004609	0,080	0,00034	0,059
Specialty	0,2682441	0,221	0,7943984	0,031*	0,6115661	0,145	0,3375354	0,407
Hospital:								
Capuchos	0,9882828	0,003*	-0,883198	0,085	1,350033	0,003*	1,274743	0,006*
Curry e Cabral	0,2699523	0,287	-0,5985265	0,274	-1,090596	0,216	0,3899994	0,274
S. José	-0,203284	0,381	1,306204	0,010*	0,8859	0,048	1,16593	0,020*
Estefânia	1,837315	0,00*	1,093221	0,205	2,447538	0,004*	1.318.677	0,090
Alfredo da Costa	1,423578	0,224	5,448322	0,981	-	-	-3,886876	0,989

Prescribing in HSAC is significant in the overall period of analysis and in two semesters, associated with positive coefficients. The medical specialty of the physician prescribing is significant in the 1st semester of 2012.

In the unordered probit case, the sign of each coefficient gives the direction of the effect, but not its marginal effect.³⁰ The marginal effects of changes in the explanatory variables are given by: $\frac{\partial \Pr[y_i=j]}{\partial x_i} = \{F(\alpha_j - x_i\beta) - F(\alpha_j - x_i\beta)\}\beta$.³¹

Table 3 – Probability of being an outlier with a below the norm expenditure level - Pr[yi = -1]

Conditional marginal effects	All Period of Analysis		2012 1st-2nd Quarter		2012 3rd-4th Quarter		2013 1st-2nd Quarter	
	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values
N° Prescriptions	-9,36e-06	0,353	-0,0000195	0,462	-0,0001081	0,082	-0,0001644	0,671
Specialty	-0,1045788	0,166	-0,2799172	0,306	-0,1911433	0,136	-0,0935281	0,727
Hospital:								
Capuchos	-0,250794	0,004*	-0,4277048	0,268	-0,3351039	0,010*	-0,2820507	0,672
Curry e Cabral	0,1529581	0,123	-0,0341375	0,856	-0,1393592	0,399	-0,2007085	0,681
S. José	-0,951416	0,265	-0,0980718	0,556	-0,240762	0,061	-0,1279824	0,698
Estefânia	-0,124534	0,538	-0,570009	0,331	-0,7038155	0,008*	-0,2268341	0,701
Alfredo da Costa	0,158819	0,402	1,450214	0,988	-	-	1,887503	0,986

Table 4 - Probability having an expenditure level according to the norm - Pr[yi = 0]

Conditional marginal effects	All Period of Analysis		2012 1st-2nd Quarter		2012 3rd-4th Quarter		2013 1st-2nd Quarter	
	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values
N° Prescriptions	1,71e-06	0,452	3,27e-06	0,952	0,0000175	0,374	0,0000612	0,980
Specialty	0,018612	0,346	0,0469594	0,952	0,0309032	0,430	0,0348422	0,980
Hospital:								
Capuchos	0,0458035	0,247	0,0717526	0,952	0,0541781	0,372	0,1050728	0,980
Curry e Cabral	0,0279353	0,329	0,005727	0,954	0,022531	0,540	0,747703	0,980
S. José	0,0173761	0,403	0,0164527	0,952	-0,240762	0,061	0,0845029	0,980
Estefânia	0,022744	0,580	0,0956258	0,952	0,1137898	0,375	0,0476775	0,980
Alfredo da Costa	-0,0290056	0,492	-0,2432906	0,990	-	-	-0,7031547	0,992

Table 5 – Probability of being an outlier with an above the norm expenditure level - Pr[yi = 1]

Conditional marginal effects	All Period of Analysis		2012 1st-2nd Quarter		2012 3rd-4th Quarter		2013 1st-2nd Quarter	
	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values	Coefficients	P-values
N° Prescriptions	7,65e-06	0,354	0,0000162	0,700	0,0000906	0,097	0,0001031	0,971
Specialty	0,0834274	0,159	-0,1911433	0,136	-0,0935281	0,727	0,0586859	0,971
Hospital:								
Capuchos	0,2049901	0,005*	0,355952	0,674	0,2809254	0,011*	0,1769779	0,971
Curry e Cabral	0,1250225	0,124	0,0284105	0,866	-0,2007085	0,681	0,1259382	0,971
S. José	0,0777654	0,266	0,081619	0,719	-0,1279824	0,698	0,0803049	0,971
Estefânia	0,1017897	0,538	0,4743829	0,680	-0,2268341	0,701	0,1423312	0,971
Alfredo da Costa	-0,1298126	0,400	-1,206922	0,987	-	-	-1,184348	0,976

The main result is that physicians prescribing in HSAC are more likely to be outliers incurring into expenditures above the norm (20 percentage points in the overall period and 28 percentage points in the 2nd semester of 2012) and less likely to be outliers incurring into expenditures below the norm (25 percentage points in the overall period and 34 percentage points in the 2nd semester of 2012), in comparison to those prescribing in HSM (the omitted variable).

