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1 **Improving Epilepsy Care in Ontario, Canada: the Impact of a Provincial Strategy for Epilepsy** 2 **Care**

3 Tresah C. Antaya, MPH;¹ Brooke Carter, MSc;² Salimah Z. Shariff, PhD;² Lysa Boissé
4 Lomax, MD;³ Elizabeth Donner, MD;^{4,5} Kirk Nylen, PhD;⁵ O Carter Snead III, MD;^{4,5} Jorge
5 G. Burneo, MD^{1,2*}

6 ¹Epilepsy Program and Neuroepidemiology Research Unit, Western University, London,
7 Ontario, Canada

8 ²ICES Western, London Health Sciences Research Institute, London, Ontario, Canada

9 ³Queen's University, Kingston, Ontario, Canada

10 ⁴Hospital for Sick Children, Toronto, Ontario, Canada

11 ⁵University of Toronto, Toronto, Ontario, Canada

12 ***Corresponding author:** Email addresses: TCA: tantaya2@uwo.ca; BC:

13 brooke.carter@ices.on.ca; SZS: Salimah.shariff@ices.on.ca; LBL:

14 lysa.boisselomax@kingstonhsc.ca; ED: elizabeth.donner@sickkids.ca; KN:

15 kirk@baszuckigroup.org; OCS: carter.snead@sickkids.ca; JGB: jburneo2@uwo.ca

16 **ORCID**

17 TCA: <https://orcid.org/0000-0002-8612-5976>

18 JGB: <https://orcid.org/0000-0002-3644-2826>

19 **Highlights**

- 20 • We assessed whether the Provincial Strategy was associated with changes in epilepsy
21 surgery and presurgical assessment rates.
- 22 • The Provincial Strategy was associated with a level increase in the rate of epilepsy
23 surgery and slope decreases in the rates of other healthcare use for epilepsy.
- 24 • Other regions with low rates of epilepsy surgery may benefit from similar interventions.

25 **Abstract**

26 **Objective**

27 In 2016, the Ontario Ministry of Health and Long-Term Care implemented the Provincial
28 Strategy for Epilepsy Care to increase epilepsy surgery use in Ontario, Canada. The
29 objectives of this study were to assess whether the use of (1) epilepsy surgery, including (a)
30 its receipt and (b) assessments for candidacy, and (2) other healthcare for epilepsy, including
31 (a) neurological consultations, (b) emergency department (ED) visits, and (c) hospital
32 admissions, changed since its implementation.

33 **Methods**

34 We used linked health administrative data and an interrupted time series design. Annual
35 cohorts were created for July 1st to June 30th of each year from 2007 to 2019, comprising
36 patients with drug-resistant epilepsy eligible for publicly-funded prescription drug coverage
37 with no cancer history. We used segmented Poisson regression models to assess whether the
38 annual rates of each outcome changed between the period before the Provincial Strategy was
39 implemented (July 2007 to June 2016) and the period after.

40 **Results**

41 There was a level increase in the rate of epilepsy surgery of 48% (95% CI: 0%, 118%) and
42 slope decreases in the rates of neurological consultations, ED visits, and hospital admissions
43 for epilepsy of 10% (95% CI: -15%, -5%), 10% (95% CI: -20%, 1%), and 7% (95% CI: -
44 12%, -1%) per year, respectively, associated with the Provincial Strategy.

45 **Conclusion**

46 The Provincial Strategy may be associated with an increased rate of epilepsy surgery and
47 reduced rates of other healthcare use for epilepsy. Other regions experiencing low epilepsy
48 surgery rates may benefit from similar interventions.

49 **Keywords:** health services; presurgical evaluation; epidemiology; pharmaco-resistant;
50 refractory

