A sampling design for a Sentinel General Practitioner Network

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Introduction

Sentinel networks are an increasingly useful tool in epidemiological surveillance and research. Since networks were first set up in Holland1 and the United Kingdom2 in the 1960s, their numbers have grown worldwide and they are used in fields as diverse as communicable3–7 and non-communicable disease surveillance8–11 and economic assessment studies12.

In sentinel networks, a group of physicians reports cases of diseases or processes of interest, on a regular basis, generally weekly.

The Spanish National Health System offers universal health care coverage, and consequently the majority of the population tends to use public medical services. This provides a good opportunity to study population-based primary health care demand through a probabilistically obtained sample of physicians. The population attended by the sentinel physicians must be representative of the general population, if the information obtained by such a network is intended to be at all generalisable to the latter13.

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At present, sentinel networks are in place in most of Spain's Autonomous Regions and under different names share objectives and methodological aspects adapted to the Spanish National Health System. Since first initiatives took with the sentinel networks in Castilla y León and the Basque Country, the rest of Autonomous Regions have developed network for the surveillance of several different diseases, and they have tackled common projects.

The Madrid Autonomous Region Sentinel General Practitioner Network came into operation in 1991, and the reporters are volunteer physicians. Although the sampling in 1991 took into account geographic and demographic criteria, the final selection was done by requesting the collaboration from all physicians. Since its creation the network has suffered leavings and substitutions of reporting physicians; these factors, along demographic changes in population, make necessary a reviewing of the design and composition of the network.

The stated aim of this study was to develop a design for probabilistic sampling of reporting physicians in the Madrid Autonomous Region Sentinel General Practitioner Network (SGPN), to ensure the representativeness of the population covered by the network, in order to generalize the design for similar populations.

Methods

Data source: Information on general practitioners and paediatricians and their patients, broken down by age and sex, was obtained from the Madrid Health Institute's Health-Card Database for the 2005 period.

The designated geographic health care unit of reference was the Basic Health Zone (BHZ). There are 247 Basic Health Zones in the Madrid Autonomous Region, each of which represents the catchment area of a given Health Centre and has a median population size of 21,103 persons (range 2522–72,662).

Demographic, socio-cultural and socio-work variables were used to describe each BHZ (Table 1). Variables were obtained from the data base of the Madrid Statistics Institute for municipal register of 2005. To reduce the number of variables without losing relevant information, a principal components analysis was performed, using the Varimax method. When extracting factors, account was taken of their eigenvalues and the values obtained in the matrix of rotated components.

Once the factors had been obtained, a cluster analysis was performed to group the 247 BHZs into homogeneous strata. Distance between units (BHZs) was measured in terms of the squared Euclidean distance, and the grouping method chosen was Ward's Method, because it is a hierarchical agglomerative method and it tends to make compact, similar sized clusters. The clustering history and dendrogram were then used to obtain the adequate number of clusters after different options were studied. The available statistical package used didn't offer other tools to determinate the number of clusters, but some other can be used, as pseudo-f, pseudo-t-square or cubic cluster criterion (CCC).

To test whether the strata were homogeneous and mutually different, we conducted a discriminant analysis, with the stratum or cluster of membership as the grouping variable, and the 5 factors yielded by the principal components analysis as the independent variables. Using variance analysis and Wills' Lambda statistic, we evaluated whether the factors employed were different in the strata by means of the equality of means test. To assess the discriminant power of the discriminant functions obtained, we examined the eigenvalues and canonical correlation coefficients of such functions (which, when squared, express the percentage of variability explained by the differences between groups), and performed Wills' Lambda test for each function. The latter can also be converted into a Chi-squared test, with its associated p-value of statistical significance, to verify the null hypothesis that there are no differences among strata by reference to the factors used. In addition, each BHZ's real stratum of membership was compared against its a priori probability of classification, through crossed validation, thereby enabling us to observe how many BHZs had been erroneously classified.

