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## ORIGINAL ARTICLE

# Global prevalence of cerebral palsy: A systematic analysis

Sarah McIntyre<sup>1</sup>  | Shona Goldsmith<sup>1</sup>  | Annabel Webb<sup>1</sup>  | Virginie Ehlinger<sup>2</sup>  |  
 Sandra Julsen Hollung<sup>3</sup>  | Karen McConnell<sup>4</sup>  | Catherine Arnaud<sup>5</sup>  |  
 Hayley Smithers-Sheedy<sup>1</sup>  | Maryam Oskoui<sup>6</sup>  | Gulam Khandaker<sup>7</sup>  |  
 Kate Himmelmann<sup>8</sup>  | on behalf of the Global CP Prevalence Group\*

<sup>1</sup>Cerebral Palsy Alliance Research Institute, Specialty of Child and Adolescent Health, Sydney Medical School, Faculty of Medicine and Health, The University of Sydney, Sydney, Australia

<sup>2</sup>Center for Epidemiology and Research in Population health (CERPOP), Inserm, University of Toulouse, Toulouse, France

<sup>3</sup>Norwegian Quality and Surveillance Registry for Cerebral Palsy (NorCP), Vestfold Hospital Trust, Tønsberg, Norway

<sup>4</sup>School of Health Sciences, Northern Ireland, UK

<sup>5</sup>Public Health Department, University of Toulouse, Toulouse, France

<sup>6</sup>Department of Pediatrics, Faculty of Medicine and Health Sciences, McGill University, Montreal, Canada

<sup>7</sup>Central Queensland Hospital and Health Service, Rockhampton, Australia

<sup>8</sup>Department of Pediatrics, Institute of Clinical Sciences, Sahlgrenska Academy, University of Gothenburg, Gothenburg, Sweden

## Correspondence

Sarah McIntyre, Cerebral Palsy Alliance Research Institute, University of Sydney, Level 4 Building M02C, 88 Mallett Street, Camperdown, NSW 2050, Australia.  
 Email: [smcintyre@cerebralpalsy.org.au](mailto:smcintyre@cerebralpalsy.org.au)

## Abstract

**Aim:** To determine trends and current estimates in regional and global prevalence of cerebral palsy (CP).

**Method:** A systematic analysis of data from participating CP registers/surveillance systems and population-based prevalence studies (from birth year 1995) was performed. Quality and risk of bias were assessed for both data sources. Analyses were conducted for pre-/perinatal, postnatal, neonatal, and overall CP. For each region, trends were statistically classified as increasing, decreasing, heterogeneous, or no change, and most recent prevalence estimates with 95% confidence intervals (CI) were calculated. Meta-analyses were conducted to determine current birth prevalence estimates (from birth year 2010).

**Results:** Forty-one regions from 27 countries across five continents were represented. Pre-/perinatal birth prevalence declined significantly across Europe and Australia (11 out of 14 regions), with no change in postneonatal CP. From the limited but increasing data available from regions in low- and middle-income countries (LMICs), birth prevalence for pre-/perinatal CP was as high as 3.4 per 1000 (95% CI 3.0–3.9) live births. Following meta-analyses, birth prevalence for pre-/perinatal CP in regions from high-income countries (HICs) was 1.5 per 1000 (95% CI 1.4–1.6) live births, and 1.6 per 1000 (95% CI 1.5–1.7) live births when postneonatal CP was included.

**Interpretation:** The birth prevalence estimate of CP in HICs declined to 1.6 per 1000 live births. Data available from LMICs indicated markedly higher birth prevalence.

This original article is commented on by Paneth and Yeargin-Allsopp on pages 1436–1437 of this issue.

**Abbreviations:** ACPR, Australian Cerebral Palsy Register; HIC, high-income country; LMIC, low- and middle-income country; SCPE, Surveillance of Cerebral Palsy in Europe.

\*Members of the Global CP Prevalence Group are listed in the Acknowledgements.

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Cerebral palsy (CP) is an umbrella term for a group of disorders of movement and posture, caused by a non-progressive interference in the developing brain. Risk factors for CP span the periods before and around the time of conception, during pregnancy, the perinatal period, and up to 2 years of age. Known risk factors and conditions that can combine into causal pathways to CP include genetic variants, congenital anomalies, preterm birth, kernicterus, intrauterine growth restriction and infection, hypoxic ischaemia and cerebrovascular insults during pregnancy and in infancy, and accidental and non-accidental brain injury.<sup>1</sup>

Population-based CP registers and prevalence studies have monitored the birth prevalence of CP for more than 60 years.<sup>2</sup> The most recent systematic review and meta-analysis of birth prevalence, which mostly included births in the 1980s and 1990s, found prevalence was 2.1 per 1000 live births.<sup>3</sup> Historically, temporal fluctuations have been reported within high-income country (HIC) regions as some causal pathways become preventable, such as kernicterus, and others arose such as increased survival of infants born very preterm with the advent of neonatal intensive care units. In recent years, significant and sustained declines in the birth prevalence of CP in HIC regions of Europe, Australia, and Japan have been reported.<sup>4–7</sup> While the causes for this decline are complicated, declines are being attributed to an array of clinical improvements in public health, maternal, and perinatal care, particularly for infants cared for in a neonatal intensive care unit at highest risk of CP such as those born very preterm or at term with hypoxic–ischaemic encephalopathy.<sup>8</sup> It is therefore important to continue to monitor how improvements in care affect the current birth prevalence of CP across the world and draw attention to recent trends in CP. It is also important to determine whether trends are being seen in all HIC regions, as well as to establish the current overall birth prevalence of CP in these regions.

Population-based data are also now emerging from regions of low- and middle-income countries (LMICs) where higher rates of CP are being reported.<sup>9–11</sup> The aetiological pathways to CP in these countries seem to differ from HICs.<sup>12</sup> As most births worldwide occur in LMICs, it is imperative that an update in the prevalence of CP includes data from these regions where possible.