51 **Introduction**

52 Epilepsy is one of the most common neurological disorders,(1) and significantly affects
53 quality of life(2) and increases morbidity and mortality risk.(3,4) Although epilepsy is a
54 complex disease requiring a holistic approach to treatment,(2) seizure frequency and severity
55 are major determinants of health-related quality of life in patients with epilepsy.(5,6)
56 Therefore, seizure control is an important objective of epilepsy treatment. However, a third of
57 patients with epilepsy continue to experience seizures despite appropriate treatment with
58 antiseizure medications (ASMs)(7) and are diagnosed with drug-resistant epilepsy (DRE).(8)
59 Fortunately, epilepsy surgery is an effective treatment alternative for patients with DRE who
60 are eligible.(9) Epilepsy surgery typically involves the resection of the epileptogenic zone,
61 the cortical area in which the patient's seizures originate.(10) It is recommended that patients
62 with DRE are assessed for epilepsy surgery eligibility upon diagnosis,(11–14) which is
63 determined by the identifiability of the epileptogenic zone and the risk of post-operative
64 morbidity.(10) However, epilepsy surgery is underutilized in the province of Ontario, where
65 approximately 39% (15.1 million) of Canadians reside.(15) Between 2001 and 2010, just
66 1.2% of patients received epilepsy surgery within two years of DRE diagnosis in Ontario, and
67 only 2.2% were assessed for candidacy.(16) These figures indicate that many people with
68 epilepsy in our region who may have benefited were not being considered for the
69 procedure.(9)

70 In 2013, the Ontario Ministry of Health and Long-Term Care (OMHLTC) approved the
71 Provincial Strategy for Epilepsy Care, herein referred to as the Provincial Strategy, to
72 increase epilepsy surgery rates. The OMHLTC committed to adding 21 new Epilepsy
73 Monitoring Unit (EMU) beds across the province, allocated funding specifically for epilepsy
74 surgeries (including neurostimulation), and commissioned the Epilepsy Implementation Task
75 Force (EITF) to implement the remaining components of the Provincial Strategy.(17) The
76 EITF, comprising senior leaders of the epilepsy community, was tasked with coordinating
77 resources and waitlists, establishing standardized protocols across hospitals for epilepsy
78 diagnosis and surgery, and developing resources for community neurologists and primary
79 care providers.(17) These standardized protocols and resources were developed as guidelines,
80 accessible from the Ontario Epilepsy Guidelines website.(18)

81 For logistical reasons, only 19 of the 21 EMU beds were added, for a total of 40 (28 adult and
82 12 pediatric) in the province. Eleven beds were distributed between three cities that did not

83 previously have EMUs. These facilities, now called District Epilepsy Centres (DECs), cannot
84 perform epilepsy surgeries but can assess patients for eligibility. The other eight new EMU
85 beds were distributed between the two cities with existing EMUs. These facilities can
86 perform epilepsy surgeries and are called Regional Epilepsy Surgery Centres of Excellence
87 (RESCs). Fig. 1 depicts the current locations of DECs and RESCs in Ontario.(19,20)

88 The effectiveness of the Provincial Strategy has not yet been evaluated using a population-
89 based sample of patients with DRE. Therefore, our primary objective was to assess whether
90 the Provincial Strategy was associated with changes in the rates of epilepsy surgery and
91 assessments for candidacy. Our secondary objective was to assess whether the rates of other
92 healthcare for epilepsy, including neurological consultations, emergency department (ED)
93 visits, and hospital admissions, changed between the period before the Provincial Strategy
94 was implemented and the period after.

95 **Materials and methods**

96 **Study Design and Data Sources**

97 We conducted a population-based study with an interrupted time series design using data
98 routinely collected on the health services provided within Ontario's publicly-funded
99 healthcare system. The databases used include the Canadian Institute for Health Information
100 (CIHI) Discharge Abstract Database (DAD), CIHI Same-Day Surgery (SDS), CIHI National
101 Ambulatory Care Reporting System (NACRS), Ontario Health Insurance Plan (OHIP),
102 Ontario Drug Benefit Program (ODB), Registered Persons Database (RPDB), Drug
103 Identification Number (DIN), Local Health Integration Network (LHIN), Postal Code
104 Conversion File (PCCF), and Ontario Cancer Registry (OCR).

105 These datasets were linked using unique encoded identifiers and analyzed at ICES. ICES is
106 an independent, non-profit research institute whose legal status under Ontario's health
107 information privacy law allows it to collect and analyze health care and demographic data,
108 without consent, for health system evaluation and improvement. Brief descriptions of the
109 databases used in this study are available in Supplementary Table 1.

