Identifying geographical regions serviced by hospitals to assess laboratory-based outcomes

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ABSTRACT
Objective: To define geographical regions (forward sortation areas; FSAs) in Southwestern Ontario, Canada from which patients would reliably present to a hospital with linked laboratory data if they developed adverse events related to medications dispensed in outpatient pharmacies.

Design: Descriptive research.


Participants: Patients aged 66 years and older who received an outpatient prescription for any drug and presented to the emergency department in the subsequent 120 days.

Main outcome measure: The proportion of patients in a given FSA presenting to an emergency department at a hospital with linked laboratory data versus a hospital without linked laboratory data. To be included in the catchment area at least 90% of emergency department visits in an FSA must have occurred at laboratory-linked hospitals in a given year.

Results: Over the study period, there were 649,713 emergency department visits by patients with recent prescription claims from pharmacies in 1 of 118 FSAs. In total, 141,302 of these patients presented to an emergency department at a laboratory-linked hospital. For the year 2003, 12 FSAs met our criteria to be in the catchment area and this number grew to 25 FSAs by the year 2009.

Conclusions: The relevant geographical regions for hospitals with linked laboratory data have been successfully identified. Studies can now be conducted using these well-defined areas to obtain reliable information on the incidence and absolute risk of presenting to hospital with laboratory abnormalities in older adults dispensed commonly prescribed medications in outpatient pharmacies.

BACKGROUND
Linked health administrative databases are powerful tools for conducting population-based observational studies. Initially intended for administrative purposes, the use of these databases has become increasingly popular in the field of health services research.1 Linked databases contain a wide range of patient-related information at various levels (eg, national or provincial level). Typically, records include information on patient demographics, hospitalisations and ambulatory visits identified by diagnostic or procedural codes assigned during the encounter, and outpatient drug dispensations from pharmacies.2

Postmarketing drug studies have become important in understanding the real-world impact of commonly used medications in outpatient settings.3–6 Drug safety studies are especially useful when exploring the effect of a drug on well-coded outcomes, such as skeletal fracture and acute myocardial infarction. Diagnostic codes for these outcomes are highly accurate with a sensitivity ≥89% and
positive predictive value ≥87%. Certain drugs can also lead to adverse laboratory-based disorders such as hypo-natraemia, hyperglycaemia or acute kidney injury. However, diagnostic codes for these conditions are less than ideal. The sensitivity of the International Classification of Diseases (ICD)-9 and ICD-10 codes for hyponatraemia ranges from only 3 to 7%,9 10 which causes underestimation of the true event rates and absolute risk differences when comparing two or more drugs. However, this could be improved by linking hospital-based laboratory data to the other data sources to provide better estimates of risk.

The use of linked healthcare administrative databases to estimate the risk of an outcome of interest is straightforward when considering a well-defined region such as the province of Ontario—the numerator is the number of patients suffering the outcome and the denominator is the entire registered population. However, when only a portion of hospitals have linked laboratory data, defining the denominator (ie, those patients at risk for both developing the outcome and presenting to a particular hospital) becomes more challenging. The goal of this project was to assign the laboratory-linked hospitals in Southwestern Ontario the regions from which its patients receiving medications from outpatient pharmacies would reliably arise.

METHODS

Setting

We conducted this study using several linked health administrative databases in Ontario, Canada. Ontario is the most populous Canadian province, with approximately 13 million residents in the year 2010, of whom 1.8 million were older than 65 years.11 All residents received universal access to hospital and physician services, and elderly residents received coverage for prescription medications. Coverage for medical services and medications from a single provincial payer provided a comprehensive set of health administrative data. We completed the study according to a pre-specified protocol which was approved by the research ethics board at Sunnybrook Health Sciences Centre (Toronto, Ontario, Canada). The relevant datasets and the analyses were held and conducted at the Institute for Clinical Evaluative Sciences (ICES). The reporting of this study follows guidelines set out for observational studies (see online supplementary appendix 1).12

Overview

In Southwestern Ontario, we specified 118 geographical regions by postal FSAs and mapped a total of 45 hospitals with emergency departments, 12 of which had laboratory-linked data by the year 2009.13 14 The 33 non-laboratory-linked hospitals were selected based on their proximity to the laboratory-linked hospitals (see online supplementary appendix 2). For each adult aged 66 years or older who visited an emergency department at one of the 45 hospitals from 1 June, 2003 to 31 December, 2009, we identified the Southwestern Ontario FSA of the pharmacy that dispensed their most recent outpatient prescription in the prior 120 days. We then classified a region as eligible for inclusion in the catchment area if at least 90% of emergency room visits in that region were to laboratory-linked hospitals in a given year.