Prevalence of CP is not static and can be expected to continuously change as a result of medical advancements, and social and economic development. This study is the result of an international collaboration which aimed to provide a snapshot of recent changes, and current birth prevalence and period prevalence (complementary indicators). Specifically, this systematic analysis of CP register data and published literature aimed to identify the following: (1) trends in birth prevalence for CP of pre- or perinatal origin, postneonatal CP, and overall CP (live births) by region and combined for two major networks—the Surveillance of Cerebral Palsy in Europe (SCPE) and the Australian Cerebral Palsy Register (ACPR) since birth year 1995; (2) most recent birth prevalence estimate (live births) and period prevalence estimate

### What this paper adds

- Birth prevalence of pre-/perinatal cerebral palsy (CP) in high-income countries (HICs) is decreasing.
- Current overall CP birth prevalence for HICs is 1.6 per 1000 live births.
- Trends in low- and middle-income countries (LMICs) cannot currently be measured.
- Current birth prevalence in LMICs is markedly higher than in HICs.
- Active surveillance of CP helps to assess the impact of medical advancements and social/economic development.
- Population-based data on prevalence and trends of CP are critical to inform policy.

(children living in a region) of CP by region and combined for those with data available from birth year 2010 for a current prevalence estimate.

## METHOD

We sought to maximize the representation of geographical regions around the world and use the most contemporary data available. We conducted a systematic analysis of population-based data from two sources: (1) CP registers/surveillance systems and (2) published prevalence studies.

### Study population

The study population included children with CP (numerator) born from 1995 in regions of the world with population-based data, and the population in which they either were born (total live births) or resided (total children of the same age living in the same region) (denominator). Regions were classified by their country's World Bank income classification (low, lower middle, upper middle, high). Timing of CP was categorized as follows: (1) pre- or perinatal CP—brain injury/maldevelopment during the pre-, peri-, or neonatal period up to 28 days of life, or unknown aetiology; (2) postneonatal CP—a known brain-damaging event unrelated to factors in the ante-, peri-, or neonatal periods, sustained after the neonatal period (28 days of life) but before the age of 2 years; or (3) overall CP—all pre- or perinatal CP and postneonatal CP.

### CP registers/surveillance systems

In 2020, invitations to participate were sent to representatives from 30 population-based CP registers known to the study investigators. Registers provided the

aggregated number of children with confirmed CP born/living in their region, for each birth year from 1995, by timing of CP, along with equivalent live birth or population denominator data. Data collection was performed during 2020 to 2021.

CP was confirmed at a minimum age of 4 years.<sup>13</sup> For the study, the definition of CP<sup>14</sup> included the five criteria agreed on by SCPE and the ACPR: (1) is an umbrella term for a group of disorders; (2) is a condition that is permanent but not unchanging; (3) involves a disorder of movement and/or posture and of motor function; (4) is due to a non-progressive interference, lesion, or abnormality, and (5) the interference, lesion, or abnormality originates in the immature brain.<sup>15,16</sup> Registers/surveillance systems providing data included children with a diagnosis of CP at the age of 2 years, but who died before age 4 or 5 years, but excluded children with a diagnosis of CP who died before the age of 2 years.

We requested that registers provide descriptive data about the geographical region represented, size of region, continuity of data collection, numerator and denominator definitions, definition of CP, data sources and methods of data acquisition, and consent requirements to confirm inclusion. To be included in the trends analysis, CP registers/surveillance systems required a minimum of 10 consecutive years of data and ongoing data collection.

## Published literature on prevalence of CP

A broad systematic literature search strategy was designed with an academic librarian, on the basis of the search originally used by Oskoui et al.<sup>3</sup> Searches were conducted in MEDLINE and EMBASE in November 2020, along with handsearching. There were no limits on language of publication, but the search was limited to papers published from 2011, to include papers published since the systematic review by Oskoui et al.<sup>3</sup> Abstracts and titles were exported into referencing software, and automatic and manual de-duplication was performed (using Covidence systematic review software, Veritas Health Innovation, Melbourne, Australia; available at [www.covidence.org](http://www.covidence.org)). Titles and abstracts were screened for possible inclusion by one investigator (SG), then the full text of potential articles was retrieved and reviewed by two (SG and SM).

Original research articles were included if they reported population-based prevalence of CP from birth year 1995 with an internationally agreed definition of CP (denominators defined as live births or children aged between 0 and 18 years in the region). The following studies were excluded: (1) abstract available only; (2) studies describing a subgroup of CP only (e.g. severe motor involvement); (3) studies including people with CP outside the target age range (e.g. born before 1995; 50% of children younger than 4 years); and (4) studies from a region already represented in the current study with newer/equivalent data from a participating CP register/surveillance system or literature.

Sets of two investigators independently reviewed each article meeting all eligibility criteria (SG, SM, HSS, SJH, GH, KH, KM). Methods and results data were extracted using data extraction sheets designed a priori for the study (including reference, year of publication, geographical location of the study, birth cohorts included, study method, data sources, definition of CP and diagnostic criteria used, age at diagnosis/confirmation of diagnosis, definition and inclusion of postneonatal CP, and numerator and denominator definitions). The corresponding author was contacted, as required, to clarify information or data, and authors were asked to provide data from 1995 only. We preferentially extracted case and denominator data for live births, rather than children living in the region. When multiple prevalence rates were reported for children at different ages, data were extracted for the age group closest to age 5 years, when a diagnosis of CP is usually confirmed/verified. For aetiological timing, CP was categorized as pre-/perinatal if postneonatal CP was explicitly excluded; otherwise, data were categorized as 'overall' CP. If not reported, a denominator was estimated from the number of cases and prevalence of CP reported and noted in the accompanying tables.

## Quality and risk of bias assessment

CP registers/surveillance systems and included publications were critically appraised for quality and risk of bias. The JBI checklist for prevalence studies<sup>17</sup> was used, which includes nine quality items (marked as yes, no, unclear, or not applicable) and an overall appraisal to 'include' or 'exclude' the study for meta-analysis. Sets of two reviewers independently assessed each data source; discrepancies were resolved with an independent third reviewer (SG, SM, HSS, SJH, GH, KH, KM).