110 **Study Population**

111 We assembled annual cohorts for July 1st to June 30th of each year from 2007 to 2019,
112 comprising all individuals with prevalent DRE who were eligible for publicly-funded

113 prescription drug coverage under the ODB program and had no history of cancer (Fig. 2).
114 The ODB program covers most of the cost of prescription medications for eligible individuals
115 in Ontario.(21) Eligible individuals include those who are 65 or older, live in long-term or
116 special care homes, receive home or community care, receive disability or income support, or
117 have high prescription drug costs relative to their income.(21) In 2018, individuals 24 years
118 and younger also became eligible, which was changed in 2019 to exclude those with private
119 prescription drug coverage.(22)

120 Epilepsy was defined using a validated algorithm of ICD-10 and OHIP physician billing
121 codes.(23) DRE was identified among patients with epilepsy as those who were prescribed at
122 least two antiseizure medications (ASM), each with at least 90 days duration, followed by a
123 third ASM or a seizure. The study population was restricted to those eligible for the ODB
124 program because drug prescription data was needed to apply our definition of drug-resistant
125 epilepsy and is only available for these individuals. The specific ICD and OHIP codes used to
126 define the inclusion and exclusion criteria are available in Supplementary Table 2, and the
127 eligible ASMs used to identify those with DRE are listed in Supplementary Table 3.

128 **Outcome Definitions**

129 Epilepsy surgery was identified using physician billing codes in the OHIP database. Eligible
130 surgical procedures included resections of any size, commissurotomies, callosotomies,
131 hemispherectomies, and implantations of deep brain and vagus nerve stimulators. We did not
132 include responsive neurostimulation, laser interstitial thermal therapy, or other ablation
133 techniques in this definition as they were unavailable in Canada before 2020. Stimulator
134 implantation was included in this definition since it indicates the patient received appropriate
135 care. Epilepsy surgery candidacy assessment was also identified using physician billing codes
136 in the OHIP database. We considered a patient to have been assessed for surgical candidacy if
137 they received scalp video-electroencephalography (video-EEG) or were implanted with
138 intracranial (subdural or depth) electrodes, whichever occurred first.

139 Neurological consultations for epilepsy were identified using the OHIP database, where we
140 required one physician billing code for an outpatient neurological consultation billed with
141 epilepsy as the reason. We used ICD-10 codes for epilepsy, status epilepticus, and Landau-
142 Kleffner syndrome in the NACRS database to identify ED visits and in the DAD to identify
143 hospital admissions for epilepsy. Specific epilepsy syndromes other than Landau-Kleffner
144 could not be included in this definition as ICD-10 codes for these syndromes are unavailable

145 in the databases used. The specific OHIP and ICD codes used to define all outcomes are
146 available in Supplementary Table 2.

147 **Research Ethics Statement**

148 ICES is a prescribed entity under Ontario's Personal Health Information Protection Act
149 (PHIPA). Section 45 of PHIPA authorizes ICES to collect personal health information,
150 without consent, for the purpose of analysis or compiling statistical information with respect
151 to the management of, evaluation or monitoring of, the allocation of resources to or planning
152 for all or part of the health system. Projects that use data collected by ICES under section 45
153 of PHIPA, and use no other data, are exempt from REB review. The use of the data in this
154 project is authorized under section 45 and approved by ICES' Privacy and Legal Office.

155 **Statistical Analyses**

156 We used segmented Poisson regression models to assess whether the annual incidence of
157 each outcome changed between the period before the Provincial Strategy was initiated (July
158 2007 to June 2016) and the period after (July 2016 to June 2019). Segmented regression is a
159 method of analyzing interrupted time series data where separate estimates are obtained for the
160 model intercept and slope in each period pre- and post-intervention.(24) The intervention is
161 assessed using the difference between the actual post-intervention intercept (the level change)
162 and slope (the slope change) from those expected if the pre-intervention trend had continued
163 unchanged into the post-intervention period.(25) We selected June 30th, 2016, as the
164 interruption point as this was the approximate date that all EMU beds became available. We
165 expected that adding EMU beds would have the most significant effect of all the strategy
166 components since this would directly increase institutional capacity to assess patients for
167 surgical candidacy. We checked the assumptions of our segmented regression models (i.e.
168 overdispersion, autocorrelation, and non-stationarity) to ensure they were satisfied. A p-value
169 of 0.05 was used to determine statistical significance, and all analyses were conducted by BC
170 using SAS software version 9.4 (SAS Institute Inc., Cary, NC, USA).