The number of persons needed to estimate the incidence rate of the most infrequent process of those monitored in Madrid in 2004 (varicella, herpes zoster, influenza and asthma) was ascertained by assuming that this disease followed a Poisson distribution and that the expected number of cases was greater than 10. Herpes zoster was the selected disease for this

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Table 1

<table>
<thead>
<tr>
<th>Demographic variables</th>
<th>Factor 1</th>
<th>Factor 2</th>
<th>Factor 3</th>
<th>Factor 4</th>
<th>Factor 5</th>
</tr>
</thead>
<tbody>
<tr>
<td>Percentage of women</td>
<td>0.732</td>
<td>0.486</td>
<td>0.130</td>
<td>0.037</td>
<td>-0.042</td>
</tr>
<tr>
<td>Proportion of subjects aged under 15 years</td>
<td>-0.725</td>
<td>0.100</td>
<td>-0.122</td>
<td>0.120</td>
<td>0.613</td>
</tr>
<tr>
<td>Proportion of subjects aged over 64 years</td>
<td>0.981</td>
<td>-0.019</td>
<td>0.071</td>
<td>-0.061</td>
<td>-0.046</td>
</tr>
<tr>
<td>Proportion of subjects aged over 84 years</td>
<td>0.829</td>
<td>0.232</td>
<td>0.231</td>
<td>-0.015</td>
<td>-0.066</td>
</tr>
<tr>
<td>Indenpendency index (quotient between population aged under 15 years and over 64 years, and population aged 15 to 64 years)</td>
<td>0.838</td>
<td>0.033</td>
<td>-0.011</td>
<td>-0.004</td>
<td>0.375</td>
</tr>
<tr>
<td>Replacement index (quotient between population aged 15 to 39 years, and population aged 40 to 64 years)</td>
<td>-0.325</td>
<td>-0.296</td>
<td>0.050</td>
<td>0.782</td>
<td>0.299</td>
</tr>
<tr>
<td>Ratio of progressiveness (quotient between population aged 0 to 4 years, and population aged 5 to 9 years (x 100))</td>
<td>0.027</td>
<td>0.199</td>
<td>0.055</td>
<td>0.892</td>
<td>-0.140</td>
</tr>
<tr>
<td>Percentage of foreigners</td>
<td>0.265</td>
<td>0.076</td>
<td>0.921</td>
<td>0.116</td>
<td>-0.072</td>
</tr>
<tr>
<td>Percentage of exclusively foreign homes</td>
<td>0.068</td>
<td>0.007</td>
<td>0.943</td>
<td>-0.004</td>
<td>-0.090</td>
</tr>
<tr>
<td>Percentage of mixed homes</td>
<td>-0.001</td>
<td>0.564</td>
<td>0.671</td>
<td>-0.061</td>
<td>0.305</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Socio-cultural variables</th>
<th>Factor 1</th>
<th>Factor 2</th>
<th>Factor 3</th>
<th>Factor 4</th>
<th>Factor 5</th>
</tr>
</thead>
<tbody>
<tr>
<td>Percentage of separated or divorced subjects</td>
<td>0.529</td>
<td>0.367</td>
<td>0.478</td>
<td>0.117</td>
<td>0.162</td>
</tr>
<tr>
<td>Percentage of illiterate subjects</td>
<td>0.152</td>
<td>-0.796</td>
<td>-0.088</td>
<td>-0.129</td>
<td>0.387</td>
</tr>
<tr>
<td>Percentage of subjects with elementary or primary education</td>
<td>-0.047</td>
<td>-0.959</td>
<td>-0.118</td>
<td>0.013</td>
<td>-0.141</td>
</tr>
<tr>
<td>Percentage of subjects with secondary education (school leaving certificate)</td>
<td>-0.481</td>
<td>0.698</td>
<td>-0.175</td>
<td>0.240</td>
<td>-0.146</td>
</tr>
<tr>
<td>Percentage of subjects with university education (diplomas, degrees, doctorates)</td>
<td>0.120</td>
<td>0.930</td>
<td>0.140</td>
<td>-0.080</td>
<td>0.132</td>
</tr>
</tbody>
</table>