Data sources

We identified emergency department visits to one of the 45 hospitals using the National Ambulatory Care Reporting System database that is maintained by the Canadian Institutes of Health Information. We characterised each hospital by a unique ambulatory-care institution number. We identified prescription drug claims using the Ontario Drug Benefits (ODB) database. The ODB programme provides Ontario residents 65 years of age and older with coverage for prescription medications. To ascertain the specific regions to be included in the catchment area, we used the ODB database to obtain postal information related to the dispensing pharmacy. Specifically, this geographical information is known as the FSA. In Canada, postal geography for each province begins with an FSA, which represents the first three characters of a postal code.15 16 These geographical units were created by the Canada Post Corporation and assigned to regions in the province to facilitate the delivery of mail to businesses and households. Each character signifies important mailing information, including the postal district (first character), whether a particular region is urban or rural (second character), and specific areas within that region (third character).16 See online supplementary appendix 3 for graphical presentation of FSAs. We preferentially assigned the pharmacy FSA to a patient instead of their home FSA since postal address information for a patient may be outdated (eg, if a patient still uses an older health card or has moved). Cerner (Kansas City, Missouri, USA) is a system that keeps patient electronic medical records, including laboratory test results for participating hospitals in one repository.17 We recently linked a portion of the Cerner holdings for Southwestern Ontario hospitals with other healthcare datasets that are housed at the ICES in Ontario, Canada. This portion contained lab data including serum creatinine, potassium, sodium and glucose results for emergency department, inpatient and outpatient visits. Other information included the date and time of testing. The implementation of Cerner within Southwestern Ontario (sponsored by the provincial government) occurred in stages, with the number of hospitals in Ontario using the system increasing over time (see online supplementary appendix 4). In 2003, there were three laboratory-linked hospitals while in 2009, there were 12. We obtained patient demographic data from the Registered Persons Database, which contains demographic information on all Ontarians ever issued a health card. All these databases were reliably linked using a
unique identifier. We used census data (ie, linked to the FSA) to compare characteristics between our catchment area population and the Ontario population. Authorizations for use of this data were obtained from the London Health Sciences Centre (laboratory-linked hospital information) and ICES (all other datasets).

**Cohort selection**

Ontario residents 66 years of age and older were eligible for inclusion in the cohort. We excluded residents in their first year of eligibility of prescription drug coverage (age 65 years) to ensure at least one full year of available medication records. We also excluded patients who had a missing date of birth or missing sex. We enrolled patients into the cohort based on a two-step process for each year in the period of interest. First, we identified patients by their first unplanned visit to an emergency department for any cause at 1 of the 45 Southwestern Ontario hospitals. We denoted the date of this visit as the index date. We then looked back 120 days prior to the index date for the most recent outpatient prescription claim for any drug and identified the location of the dispensing pharmacy by the FSA. We excluded a small number of patients who had two or more prescriptions at two or more pharmacies on the day of the most recent prescription. If the identified FSA was not part of the Southwestern Ontario region (ie, not 1 of the 118 prespecified FSAs), we excluded these patients.

**FSA selection**

We calculated the proportion of patients who presented to laboratory-linked hospitals versus non-laboratory-linked hospitals in a given pharmacy FSA to determine the eligibility of FSAs. Because the number of laboratory-linked hospitals increased from year to year, we repeated the calculations for each year of interest. For an FSA to be included in the catchment area, at least 90% of emergency department visits in that FSA had to occur at laboratory-linked hospitals in a given year. FSAs in which more than 10% of emergency visits were to non-laboratory-linked hospitals were not included in the catchment area as we could not be confident that a medication-related lab-based disorder would prompt presentation to a laboratory-linked hospital. (Note: privacy regulations prohibit us from specifying cell sizes less than six; when this occurred, we treated these cell sizes as zero in all calculations.)