171 **Data availability**

172 The dataset from this study is held securely in coded form at ICES. While legal data sharing
173 agreements between ICES and data providers (e.g., healthcare organizations and government)
174 prohibit ICES from making the dataset publicly available, access may be granted to those
175 who meet pre-specified criteria for confidential access, available at www.ices.on.ca/DAS

176 (email: das@ices.on.ca). The full dataset creation plan and underlying analytic code are
177 available from the authors upon request, understanding that the computer programs may rely
178 upon coding templates or macros that are unique to ICES and are therefore either inaccessible
179 or may require modification.

180 **Results**

181 We included 79,462 participants with DRE in the period before the Provincial Strategy was
182 implemented and 34,686 in the period after. The total and annual mean number of each
183 outcome identified by period are available in Table 1. The results of the segmented Poisson
184 regression analyses are summarized in the following sections, with all specific estimates
185 available in Supplementary Table 4 and the plotted rates and fitted segmented regression
186 lines in Fig. 3.

187 **Epilepsy Surgeries and Assessments for Candidacy**

188 The Provincial Strategy was associated with a level increase in the rate of epilepsy surgery of
189 48% (95% CI: 0%, 118%). The slope change in this rate was minimal, decreasing by 2%
190 (95% CI: -17%, 15%) per year. The Provincial Strategy was not associated with level or
191 slope changes in the rate of assessments for epilepsy surgery candidacy, with a level increase
192 of 5% (95% CI: -18%, 12%) and a slope decrease of 4% (95% CI: -10%, 3%) per year.

193 **Other Healthcare for Epilepsy**

194 The Provincial Strategy was not associated with level changes in the rates of neurological
195 consultations (8%, 95% CI: -6%, 23%), emergency department visits (-5%, 95% CI: -28%,
196 25%), or hospital admissions for epilepsy (10%, 95% CI: -4%, 25%). However, the
197 Provincial Strategy was associated with slope decreases in the rates of these outcomes of 10%
198 (95% CI: -15%, -5%), 10% (95% CI: -20%, 1%), and 7% (95% CI: -12%, -1%) per year,
199 respectively.

200 **Discussion**

201 We found that the Provincial Strategy was associated with a 48% level increase in the rate of
202 epilepsy surgeries. We also found that the Provincial Strategy was associated with slope
203 decreases in the rates of neurological consultations, emergency department visits and hospital

204 admissions for epilepsy of 10%, 10%, and 7% per year, respectively. However, the estimate
205 for emergency department visits was not statistically significant.

206 These findings indicate that the Provincial Strategy may be associated with an increase in the
207 use of epilepsy surgery in Ontario. The estimate for the level increase in the rate of surgeries
208 was not statistically significant; however, the lower confidence limit was equal to the null
209 value of zero. Both the number of events per data point and the number of data points in the
210 periods before and after the interruption point affect power in segmented regression
211 analyses.⁽²⁶⁾ Considering that there was an annual average number of epilepsy surgeries of
212 49.2, there were only three data points post-interruption and that the lower confidence limit
213 was equal to the null value, we likely had insufficient power to reject the null hypothesis.
214 Therefore, we believe the observed increase likely reflects a true increase in this outcome.
215 This finding is important not only for patients, but also for Ontario's healthcare system. The
216 Provincial Strategy was shown to be cost-effective prior to implementation,⁽²⁷⁾ and the cost-
217 effectiveness of epilepsy surgery in Ontario in general has also been demonstrated.^(28–30)

218 It was surprising that we observed no corresponding increase in assessments for surgical
219 candidacy. The long waitlists for epilepsy surgery preceding the availability of additional
220 funding, and a corresponding increase in operating room resources for epilepsy surgery, may
221 have caused the sudden increase in epilepsy surgeries but not assessments. Additionally,
222 EMU beds in some DEC and RESCs are used to treat conditions other than epilepsy if
223 needed, which may have contributed to the lack of increase in assessments.

224 A previous study found a slope decrease in epilepsy surgeries at the Toronto RESC among
225 pediatric patients between 2013 and 2019, relative to 2001 to 2012.⁽³¹⁾ This study also found
226 that in both periods, there was an increasing trend in assessments for epilepsy surgery
227 candidacy. These findings contrast those of the present study; however, these studies have
228 several methodological differences. The previous study selected 2013 as the interruption
229 point instead of 2016, as it was consistent with a previous study by the authors and with the
230 timing of the Provincial Strategy's approval by OMHLTC. We selected 2016 as it was
231 consistent with the approximate date that the added EMU beds at the RESCs became
232 available. Additionally, the previous study included patients at just one specialized epilepsy
233 centre in Ontario, which may not reflect the trends observed in the entire province.