## Statistical analysis

### Objective 1a: recent temporal trends in each region

The temporal trend in the number of pre-/perinatal CP cases per 1000 live births, and the number of postneonatal CP cases per 10000 live births between 1995 and 2014 in each region was classified as increasing, decreasing, heterogenous, or no change. For each region, this classification was determined through a two-step process. First, a Mann–Kendall test<sup>15,16</sup> for monotonic trends was used to determine whether the birth prevalence rate for a given region was monotonically increasing or decreasing. The trend for a given region was classified as increasing if the resultant Kendall's  $\tau$  coefficient was positive and significant ( $p < 0.05$ ), and was classified as decreasing if the coefficient was negative and significant ( $p < 0.05$ ). Second, if the trend in a given region could not be classified as either increasing or decreasing (Kendall's  $\tau$  coefficient giving  $p > 0.05$ ), then a Poisson regression model with an offset term for live births and a smoothing spline term for birth year was

used to distinguish between a heterogenous temporal trend and the presence of no change in the birth prevalence rate. Cubic B-splines were used for all models, and the degree of smoothness was determined using the restricted maximum likelihood method. The presence of overdispersion in the Poisson regression was inspected. Regions where the smoothing spline term for birth year was significantly different from zero were classified as heterogenous, while regions where the spline term was not significantly different from zero were classified as no change. A smoothed trend line for each region was plotted to visualize birth prevalence trends.

### Objective 1b: combined recent temporal trends (register networks)

Temporal trends in the birth prevalence of pre-/perinatal and postneonatal CP using data from two large CP register networks, the SCPE and the ACPR, were analysed by Poisson regression models with an offset term for live births. Data from the two networks were pooled after testing for any difference in trends between them. Orthogonal polynomial terms for birth year up to the fourth degree were considered, and the final form of birth year in the model was selected using Akaike information criteria. A quadratic model (polynomial up to the second degree) was ultimately selected.

### Objective 2a: most recent prevalence in each region

Recent prevalence of pre-/perinatal, postneonatal, and overall CP were calculated for each region, with 95% confidence intervals (CI). Data for the two most recent birth years were used for pre-/perinatal CP, while data for any number of birth years from 2010 were used for postneonatal CP given the small number of individuals with postneonatal CP. The 95% CIs were calculated using approximation to the normal distribution after proportions were transformed using Freeman–Tukey double arcsine transformation.<sup>17</sup> Results are presented as prevalence rates after the pooled estimates were back transformed.

### Objective 2b: current combined global prevalence

Meta-analyses of ‘current’ prevalence of pre-/perinatal CP per 1000 live births, postneonatal CP per 10000 live births, and overall CP per 1000 live births were performed for regions with prevalence data for at least two consecutive birth years from 2010. The current birth prevalences of CP (pre-/perinatal, postneonatal, overall) were derived using univariate meta-analysis of proportions methods. Random effects meta-analyses using the DerSimonian and Laird method were preferred to fixed effects meta-analysis because of the anticipated heterogeneity of CP prevalence between regions. Heterogeneity between regional estimates

was assessed using the coefficients  $\tau^2$  and  $I^2$ .  $\chi^2$  tests of heterogeneity were also performed. These tests were only used descriptively.

Statistical analyses were performed using R version 4.1 (packages Kendall version 2.2,<sup>18</sup> gam version 1.2,<sup>19</sup> ggplot2 version 3.3.5,<sup>20</sup> meta version 5.2,<sup>21</sup> dmetar version 0.0.09,<sup>22</sup> mgcv version 1.8;<sup>23</sup> R Foundation for Statistical Computing, Vienna, Austria) and STATA version 14.2 (packages metaprop\_one; StatCorp, College Station, TX, USA).

### Ethical review

In accordance with the National Health and Medical Research Council National Statement on Ethical Conduct in Human Research (Australia), ethical review was not required for this study as it posed negligible risk and involved the use of existing collections of non-identifiable data (tabulated register data and published literature).