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<thead>
<tr>
<th>Socio-work variables</th>
<th>Factor 1</th>
<th>Factor 2</th>
<th>Factor 3</th>
<th>Factor 4</th>
<th>Factor 5</th>
</tr>
</thead>
<tbody>
<tr>
<td>Percentage of unemployed persons</td>
<td>-0.173</td>
<td>-0.721</td>
<td>0.074</td>
<td>0.286</td>
<td>0.133</td>
</tr>
<tr>
<td>Percentage of pensioners</td>
<td>0.970</td>
<td>-0.108</td>
<td>0.103</td>
<td>-0.046</td>
<td>-0.066</td>
</tr>
<tr>
<td>Percentage of gainfully employed subjects</td>
<td>-0.906</td>
<td>0.161</td>
<td>-0.047</td>
<td>0.235</td>
<td>0.156</td>
</tr>
<tr>
<td>Percentage of housewives</td>
<td>-0.074</td>
<td>0.811</td>
<td>-0.281</td>
<td>-0.259</td>
<td>-0.155</td>
</tr>
</tbody>
</table>
calculation, as it was the most infrequent among the monitored processes. Assuming all these factors, the 95% confidence interval of the needed number of cases could be ascertained by the next approximation:

\[ A \pm Z_{1-\alpha/2} \cdot \sqrt{a} \]

The accuracy of this estimation is the relative accuracy (\( p \)) that we want to estimate the rates, and multiplied by the needed number of cases:

\[ Z_{1-\alpha/2} \cdot \sqrt{a} = p \cdot a \]

So, the needed number of cases to estimate a rate with a determined accuracy will be:

\[ d = \frac{Z_{1-\alpha/2}^2}{p^2} \]

We calculated the number of cases needed to estimate a rate with an accuracy of 10%, and an \( \alpha \) of 0.05, taking into account a design effect of 1.5\(^2\). Once the necessary number of cases had been established, we then obtained the population required for monitoring purposes based on the herpes zoster incidence rate in Madrid in the preceding year. The resulting population was allocated to the strata in proportion to their respective populations.

In the Spanish National Health System, Primary Health Care is given by general practitioners or paediatricians, depending on the age of patients, so it is necessary that sentinel network includes both of them. As the SGPN is made up of Primary Care general practitioners and paediatricians, the population required for monitoring purposes was divided into adult and childhood populations, according to the proportion of population aged under 15 years drawn from the 2005 Madrid Regional population register.

In each stratum, general practitioners and paediatricians were consecutively selected by simple random sampling, until the pre-established required minimum populations were reached.

The representativeness of the population monitored by the selected physicians was studied, by age group and sex, with respect to the population of the Madrid Autonomous Region, in line with the 2005 population register. To this end, the proportion of the population in each sex and age group over the total sample selected was compared to the same proportion in the Madrid Autonomous Region.

The software used was the statistical package SPSS\textsuperscript{*} 13.0.

Results

Factorial analysis

Five factors were extracted using the principal-components-analysis method. The first 4 had eigenvalues of over 1 and accounted for 81.5% of the variance. The fifth factor had an eigenvalue very close to 1. As these 5 factors would together account for 86.6% of the variance, we decided to include the fifth factor (Table 2).

The contribution of the variables to the 5 factors can be seen in the matrix of rotated components (Table 1).

Factor 1 (eigenvalue: 6.793) was related to demographic variables, registering high values in the most elderly populations, characterised by a high percentage of pensioners and a low percentage of gainfully employed subjects.