³⁰ For further details see section B.1. of the Extra Annexes

³¹ For further details see section C.1.5. of the Extra Annexes

4.2- Oncology

4.2.1- Comparisons among physicians

This dataset contains 193 physicians prescribing oncology related pharmaceutical drugs. However, to use the relative distribution model one needs to only include those that have prescribed at least more than four times, therefore only 135 physicians were analysed.

The density of outliers incurring into higher expenditure levels than the average seems to be more or less constant across the three semesters.

HSJ shows a decrease in outlier physicians prescribing levels above the norm throughout the three semesters. HSAC consistently has a great density of outlier physicians prescribing below the norm expenditure level. On the other hand, HCC displays great density outlier physicians prescribing levels above the norm throughout the three semesters.³²

There is also great divergence between medical specialties.³³ Considering the overall period, there are six specialties which physicians are all outliers prescribing above the norm level and one where they are not outliers. However, in what concerns cancer, different medical specialties are related to several types of cancer.³⁴ Different types of cancer require different a treatment so, costs are not comparable across the different pathologies.

4.2.2- Outlier physicians across semesters

In what concerns physicians identified as outliers because they have incurred into higher expenditure levels, on average, than their colleagues, most of them were showing this behaviour throughout the three semesters. By analysing the physicians that were identified as outliers that incurred into lower expenditure levels, than their colleagues, the majority also showed this pattern during the three semesters. Thus, most physicians have been maintaining their prescribing patterns unchanged.³⁵

4.2.3- Expenditure levels across semesters

The expenditure level of oncology related pharmaceutical products has increased from the 1st to the 2nd quarter of 2012, from the second to the third quarter of 2012 and from the 4th quarter of 2012 to the 1st quarter of 2013. On the other hand, from the 3rd to the 4th quarter of 2012 and from the 1st to the 2nd quarter of 2013 there is a greater density for physicians incurring into lower expenditure levels in the latter period. This might be related to the fact that since August (in the 3rd quarter of 2012) physicians started to have feedback about their and the total expenditure level in pharmaceuticals. Furthermore, since the 1st of March of 2013, the reference prices for pharmaceutical products changed. The newly selected countries were chosen based on their lower pharmaceutical products' price. With respect to the quantity and the unit price of the pharmaceutical products prescribed, there are no major changes across these 1,5years. In some periods, there is a small density of pharmaceutical products which unit price is extremely high (more than 1000€), which may be related to very acute episodes.

4.2.4- Significant Characteristics

In what concerns cancer, different medical specialties are related to several types of cancer, as it was stated before. Across each semester and in the overall period the only significant variables are those concerning the physician's medical specialty. However, one cannot do such comparison. In what concerns the

³² More details are available in the extra annexes, in the C.2.2. section

³³ More details are available in the extra annexes, in the C.1.3. section

³⁴ For further details see section D.1.

³⁵ More details are available in the extra annexes, in the C.2.4. section

number of pharmaceutical products that were prescribed and the hospital where they were prescribed, no coefficient is significant.³⁶

4.3- Rheumatism

4.3.1- Comparisons among physicians

In what concerns physicians prescribing for rheumatology, they are concentrated in three out of the six hospitals: HCC, HSAC, and HDE. HCC seems to have more physicians incurring into higher expenditure levels than the norm. This pattern is consistent throughout the different semesters. The opposite happens with HSAC and HDE, which have a higher proportion of physicians incurring into lower expenditure levels than the norm.³⁷

General Medicine is the medical specialty with more physicians with expenditure levels above the norm, when considering the overall period. In what concerns the specialty of the medical physicians below the norm, the results show that the majority are specialists in *Anesthesiology* and *Pediatrics*. This is true both for the overall period and for each semester.³⁸

4.3.2- Outlier physicians across semesters

Regarding physicians identified as outliers because they incurred into higher expenditure levels, on average, than their colleagues, the majority were showing this behaviour throughout two semesters followed by those showing this behaviour during three semesters. There was no physician showing this pattern only in one semester.

By analysing the physicians that were identified as outliers that incurred into lower expenditure levels than their colleagues, the majority showed this pattern during one semester, followed by those showing this behaviour during three semesters.