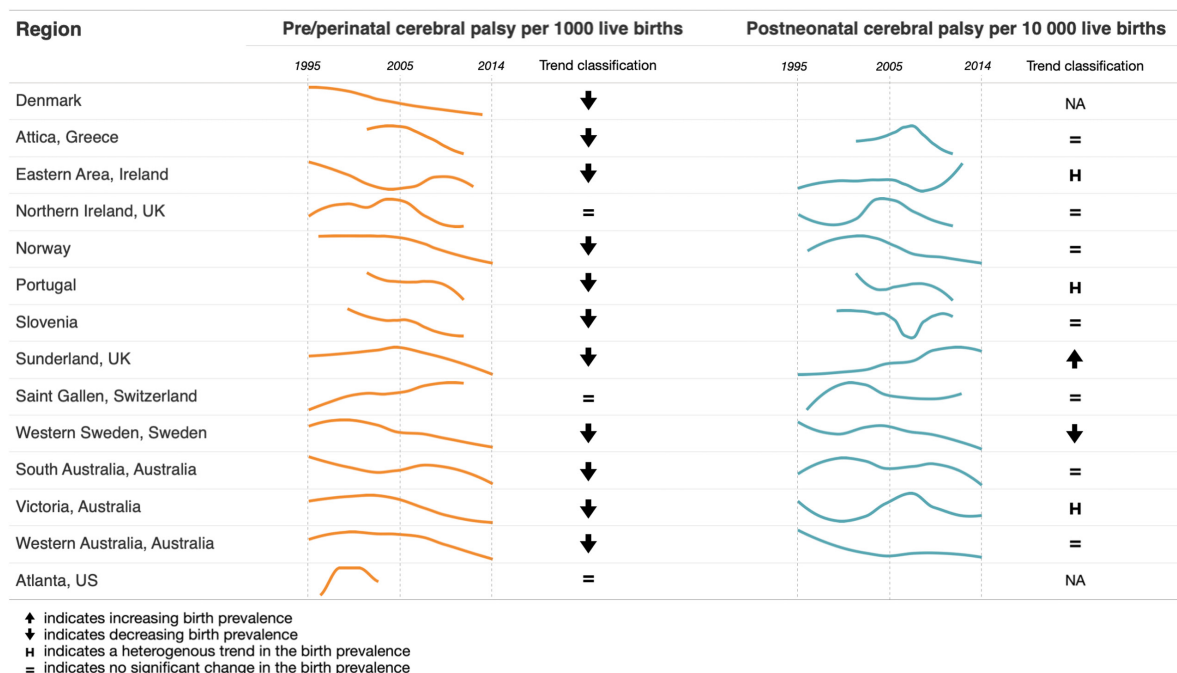
## RESULTS

Data from 41 regions of 27 countries were included. CP registers contributed data representing 19 regions from 15 countries, all from Europe and Australia, and classified as regions from HICs. Data from two register networks of Australia and Europe were also received (ACPR and SCPE). Published literature provided data from an additional 22 regions from 12 countries: Africa ( $n = 2$  regions from LMICs), Asia ( $n = 4$  regions from LMICs and  $n = 5$  regions from HICs), Europe ( $n = 1$  region from LMICs and  $n = 3$  from HICs), North America ( $n = 7$  regions from HICs) (Table 1 and Figure S1). No registers or studies were assessed as having a high risk of bias; therefore, all were included in at least one analysis (Tables 1, S1, and S2). Data sources not from registers ranged from face-to-face clinical assessments to administrative data linkages (Table 1).

All regions that were able to provide data for trend analyses were from HICs. Data for trend analyses were provided by regions from CP registers (13 of 14 regions). The remaining region from the USA used 1-year survivors as its denominator, and reported results from a surveillance system.<sup>24</sup> From the 14 regions covering over 8 million live births that contributed to the pre-/perinatal trend analysis, 79% showed a statistically significant decline. The regions reporting through to 2014 all showed a decline. The remaining three regions showed no change in the time period reported (Figure 1 and Table S3). However, the most recent data available for the USA were from the early 2000s, the Swiss region represents a very small population, and there have been recent declines in Northern Ireland, but not for the entire period for which data were available for this study (1995–2011).

From the 12 regions that were able to provide data for postneonatal CP, the pattern was mixed, with one region increasing, one decreasing, three being heterogenous, and seven showing no change. This mixed pattern was also seen in those that provided data through to 2014 (Figure 1 and Table S3).





**FIGURE 1** Birth prevalence trends of cerebral palsy.

There was no difference in the trends between SCPE and ACPR (test for interaction between birth year and register  $p = 0.67$ ); therefore data were combined for the two register networks. Figure 2 shows the combined trend line and 95% CIs for pre-/perinatal CP across Europe and Australia with a statistically significant declining trend ( $p = 0.012$ ). There was no change for postneonatal CP (figure not shown).

Birth prevalence estimates with 95% CIs were calculated for two CP register networks and 25 regions for pre-/perinatal CP, 21 regions for postneonatal CP (Figure 3), and 23 regions for overall CP (Figure S2). Most recent birth years were included in the analysis; however, they ranged from 1995 to 2014. Variation across regions reflect different birth years, size of the denominator population, and World Bank income levels. Two regions from LMIC had high birth prevalence estimates of 3 and 3.4 per 1000 live births, one of these regions also had a high postneonatal CP estimate.

Meta-analysis was restricted to regions with data since 2010 to obtain an estimate for current birth prevalence. A total of 17 regions were included in the analysis for both pre-/perinatal CP, postneonatal CP, and overall CP, all of which were HICs and from CP registers. Heterogeneity does exist between regions; however, for pre-/perinatal CP there was a combined estimate of 1.5 per 1000 (95% CI 1.4–1.6) live births ( $\tau^2 < 0.001$ ,  $I^2 = 69.4\%$ ) (Figure 4a). For postneonatal CP, the estimate of current birth prevalence was 0.8 per 10000 (95% CI 0.6–1.0) live births ( $\tau^2 < 0.001$ ,  $I^2 = 70.1\%$ ) (Figure 4b). For overall CP, the estimate of current birth prevalence was 1.6 per 1000 (95% CI 1.5–1.7) live births, seen in Figure S3 ( $\tau^2 < 0.001$ ,  $I^2 = 72.9\%$ ).

Twelve regions reported period prevalence of CP in children (denominator being children the same age living in

the area) (Figure 5). Estimated birth years for these studies ranged from 1995 to 2016 and covered over 7 million living children (Table S2). Four of the regions were from LMICs and prevalence ranged from 2.3 to 3.7 per 1000 children. Regions from HICs ranged from 1.6 to 2.9 per 1000 children; those with higher estimates included much earlier birth years.

## DISCUSSION

Before this paper, the most recent international CP birth prevalence study was published in 2013.<sup>3</sup> As such, the great majority of data included were from HICs and birth years from the 1980s and 1990s. At that time the overall birth prevalence was stable, and the estimate was 2.1 per 1000 live births. Since then, several studies have been published that suggest declines in birth prevalence have occurred in the 2000s.<sup>4,6</sup> This study was therefore undertaken to update our understanding of the global prevalence of CP by using contemporary data from CP registers and surveillance systems as well as published literature.

Our study confirmed that pre-/perinatal CP is declining in high-income regions in Europe and Australia. The trend was similar for individual regions and for the two major CP register networks, SCPE and ACPR. Only one other high-income region outside these networks was able to be included;<sup>24</sup> however, reporting for this study concluded in birth year 2002, and the same declining trend was not noted. Unfortunately, no registers in LMICs are yet able to report on trends, as at least a decade of population data is required to be meaningful. However, a recent systematic review, which used novel methods to predict trends, reported a concerning increasing trend in China.<sup>25</sup>