234 Although the relative increase in epilepsy surgeries was substantial, the absolute rate of
235 epilepsy surgery remains low after the Provincial Strategy was implemented. This finding

236 suggests that most patients with DRE in Ontario are still not receiving appropriate epilepsy
237 care. Considering that there remain waitlists for EMU admission, intracranial-EEG, and
238 epilepsy surgery, increasing the number of EMU beds and the availability of neurosurgeons
239 and surgical facilities is likely needed to further increase the Ontario healthcare system's
240 capacity to perform epilepsy surgeries.

241 We also found slope decreases in the rates of neurological consultations, ED visits, and
242 hospital admissions for epilepsy after the Provincial Strategy was implemented. Although the
243 decrease in ED visits was not statistically significant, the upper confidence limit was close to
244 the null value. Considering that the estimate is not very precise, its lack of statistical
245 significance may also be due to insufficient power.

246 The slope decrease in neurological consultations can be interpreted as a positive or negative
247 outcome of the Provincial Strategy, depending on whether it is associated with improved
248 seizure control. If the decrease reflects fewer neurological consultations not associated with
249 improved seizure control, it indicates that patients received less frequent care from a
250 neurologist without a significant improvement in their symptoms. However, we have
251 evidence that epilepsy surgery rates may have increased and that ED visits and hospital
252 admissions for epilepsy may have decreased, both suggesting that seizure control likely
253 improved in this population. Therefore, considering these findings and that improved seizure
254 control is typically associated with less frequent neurologist visits, we believe this decrease is
255 a positive outcome of the Provincial Strategy.

256 There are some limitations of this research. One limitation is our likely lack of sufficient
257 power in the analyses estimating the associations of the Provincial Strategy with epilepsy
258 surgeries and possibly with ED visits. We could not increase power in this study, as we
259 included all individuals who met our inclusion criteria and resided in Ontario. We also could
260 not include the annual periods after 2019 because epilepsy surgeries were affected by the
261 COVID-19 pandemic, which began in March 2020 in our region.

262 The identification of our study sample also has some potential limitations. Our definition of
263 drug-resistant epilepsy may not have been sufficiently specific, as we could not determine
264 whether patients tried a second antiseizure medication due to intolerability. However, we
265 attempted to mitigate this limitation by including a minimum duration of 90 days for each
266 antiseizure medication trial, as we expected medication changes due to intolerability to occur
267 within the first 90 days. Another limitation is our ability to include only individuals eligible

268 for ODB coverage. Therefore, we could not include all patients with DRE in the province.
269 However, since those who receive disability benefits are eligible for ODB and many patients
270 with DRE receive disability benefits, we likely included a significant proportion of patients
271 with DRE in the province. Additionally, the ODB eligibility criteria changed during the study
272 period. In 2018, the Ontario government extended eligibility to individuals 24 years and
273 younger, and in 2019, to those 24 years and younger without private prescription drug
274 coverage.(22) Therefore, this extension of eligibility may have introduced selection bias if
275 young people who would not have been covered before this change had systematically
276 differing rates of our outcomes than the rest of the study population. Importantly, these
277 limitations indicate that estimates of the prevalence of drug-resistant epilepsy in Ontario
278 cannot be derived from this study.

279 Misclassification of the outcomes also likely occurred; however, the magnitude is unclear, as
280 the codes and algorithms used to identify these concepts were not previously validated.
281 Additionally, we could only use video-EEG to identify surgical assessments, as
282 neuropsychological evaluations are unidentifiable in the available data. In addition, other
283 changes in epilepsy care not related to the Provincial Strategy likely occurred in the province
284 during the study period, which may have confounded our results. To our knowledge, the only
285 significant change was the development of Project ECHO: Epilepsy Across the Lifespan, a
286 continuing medical education program for community healthcare professionals to improve
287 epilepsy care.(32) The virtual and case-based program connects community healthcare
288 providers with a multi-disciplinary team experienced in diagnosing and treating epilepsy. As
289 our study period ended in June 2019, and Project ECHO: Epilepsy was launched in 2019, it is
290 unlikely that the program confounded our findings.