Factor 2 (eigenvalue: 5.033) was mainly related to educational variables, assuming high values in the indicators of intermediate and higher education, and low values in the percentages of:

<table>
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<tr>
<th>Table 2</th>
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<tbody>
<tr>
<td><strong>Eigenvalues of factors extracted and percentage of variance explained</strong></td>
</tr>
<tr>
<td>Factor</td>
</tr>
<tr>
<td>-------</td>
</tr>
<tr>
<td>1</td>
</tr>
<tr>
<td>2</td>
</tr>
<tr>
<td>3</td>
</tr>
<tr>
<td>4</td>
</tr>
<tr>
<td>5</td>
</tr>
</tbody>
</table>

Factor 3 (eigenvalue: 2.212) was related to variables of immigration, assuming high values in the populations with most immigrants.

Factor 4 (eigenvalue: 1.441) was related to the proportion of subjects aged under the age of 15 years.

Factor 5 (eigenvalue: 0.977) was related to the proportion of subjects with primary education and without any formal education; unemployed persons; and housewives.

Persons with primary education and without any formal education; unemployed persons; and housewives.

Cluster analysis

On completion of the cluster analysis, 14 strata were obtained. The number of BHZs in these strata ranged from 4 to 34, with a median of 18.

Discriminant analysis

Following the discriminant analysis, the Wilks’ Lambda coefficient displayed low values in the equality of means test, reflecting differences among the strata for each factor. Moreover, the F statistics for each of the resulting factors in the analysis of variance (ANOVA) registered associated p-values that were statistically significant (Table 3).

A total of 5 discriminant functions were obtained: their eigenvalues ranged from 0.90 to 4.06, and their canonical correlation coefficients registered high values, with the lowest being 0.69 and the highest 0.90. As a result, Wilks’ Lambda test yielded values very close to 0 for all of the first 4 discriminant functions (Table 4). Reference to the classification matrix showed that, according to the discriminant functions, 86.6% of the health zones were correctly classified.

Sample selection

The number of cases required to estimate an incidence rate with a relative accuracy of 10% was 384. As the herpes zoster incidence rate in Madrid in 2004 was 0.00335932 year\(^{-1}\), a population of 114,309 was thus needed. Applying a design effect
of 1.5, the necessary population was 171,464 persons, representing 3% of the Madrid Autonomous Region’s population. In view of the fact that 14.3% of the Madrid population in 2003 was under the age of 15 years, the required minimum population was 146,946 adults and 24,518 children. These populations were distributed among the 14 strata in proportion to the total populations of each stratum (Table 5).

In each stratum, general practitioners and paediatricians were consecutively selected by simple random sampling, until the pre-established required minimum populations were reached. We consecutively selected 88 general practitioners and 32 paediatricians, until the pre-established required minimum populations were reached; finally general practitioners and paediatricians selected covered populations of 154,610 and 31,336 subjects, respectively.

In the representativeness study, no statistically significant differences \( (p < 0.05) \) were observed between the sample and the Madrid Autonomous Region in terms of the proportions of the population in each age group and sex, except for the groups aged 0 to 4 and 5 to 9 years in both sexes (Fig. 1).
Discussion

One of the main difficulties of sentinel networks lies in selecting the reporting physicians in such a way that the monitored population is representative of the general population. Most sentinel networks use design or quality control tools to enhance such representativeness as far as possible\(^1\).\(^2\).\(^3\).

Indeed, these types of designs, which at times encounter difficulties posed by the great extent of territory to be covered or the dispersion of the population to be monitored, are not always a simple task\(^2\).\(^4\).\(^5\).\(^6\).\(^7\).\(^8\).\(^9\).\(^10\).\(^11\).\(^12\). The reporting physicians in many highly important, traditional networks are volunteers\(^9\).\(^10\).\(^11\).