**TABLE 1** Regions included in the study

Region, country	Data or reference	Data source	World Bank income of country	Denominator	Participating in temporal trends (birth years)	Participating in most recent birth prevalence (birth years)
Cross River State, Nigeria	Duke et al. <sup>36</sup>	Key informant method + clinical assessment	Lower middle	Children	—	2003–2014
Eastern Uganda, Uganda	Kakooza-Mwesige et al. <sup>35</sup>	Door-to-door screening + clinical assessment	Low	Children	—	1998–2013
Shahjapur, Bangladesh	Khandaker et al. <sup>10</sup>	Bangladesh Cerebral Palsy Register	Lower middle	Live births	—	1998–2010
Rajshahi Division, Bangladesh	Murthy et al. <sup>43</sup>	Key informant method + clinical assessment	Lower middle	Children	—	1995–2013
Henan, China	Yuan et al. <sup>44</sup>	Clinical screening + assessment	Upper middle	Children	—	2005–2010
R.S. Pura Town, India	Raina et al. <sup>45</sup>	Door-to-door screening + clinical assessment	Lower middle	Children	—	1999–2003
Japan	Toyokawa et al. <sup>46</sup>	National health insurance claims	High	Children	—	2004–2009
Okinawa, Japan	Touyama et al. <sup>7</sup>	Okinawa Child Development Center surveillance database	High	Live births	—	1998–2007
Tochigi, Japan	Yamagishi et al. <sup>47</sup>	Survey from medical record	High	Live births	—	2009–2013
South Korea	Park et al. <sup>48</sup>	National health insurance data	High	Children	—	1999–2003
Taiwan	Chang et al. <sup>49</sup>	National health insurance data	High	Live births	—	1996–2000
Australia	Data	Australian Cerebral Palsy Register (ACPR)	High	Live births	1995–2014	2013–2014
New South Wales/Australian Capital Territory, Australia	Data	NSW/ACT Cerebral Palsy Register	High	Live births	—	2011–2012 <sup>a</sup>
Queensland, Australia	Data	Queensland Cerebral Palsy Register	High	Live births	—	2010–2011 <sup>a</sup>
South Australia, Australia	Data	South Australian Cerebral Palsy Register	High	Live births	1995–2014	2013–2014 <sup>a</sup>
Victoria, Australia	Data	Victorian Cerebral Palsy Register	High	Live births	1995–2014	2013–2014 <sup>a</sup>
Western Australia, Australia	Data	WA Register of Developmental Anomalies – Cerebral Palsy	High	Live births	1995–2014	2013–2014 <sup>a</sup>
Europe	Data	Surveillance of Cerebral Palsy in Europe (SCPE)	High	Live births	1995–2010	2009–2010
Belgium	Data	Belgian Cerebral Palsy Registry	High	Live births	—	2010–2011 <sup>a</sup>
Croatia	Data	Register of Cerebral Palsy of Croatia (RCP-HR)	High	Live births	—	2010–2011 <sup>a</sup>
Denmark	Data	Danish Cerebral Palsy Registry and National Cerebral Palsy Follow-Up Programme (CPOP)	High	Live births	1995–2013	2012–2013 <sup>a</sup>
Toulouse, France	Data	Childhood Disabilities Register of the Haute-Garonne County (RHE31)	High	Children	—	2010–2011

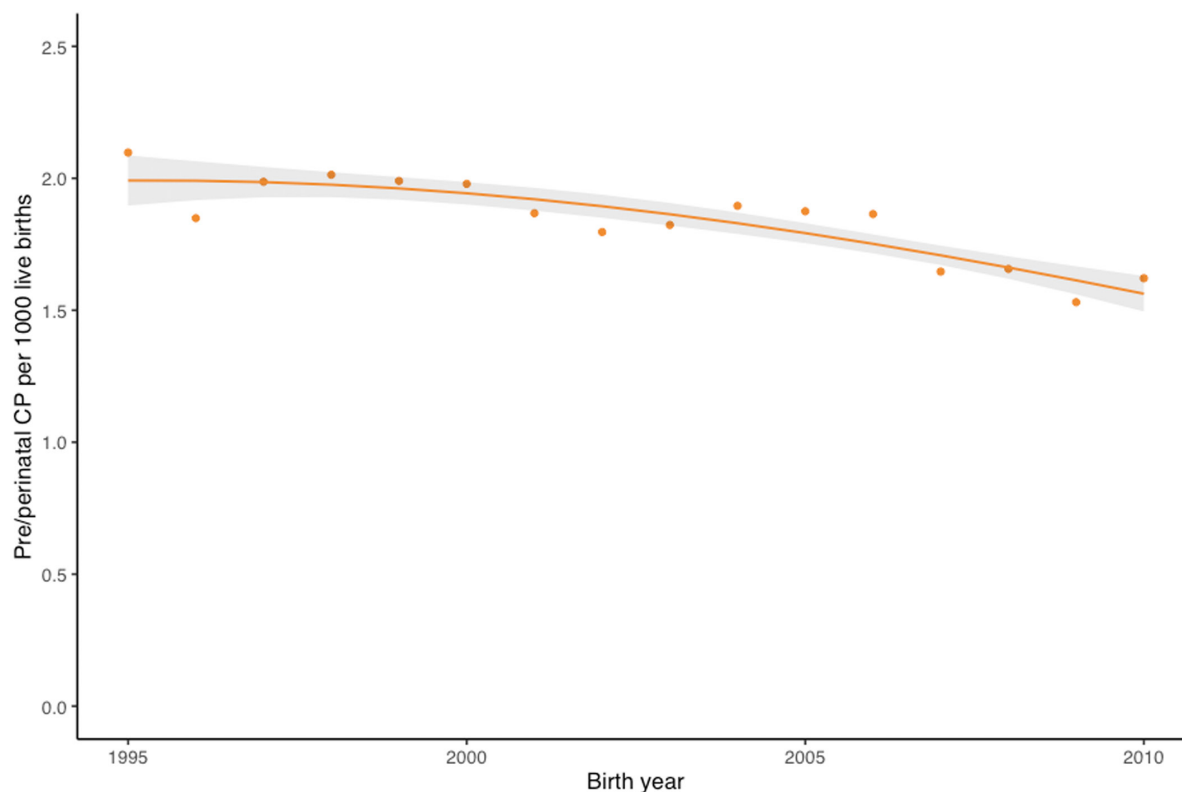
(Continues)

TABLE 1 (Continued)