**RESULTS**

A flow diagram illustrating the procedures for catchment area ascertainment is presented in figure 1. From 2003 to 2009, there were a total of 649,713 emergency department visits with a most recent prescription claim in a particular FSA, of which 141,302 (22%) were to laboratory-linked hospitals. In the year 2003, there were 12 FSAs that met our criteria and this grew to 25 FSAs by the year 2009 (see online supplementary appendix 5). The map of Southwestern Ontario in figure 2 depicts the locations of these regions. Individuals from our catchment area (year 2009) were similar to the rest of the elderly population of Ontario (table 1). As of the year 2006 (most recent census profile), almost 5% of Ontario’s elderly population (80,000 adults ≥65 years of age) resided in these 25 FSAs.

**DISCUSSION**

Determining the geographical regions that are predominantly serviced by laboratory-linked hospitals is an important first step to using hospital-based laboratory data in health outcomes research. Ours is the first study to use this methodology within Ontario’s linked health administrative databases to determine a catchment area for particular hospitals. Where there is complete connectivity of data sources (eg, electronic medical record data, provincial hospital and procedure data, drug claim information, etc), other researchers may replicate these methods to define relevant regions for their jurisdiction of interest.

Since the year 2009, the number of hospitals with laboratory-linked data has remained consistent. As such, the catchment area has been kept the same since then. Certainly, the set of eligible FSAs may fluctuate depending on resident characteristics and available healthcare facilities. However, based on the patterns we observed from 2003 to 2009, the number of eligible FSAs only increased when additional hospitals began using the Cerner system. As such, once other hospitals have registered with Cerner, we will use the same methodology to update the existing catchment area. The study also highlights how regions of interest defined by geography can change over time.

Our study had several strengths that helped mitigate sources of error. We used emergency department visits as opposed to inpatient hospital admissions to ascertain encounters at a specific hospital. Inpatient hospital admissions can be planned and, particularly for tertiary care centres with specialised services, may include patients who live both near to and far from the hospital. Conversely, the emergency department setting allowed us to detect unplanned visits from patients within the area and likely reflected the true population who would present to the hospital if they were to incur a serious sudden medication-related laboratory disorder. We had a large number of emergency department visits from all over Southwestern Ontario to form the basis of our catchment area.

Some studies might restrict their analysis to only laboratory-linked hospitals and consider those FSAs where a large number of patients visited these hospitals. However, this would fail to ascertain the number of patients presenting to nearby non-laboratory-linked hospitals. For this reason, in our analysis we considered both laboratory and non-laboratory-linked hospitals to limit the possibility of falsely classifying a given FSA as part of the catchment area.
ODB is a highly reliable database for prescription drug claims with a basic error rate under 1% (\(\sim 0.7\%), 95\% CI 0.5\% to 0.9\%\). This indicates that previous prescription use was identified with a high degree of accuracy. Also, we ensured that temporality was established in that the prescription claim predated the emergency department visit. These methods were apt to detect an outpatient medication that was filled from a pharmacy in the patient’s home region.

Our study does have some limitations. Although the ODB database was highly accurate for prescription drug claims, we did not have an indication about the validity of the pharmacy FSAs. If some pharmacy FSAs were incorrect, we might have misclassified them as being either eligible or ineligible for inclusion in the catchment area. If this was the case, we would not be capturing the true regions from which patients would present to a laboratory-linked hospital. However, this was likely not an issue since the FSAs identified for the corresponding hospitals had excellent face validity. For example, patients who visited Tillsonburg District Memorial Hospital filled a prescription, mainly, from a Tillsonburg FSA.

Since the provincial drug plan only contains drug-dispensing information on patients over the age of 65, the current methods preclude us from capturing adverse events in younger patients. However, future drug safety studies that use this catchment area will still address adverse health outcomes in an understudied, vulnerable segment of the population.