The Madrid Autonomous Region has a considerable population that is not homogeneous, with a great degree of demographic, cultural and economic diversity. Simple random sampling is an effective technique but does not ensure representativeness. However, by dividing the population into strata by means of characteristics which inform on characteristics that differ between some population groups and others, and selecting subsamples in each stratum, the representation of all in the total sample is guaranteed\(^1\).\(^2\).\(^3\). Furthermore, if the strata can be made uniform, the accuracy of the estimates obtained will be greater\(^1\).\(^2\).\(^3\).\(^4\). The choice of a probabilistic method is always desirable, but sometimes the method of selection is limited by the available resources, the health system structure, and the Primary Health Care Services features. In the method presented in this article, physicians are the selected units, since it is not feasible to select a random sample of people attended in Primary Health Care Services; the probabilistic method is not applied to persons but physicians; so every citizen have their own likelihoods to be selected depending on their assigned physicians.

In the Madrid Autonomous Region, Basic Health Zones are units of sufficiently small size to ensure that the demographic, socio-economic and socio-cultural variables within them are homogeneous. Hence, a technique such as cluster analysis can efficiently yield strata composed of various BHZs having similar characteristics. Random selection of reporting physicians in each stratum renders this design capable of producing a sample that is representative and estimators that are unbiased and accurate. A number of physicians was selected in each stratum in such a way as to guarantee that the monitored population exceeded the pre-established minimum required.

The study's representativeness highlighted the fact that the population covered by the selected sample of physicians was representative of the population of the Madrid Autonomous Region in terms of age and sex, though in the 0–4 and 5–9 age groups, the proportion of the monitored population was significantly higher than the proportion of the population of the Madrid Autonomous Region. This was due to the fact that in the smallest strata, the minimum paediatric population required was likewise very small, and that the population monitored in these strata by the first paediatrician selected was far higher. Nevertheless, the magnitudes of the differences observed were small, and there is no reason why this should in any way compromise the validity of the results obtained by the network. The good representativeness of the selected sample according to the general population of Madrid is a positive point in the sample selection process.

The data of population assigned to each physician come from Madrid Health Institute's Health-Card Database, which includes all population; however there are a percentage of people that use private health services. It is important to take into account that this fact could to influence the sampling design by underestimating the incidence rates of the monitored diseases. This bias can be partially reduced if we monitor a larger population than theoretically defined.

One of the limitations of this design might be the fact that geographic criteria were not taken into account. The Madrid Autonomous Region is a small territory, with little geographic spread, so that any possible social, economic or cultural differences are significantly close. That is the reason of include social and demographic variables, but in case of a selection of a sample in territories with different features is necessary to consider other data as population density or geographic dispersion. Accordingly, we chose to take cluster analysis into account as a technique for obtaining homogeneous units, even though the ensuing strata may not be territories that are geographically united. The choice of a mainly adult disease as herpes zoster to pre-determine the sample size does not affect the validity neither of the design nor the obtained results, as childhood diseases currently monitored are more frequent. Although a specific process as herpes zoster has supported it, we must emphasize that the aim of this work is to set the frame of a design applicable to future studies.

Although a design effect of 1.5 was applied, when a multistage method is designed, a higher design effect should be used if the design is very complex.

Another limitation of the study could be the fact of a high percentage of selected physicians refused to collaborate, as a voluntariness bias could be introduced and the representativeness of sample would be affected. Nevertheless, if a physician refused to participate, another one would be selected by simple random sampling in the same stratus. It is not foreseeable a high number of rejections, as Primary Health Care physicians keep a strong commitment with their labour, and the burden derived from the work in the sentinel network is not expected to be excessive.

In view of the fact that sentinel networks are being increasingly used in epidemiological surveillance, suitable designs are required to ensure attainment of their designated goals.

In order to be able to generalise the results obtained, a feature to be sought is the representativeness of the network vis-à-vis the general population. Cluster analysis is a method that answers this need. To this end, it is necessary to have sampling units that are as unaggregated as possible, together with information on demographic, social, economic or cultural factors of interest.

Furthermore, selection of physicians by probabilistic methods, as in this study, affords the chance of ascertaining the degree of accuracy of the estimates obtained, thereby adding value to the data collected by sentinel networks.

Acknowledgements

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