Region, country	Data or reference	Data source	World Bank income of country	Denominator	Participating in temporal trends (birth years)	Participating in most recent birth prevalence (birth years)
Grenoble, France	Data	Register for childhood disabilities and perinatal survey (RHEOP)	High	Children	—	2009–2010
Attica, Greece	Data	The Cerebral Palsy Register of Attica	High	Live births	1999–2011	2010–2011 <sup>a</sup>
Borsod, Hungary	Fejes et al. <sup>50</sup>	Hospital, education, services record review	High	Live births	—	1995–2006
Eastern area, Ireland	Data	Eastern Area Cerebral Palsy Register	High	Live births	1995–2012	2011–2012 <sup>a</sup>
Moldova	Gincota Buftac et al. <sup>51</sup>	Hospital record review	Lower middle	Live births	—	2009–2010
Norway	Data	Norwegian Quality and Surveillance Registry for Cerebral Palsy (NorCP)	High	Live births	1996–2014	2013–2014 <sup>a</sup>
Portugal	Data	Portuguese Surveillance of Cerebral Palsy Programme	High	Live births	2001–2011	2010–2011 <sup>a</sup>
Slovenia	Data	Slovenian Register of Cerebral Palsy	High	Live births	1999–2011	2010–2011 <sup>a</sup>
Western Sweden, Sweden	Data	Cerebral Palsy Register of Western Sweden	High	Live births	1995–2014	2013–2014 <sup>a</sup>
Saint Gallen, Switzerland	Data	The Cerebral Palsy Register of St. Gallen (SPRN)	High	Live births	1995–2011	2010–2011 <sup>a</sup>
Northern Ireland, UK	Data	Northern Ireland Cerebral Palsy Register	High	Live births	1995–2011	2010–2011 <sup>a</sup>
North of England, UK	Glimianaia et al. <sup>52</sup>	North of England Collaborative Cerebral Palsy Survey (NECCPS)	High	Live births	—	1996–2000
Scotland, UK	Bugler et al. <sup>53</sup>	Cerebral Palsy Integrated Pathway Scotland (CPIPS) surveillance program	High	Children	—	1997–2016
Sunderland, UK	Data	Sunderland and area study	High	Live births	1995–2014	2013–2014 <sup>a</sup>
Northern Alberta, Canada	Robertson et al. <sup>54</sup>	Canadian Cerebral Palsy Registry	High	Live births	—	2008–2010
Ontario, Canada	Ray et al. <sup>55</sup>	Pre-existing administrative linked dataset	High	Neonatal survivors	—	2002–2008
Quebec, Canada	Oskoui et al. <sup>56</sup>	Cerebral Palsy Register (REPACQ)	High	Children	—	1999–2001
USA	Zablotsky and Black <sup>57</sup>	National health interview survey	High	Children	—	1997–2015
Alabama, Georgia, Missouri, Wisconsin, USA	Durkin et al. <sup>58</sup>	Cerebral Palsy surveillance program (ADDM Network)	High	Children	—	2002
Metropolitan Atlanta, Georgia, USA	Van Naarden Braun et al. <sup>24</sup>	Cerebral Palsy surveillance program (MADDSP)	High	1-year survivors	1996–2002	2002
South Carolina, USA	Li et al. <sup>59</sup>	Medicaid services, hospital discharge abstracts, department of disabilities/special needs	High	Live births	—	2009

<sup>a</sup>Included in meta-analysis of current birth prevalence.





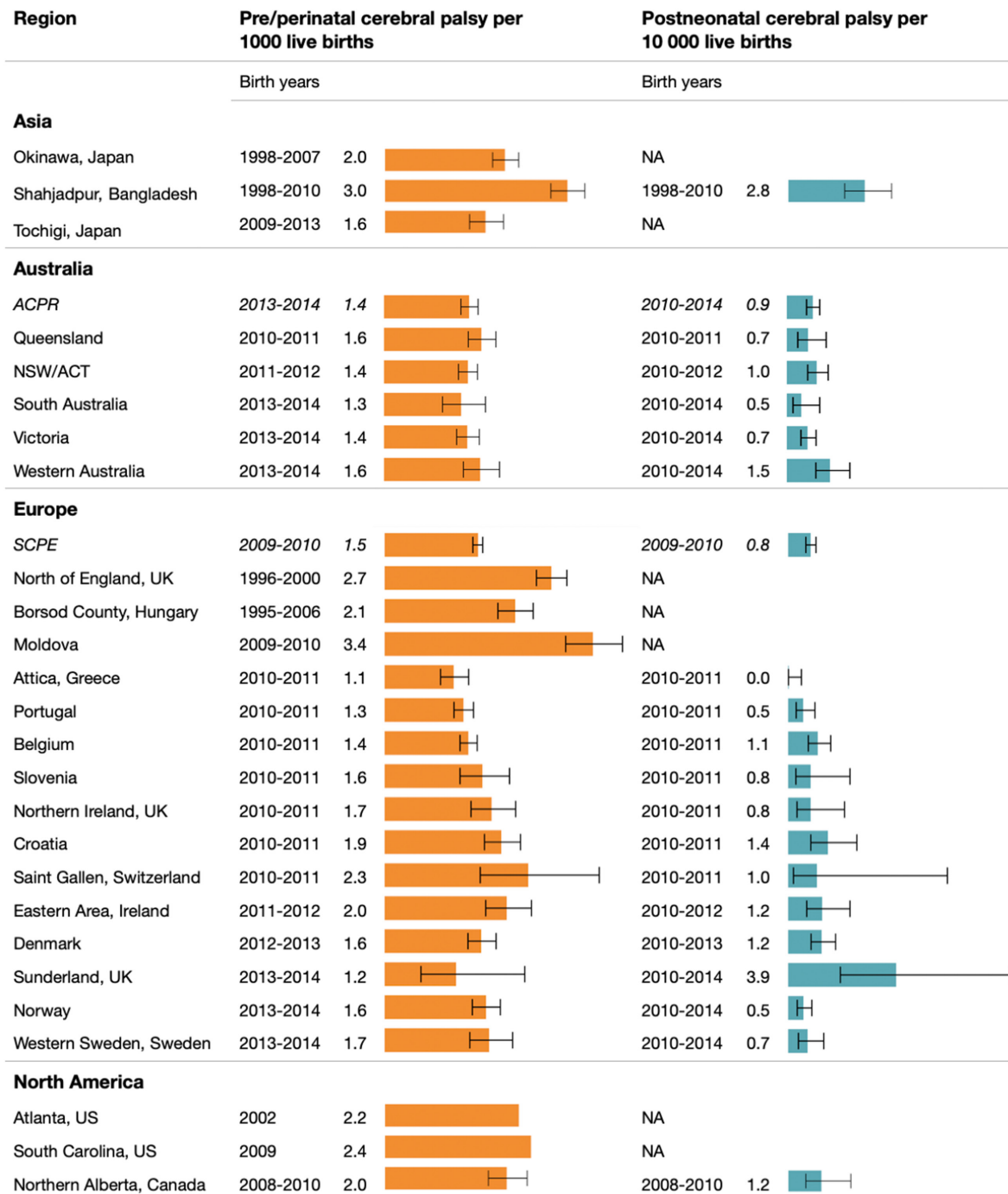
**FIGURE 2** Birth prevalence trend of pre-/perinatal cerebral palsy (CP; Surveillance of Cerebral Palsy in Europe and the Australian Cerebral Palsy Register combined).