291 The choice of interruption point may be a limitation of this study. We selected July 1st, 2016,
292 as this is the approximate date that all new EMU beds were added at the RESCs. However,
293 the guidelines and standardized protocols were published between 2014 and 2017, and the
294 coordination of resources and waitlists began in 2017. Although the funding for epilepsy
295 surgery became available in 2013, it is unlikely that it was immediately utilized, as
296 corresponding increases in personnel and surgical facilities availability were required.
297 Finally, there may be spatial heterogeneity in the effectiveness of the Provincial Strategy due
298 to the distribution of the added EMU beds and other resources. Future research should
299 explore whether there may be spatial accessibility barriers to epilepsy care in Ontario.

300 In conclusion, the Provincial Strategy was associated with a level increase in the rate of
301 epilepsy surgeries and slope decreases in the rates of neurological consultations, ED visits,
302 and hospital admissions for epilepsy. However, the estimates for epilepsy surgeries and ED
303 visits were not statistically significant. Other regions with comparable healthcare
304 infrastructure experiencing low rates of epilepsy surgery may benefit from similar
305 interventions.

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317 **Competing interests**

318 J Burneo received funding for this study from the Jack Cowin Endowed Chair in Epilepsy
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324 **Statement of Authorship**

325 T Antaya: project administration, methodology, writing – original draft preparation. B Carter:
326 project administration, methodology, data curation, formal analysis, writing – review &
327 editing. S Shariff: methodology, writing – review & editing. L Boissé Lomax:
328 conceptualization, writing – review & editing. E Donner: conceptualization, writing – review
329 & editing. K Nylen: conceptualization, writing – review & editing. OC Snead:

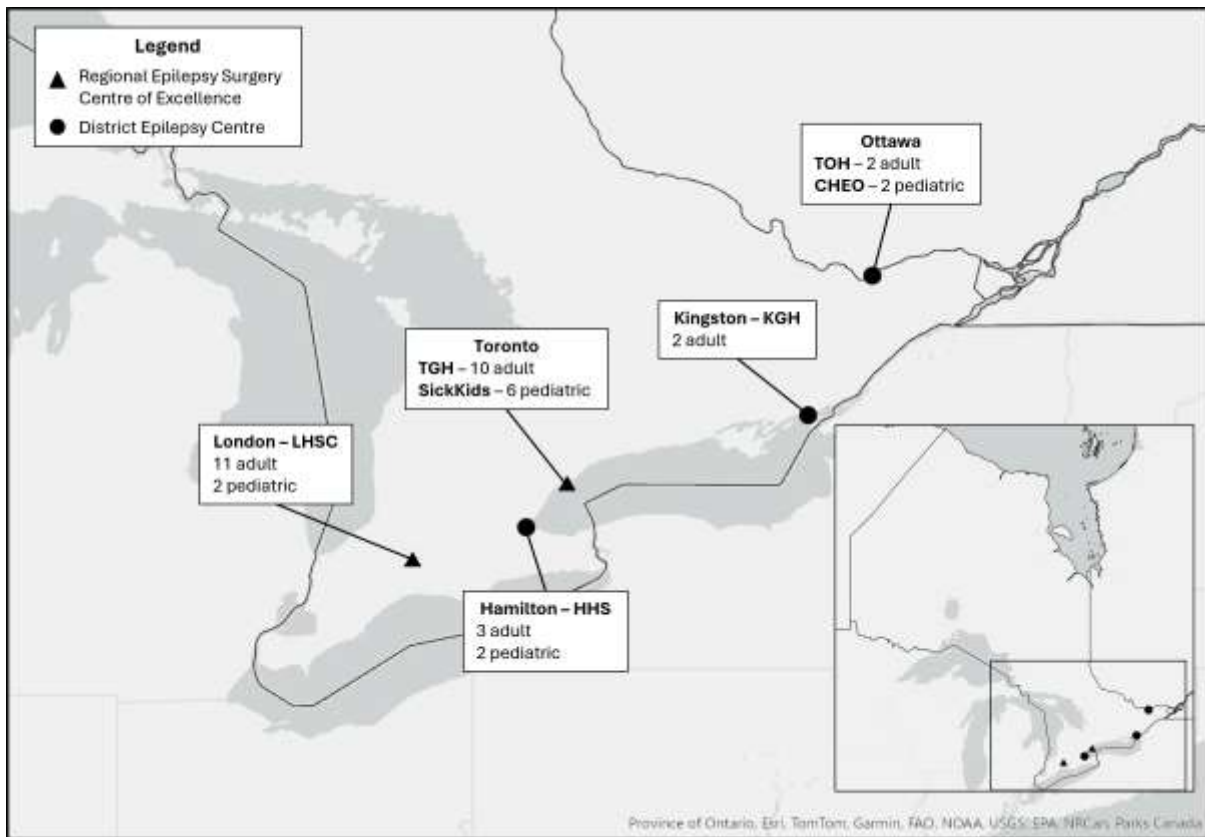
330 conceptualization, writing – review & editing. J Burneo: conceptualization, methodology,
331 funding acquisition, supervision, writing – review & editing.