4.3.3- Expenditure levels across semesters

Expenditure levels in pharmaceuticals have been decreasing in 3 out of 5 periods (from the first to the 2nd quarter of 2012, from the 2nd to the 3rd quarter of 2012 and from the 4th quarter of 2012 to the 1st quarter of 2013). Thus, expenditure levels have been decreasing over time for most physicians, except for two periods. Regarding the unit price, there is an increase in the prescription of very costly pharmaceutical products since the first two quarters of 2012. In what concerns the quantity, it seems that there are no major changes across time.³⁹

5- Conclusion

Regarding HIV, there is a great variation in prescribing patterns across hospitals. One should highlight the case of HSJ, where physicians dealing with HIV cases gather and make prescribing decisions together. During the observation period, this hospital shows a high density of physicians incurring into expenditure levels below the norm, in comparison to their colleagues. Concerning the permanence/change of the prescribing behaviour by physicians, it was found that approximately half of the physicians that were considered as outliers, incurring into higher expenditure levels than the norm tend to maintain their prescribing patterns across these 1,5 years. Expenditure levels in pharmaceutical products have been increasing over time for most physicians, except in one period. With respect to the unit price, there is an increase in the prescription of very costly pharmaceutical products since the first two quarters of 2012, however the quantity prescribed did not register major changes across time. The unordered probit model consistently shows

³⁶ More details are available in the extra annexes, in the C.2.5. section

³⁷ More details are available in the extra annexes, in the C.3.2. section

³⁸ More details are available in the extra annexes, in the C.3.3. section

³⁹ More details are available in the extra annexes, in the C.3.4. section

that physicians prescribing in HSAC have a higher probability of being outliers incurring into higher expenditures than the norm and a lower probability of being outliers incurring into lower expenditure levels than the norm, in comparison to those prescribing in HSM (the omitted variable).

Physicians prescribing oncology-related pharmaceutical products seem to have very different prescription patterns according to the hospital where they prescribe. HSJ shows a decrease in outlier physicians prescribing levels above the norm throughout the three semesters. HSAC consistently has a great density of outlier physicians prescribing below the norm expenditure level. HCC displays a great density of outlier physicians prescribing levels above the norm throughout the three semesters. Regardless of the hospital, most physicians have been maintaining their prescribing patterns unchanged. There are also great differences across medical specialties. However, different types of cancer require different treatments and thus, costs are not comparable across different pathologies. As to the stability of outlier behaviour, most physicians have not been changing their prescribing patterns. With respect to the quantity and the unit price of the pharmaceutical products prescribed, there are no major changes across the observation period. In some periods, pharmaceutical products with a extremely high unit price (more than €1.000, while almost all the others cost less than €500) are prescribed. However, they only represent a small density of the overall products. This may be related to very acute episodes, which require costlier products.

In what concerns rheumatism, physicians are concentrated in three out of the six hospitals: HCC, HSAC, and HDE. HCC seems to have more physicians incurring into higher expenditure levels than the norm. This pattern is consistent throughout the different semesters. The opposite happens with HSAC and HDE, which have a higher proportion of physicians incurring into lower expenditure levels than the norm. Most of outlier physicians that incur in higher expenditures maintain their prescribing behaviour throughout this period. In contrast, those that incur in lower expenditure levels than the norm do not maintain their prescribing behaviour. Expenditure levels have been decreasing over time for most physicians, except for two periods. Regarding the unit price, there is an increase in the prescription of very costly pharmaceutical products since the first two quarters of 2012. In what concerns the quantity, it seems that there are no major changes across time.

Overall, there is great divergence of prescribing patterns across physicians, the hospitals where they prescribe and their medical specialties. Physicians tended to maintain their prescribing patterns across these 1,5 years, whether they were outliers incurring into higher or lower expenditure levels than the norm and even after the introduction of the individual feedback. With respect to the evolution of total costs of pharmaceutical products, the price decrease of pharmaceuticals does not seem to have had a permanent impact in expenditure levels.

The results stated above show a discrepancy in relation to the Law of Large Numbers. This law states that, on average, each physician should treat similar patients, in terms of health status and related expenditure. Hence, it should be expected that prescription patterns of pharmaceuticals would be similar across hospitals and across physicians with the same medical specialty. Therefore, there is room to reduce hospital costs of pharmaceutical products, by better practice of physicians.

Nevertheless, it would be valuable to have information regarding the treated patients. Then one could assess whether physicians that are always incurring into expenditure levels above the norm were treating patients that could justify the use of costlier pharmaceutical products or not (since in some cases the application of the Law of Large Numbers might be unsuitable). If it would be found that such patients did not require costlier pharmaceutical products, then one could conclude that these physicians should change their prescribing behaviour while the other ones (incurring into below the norm expenditure levels) should be emphasised as having a better practice.

Future research should include a method, such as the differences-in-differences one, to evaluate whether physicians change their prescription patterns when they are provided with better information regarding their prescription behaviour. Furthermore, in the specific case of HIV, it would be interesting to assess the impact of physicians prescribing pharmaceutical products together, by analysing the case of HSJ.⁴⁰ More-

⁴⁰ See Extra Annexes, section B.2. for further details

over, having information about the patients' as well as physicians' characteristics (e.g. sex, age and education level.) could be beneficial for further research.

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