Neonatal intensive care units are expanding in China and other LMICs, and the increased survival of medically fragile infants born at increasingly lower gestational ages may result in an initial spike in CP prevalence, while further development of neonatal care may decrease the prevalence again—as has been seen in HICs.<sup>26</sup> A challenge for all is how to share knowledge, experiences, and lessons learnt to minimize this inevitable spike. A recent large randomized controlled trial of therapeutic hypothermia has shown that we cannot assume that standard interventions in HICs will work in the same way in LMIC settings;<sup>27</sup> evaluation of such interventions is essential before being introduced into new settings and prevention opportunities should remain a priority.<sup>28</sup>

To calculate current global birth prevalence estimates, we restricted meta-analyses to regions with more than one birth year from 2010. No LMICs were able to participate in these analyses, so these primary findings are for high-income regions only. The current pre-/perinatal CP birth prevalence is 1.5 per 1000 live births. The current overall (including post-neonatal) CP birth prevalence is 1.6 per 1000 live births. This prevalence estimate is 25% lower than the overall birth prevalence estimate reported in 2013 (2.1 per 1000),<sup>3</sup> and this updated current birth prevalence estimate for HICs should now be used. This is particularly encouraging as this decline has occurred during the same era that survival in neonatal intensive care units is improving for infants born extremely preterm.<sup>31</sup> As described earlier, we have learnt to expect that

advances in health care may lead to increases in CP prevalence, as well as decreases.

The number of CP registers and prevalence studies in LMICs is increasing, yet they remain extremely under-represented. In this study, 7 out of 41 regions were from LMICs (Nigeria, Uganda, Bangladesh [ $n = 2$ ], China, India, Moldova) compared with 3 out of 49 in the previous study (China, Kenya, Turkey).<sup>3</sup> Two regions were able to report birth prevalence using live births as a denominator, making this comparable to the high-income regions. Birth prevalence was 3.3 per 1000 overall for Shahjampur, Bangladesh, and 3.4 per 1000 for pre-/perinatal CP in Moldova. These birth prevalence estimates are more than double the findings for high-income regions in our meta-analysis. The remainder reported period prevalence (with a denominator of children living in the region) as high as 3.7 per 1000 children in Rajshahi Division, Bangladesh. Additional literature from Albania, Egypt, and Pakistan, which could not be included in this review, supports these findings.<sup>32–34</sup> Low- and middle-income regions reporting prevalence estimates suggest that these are almost certainly underestimates due to survival bias (i.e. high mortality in the early years, before a CP diagnosis and, again, high mortality in children with CP), incomplete ascertainment, and inability to include very mild cases at population level.<sup>10,35,36</sup> Collaborative efforts such as mentorship programmes with SCPE, ACPR, and the Global LMIC CP register will increase representation of LMICs.<sup>37</sup>