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427

428 **Figure 1 Current Epilepsy Monitoring Unit locations in Ontario, Canada.**

429 Footnote: LHSC: London Health Sciences Centre; TGH: Toronto General Hospital; HHS:
 430 Hamilton Health Sciences; KGH: Kingston General Hospital; TOH: The Ottawa Hospital;
 431 CHEO: Children's Hospital of Eastern Ontario

Met the diagnosis of epilepsy within 10 years before beginning of study year (e.g. 10 years before July 1, 2009 [index date])

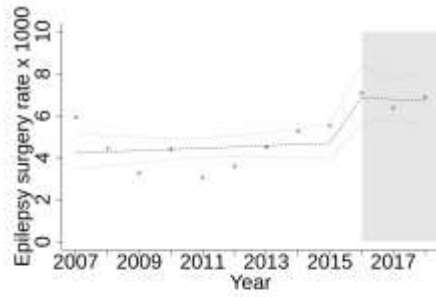
Exclusions:

- a) Missing age or sex
- b) Age >105 years on index date
- c) Death before index date
- d) Non-Ontario resident on index date
- e) Ineligible for OHIP between index date and end of study year (e.g. June 30, 2010)
- f) Death before end of study year
- g) Had cancer prior to index date
- h) Not eligible for ODB between epilepsy diagnosis date and index date
- i) Did not meet definition of drug-resistant epilepsy between epilepsy diagnosis date and index date

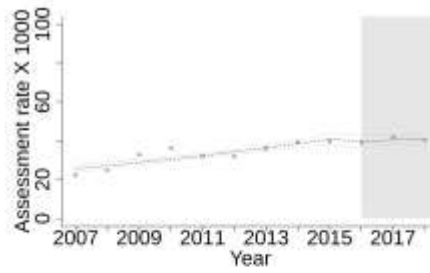
Participants included in annual cohort

432

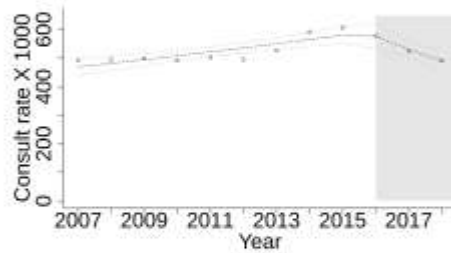
433 **Figure 2 Flow chart depicting annual cohort build.**



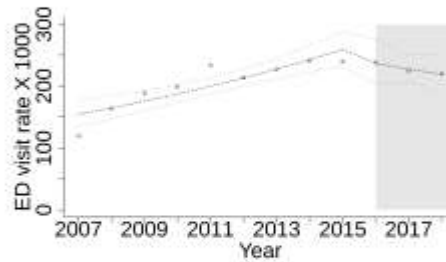
a) Epilepsy surgery.



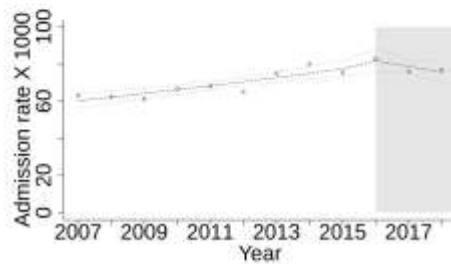
b) Assessment for epilepsy surgery candidacy.



c) Neurological consultations for epilepsy.



d) Emergency department visits for epilepsy.



e) Hospital admissions for epilepsy.

434

435 **Figure 3 Segmented regression plot for each outcome.** (A) Epilepsy surgery. (B)
 436 Assessment for epilepsy surgery candidacy. (C) Neurological consultations for epilepsy. (D)
 437 Emergency department visits for epilepsy. (E) Hospital admissions for epilepsy.