We recognise that we will not capture cases who do not present to hospital (ie, those who do not present at all or who present to an outpatient laboratory instead), or those who present to hospital but fail to have the appropriate tests. Nonetheless, we will capture a substantial number of important severe lab-based outcomes that if anything, will underestimate the true incidence. These
Figure 2  Locations of eligible regions (shaded) in Southwestern Ontario as of 2009. Boxed numbers represent laboratory-linked hospitals and starred numbers represent non-laboratory-linked hospitals. See online supplementary appendix 2 legend for further details.

Table 1  Comparison of catchment area and Ontario populations*

<table>
<thead>
<tr>
<th>FSA characteristics</th>
<th>Catchment area</th>
<th>Ontario</th>
</tr>
</thead>
<tbody>
<tr>
<td>N</td>
<td>25</td>
<td>509</td>
</tr>
<tr>
<td>Rural, N (%)</td>
<td>2 (8.0)</td>
<td>55 (10.8)</td>
</tr>
<tr>
<td>Patient characteristics</td>
<td></td>
<td></td>
</tr>
<tr>
<td>N</td>
<td>80000</td>
<td>1649055</td>
</tr>
<tr>
<td>Age, N (%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>65–69</td>
<td>21455 (26.8)</td>
<td>466295 (28.3)</td>
</tr>
<tr>
<td>70–74</td>
<td>18760 (23.5)</td>
<td>401890 (24.4)</td>
</tr>
<tr>
<td>75–79</td>
<td>16480 (20.6)</td>
<td>338825 (20.5)</td>
</tr>
<tr>
<td>80–84</td>
<td>13085 (16.4)</td>
<td>250250 (15.2)</td>
</tr>
<tr>
<td>85+</td>
<td>10220 (12.8)</td>
<td>191795 (11.6)</td>
</tr>
<tr>
<td>Women, N (%)</td>
<td>46150 (57.7)</td>
<td>931580 (56.5)</td>
</tr>
</tbody>
</table>

*2006 Census counts were applied to the 2009 catchment area.
studies will allow us to generate new information to guide optimal medication use.

We also understand that because our strict inclusion criteria has defined a population representing only 5% of Ontario’s elderly residents, findings that arise from this catchment area may not fully generalise to the entire province. With respect to age, sex and rural living, the catchment area was similar in make-up to the Ontario population. Nonetheless, studies using this area will have lower sample sizes as compared with the entire province. Depending on the number of medication users and expected event rate, this may challenge the feasibility of some studies. For this reason, it may be prudent to conduct separate analyses examining diagnostic codes first (for all of Ontario) and then lab results (for the catchment area). Observing concordant signals across these two sets of analyses would strengthen inferences about the associations under study.

Implementation

We are now using this catchment area to define hospital-based laboratory outcomes in Canadian Institutes of Health Research-funded drug safety studies. Outcomes of interest include hospitalisation with hyponatremia, hyperkalemia and acute kidney injury. For research that examines lab-based disorders from medication use, both diagnostic codes and lab data are being used to define a particular outcome in separate related analyses. Preliminary work confirms that approximately 5% of elderly medication users in Ontario are being assessed in our catchment area.

CONCLUSION

Medication-related lab-based outcomes can be accurately identified at the population level using hospital-based laboratory data. A catchment area for those regions serviced by laboratory-linked hospitals can be used in future analyses. By capturing serious events that would otherwise go undetected by diagnostic codes, researchers may better inform health policy decision-makers about potential risks of common medications in routine practice. The new knowledge created can be translated and integrated into clinical practice (eg, routine measurements of serum electrolytes after certain drugs are prescribed) so that adverse events can be mitigated or even avoided.

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Contributors SG participated in the data acquisition and linkage, coordination of the study, study design, provided interpretation of study results, and drafted the manuscript. SZS participated in the study design, performed the analysis and provided interpretation of study results. MMH, TH and GK participated in the data acquisition and linkage. MAW participated in the study design and provided feedback on the manuscript. AXG conceived of the study, participated in its design and provided feedback on the manuscript. All authors read and approved the final manuscript.

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REFERENCES


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