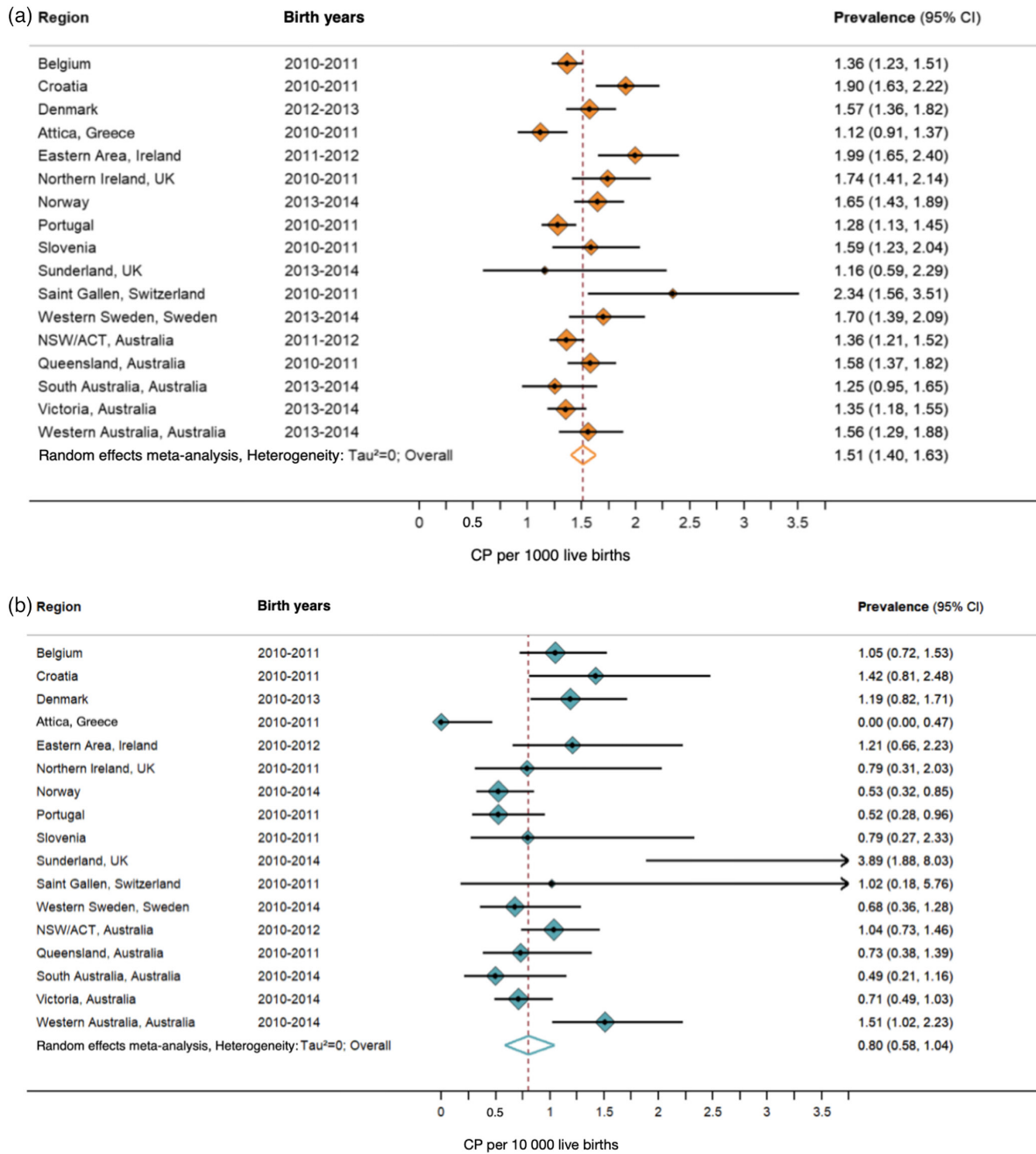


**FIGURE 3** Regional birth prevalence of cerebral palsy. ACPR, Australian Cerebral Palsy Register; ACT, Australian Capital Territory; NSW, New South Wales; SCPE, Surveillance of Cerebral Palsy in Europe.

There has been no change in postneonatal CP, and the current estimate for HICs is 0.8 per 10 000 live births with wide confidence intervals. The numbers for postneonatal CP in HICs are small, and we have less confidence about these trends, particularly for children with a brain injury closer to the age of 2 years, which may be described as an acquired brain injury rather than postneonatal CP. LMICs with higher proportions of postneonatal CP (up to 36% in Nigeria,<sup>36</sup> compared with 6% in Australia<sup>2</sup>), alert us to the

differences in aetiologies of postneonatal CP (e.g. malaria, previous nutritional status) and potentially different opportunities for prevention strategies that are specific to each region.<sup>37,38</sup>

A shared understanding of the definition and classification of CP is essential for reliable estimates and trends. Standardized and consistent approaches used by registers enable accurate monitoring of the condition over time. In situations when complete agreement cannot be reached, data



**FIGURE 4** (a) Current pre-/perinatal birth prevalence of cerebral palsy (CP) in high-income countries. (b) Current post neonatal birth prevalence of CP in high-income countries. ACT, Australian Capital Territory; CI, confidence interval; NSW, New South Wales.

can be harmonized for comparisons.<sup>39</sup> For example, in this study, data were restricted to CP that occurred in the first 2 years of life, and children who survived beyond the age of 2 years, despite variations between CP registers in these limits.<sup>13</sup> We recommend the continued use of papers such as the one by Smithers-Sheedy et al.<sup>13</sup> (including confirming diagnosis at age 4 years) and the full annotation that describes in detail the definition of CP.<sup>14</sup>

A strength of this study was our reporting on both birth prevalence and period prevalence, which is rarely done. These two indicators are complementary. While birth

prevalence is a relevant indicator of the impact of the organization of care and practices in the peri- and neonatal period, the cross-sectional approach used for period prevalence estimates is more relevant for documenting public health issues, notably the impact of CP in the community. It is generally accepted that period prevalence is higher than birth prevalence, which is consistent with our results, although comparisons are difficult (small sample size and different regions). Another strength of this study was our representation of regions without CP registers, by including published population-based studies with alternative methodologies,



**FIGURE 5** Period prevalence of overall cerebral palsy for children.

such as large surveys and administrative data. However, it is known that conditions, including CP, are coded inconsistently,<sup>40</sup> so risk of bias was higher for studies that solely relied on administrative data for identifying children with CP. Finally, we observed regional heterogeneity in our birth prevalence meta-analyses. Although there will always be a level of true variation in prevalence between regions, it is likely that under-ascertainment of children with mild CP is also a contributing factor, particularly for newly established registers.<sup>41</sup>

The declining trends in the birth prevalence of pre/perinatal CP, evidenced by CP registers in this paper, increases our understanding of the condition and the impact of improvements in ante-, peri-, and postnatal care in HICs. This global overview represents the recent and current situation in over 40 regions of the world. Sustainable registers with good ascertainment are essential for continued monitoring of trends and prevalence, and the real-world impact of changing social development and health care in low-, middle-, and high-income countries. Recognition of CP at national and international levels provides a powerful tool to potentially influence policy and services, leading to a demonstrable contribution to society and economies.<sup>42</sup>

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## DATA AVAILABILITY STATEMENT

Data sharing is available on request to the authors, and if data is from a register it would require approval from the individual register.



## ORCID

Sarah McIntyre  <https://orcid.org/0000-0002-0234-1541>  
 Shona Goldsmith  <https://orcid.org/0000-0003-3903-6142>  
 Annabel Webb  <https://orcid.org/0000-0001-8435-4436>  
 Virginia Ehlinger  <https://orcid.org/0000-0003-4992-5998>  
 Sandra Julsen Hollung  <https://orcid.org/0000-0002-7486-7454>  
 Karen McConnell  <https://orcid.org/0000-0002-5221-9800>  
 Catherine Arnaud  <https://orcid.org/0000-0002-4002-802X>  
 Hayley Smithers-Sheedy  <https://orcid.org/0000-0002-0082-2413>  
 Maryam Oskoui  <https://orcid.org/0000-0003-1042-0108>  
 Gulam Khandaker  <https://orcid.org/0000-0002-0661-4113>  
 Kate Himmelmann  <https://orcid.org/0000-0002-3959-9554>

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## SUPPORTING INFORMATION

The following additional material may be found online:

**Figure S1:** Data sources flow chart.

**Figure S2:** Overall regional birth prevalence of cerebral palsy.

**Figure S3:** Current overall birth prevalence of cerebral palsy in high-income countries.

**Appendix S1:** Search strategy.

**Table S1:** JBI/supporting data for regions represented in birth prevalence analyses.

**Table S2:** JBI/supporting data for regions represented in period prevalence analyses.

**Table S3:** Supporting data for regional trends in pre-/perinatal cerebral palsy and postneonatal cerebral palsy.

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