PREVALENCE OF PRIMARY IMMUNE THROMBOCYTOPENIA IN OKLAHOMA

Deirdra R. Terrell¹, Laura A. Beebe¹, Barbara R. Neas¹, Sara K. Vesely¹, Jodi B. Segal³, and James N. George¹,²

¹Department of Biostatistics & Epidemiology, College of Public Health, University of Oklahoma Health Sciences Center, Oklahoma City, OK ²Department of Medicine, College of Medicine, University of Oklahoma Health Sciences Center, Oklahoma City, OK ³Department of Medicine, Johns Hopkins University School of Medicine, Baltimore, Maryland

Abstract

Objective—To determine the prevalence of ITP in Oklahoma regardless of age, clinical characteristics, insurance status, and source of health care.

Study Design and Setting—Patients with ITP were identified by the administrative code ICD-9-CM 287.3 in Oklahoma hematologists’ offices for a 2-year period, 2003–2004. Prevalence was estimated separately for children (<16 years old) and adults because of their distinct clinical characteristics. Oklahoma census data for 2000 was used as the denominator.

Results—Eighty-seven (94%) of 93 eligible Oklahoma hematologists participated; 620 patients with ITP were identified. The average annual prevalences were: 8.1 (95% CI 6.7, 9.5) per 100,000 children, 12.1 (95% CI 11.1, 13.0) per 100,000 adults, and 11.2 (95% CI 10.4, 12.0) per 100,000 population. Among children and adults less than age 70 years, the prevalence was greater among women. Among adults aged 70 years and older, the prevalence was greater among men. The highest prevalence of ITP was among men age 80 years and older.

Conclusion—These data document for the first time the prevalence of ITP regardless of age, clinical characteristics, insurance status, and source of health care. The methodology developed for this prevalence analysis may be adaptable for epidemiologic studies of other uncommon disorders which lack specific diagnostic criteria and are treated primarily by medical specialists.

Keywords

primary immune thrombocytopenia; ITP; prevalence; demographics; administrative data; ICD-9-CM 287.3

INTRODUCTION

Primary immune thrombocytopenia (ITP) is an uncommon acquired autoimmune disorder characterized by isolated thrombocytopenia without an apparent alternative etiology.(1–3) Although ITP rarely causes death from bleeding,(4,5) the risks for bleeding,(6) side effects from treatments,(7) and symptoms of fatigue(8) can substantially affect the lives of patients...
with ITP and their families. (6) Although the effect of ITP on the quality-of-life of individual patients has been documented, (6,9–11) understanding the population burden of ITP requires knowledge of its prevalence. Knowledge of the prevalence of ITP is also important for the development of new treatments, as emphasized by recent pharmaceutical-supported studies of the epidemiology of ITP. (12–14)

The annual incidence of ITP is between 1.9 and 6.4 per 100,000 children and 3.3 per 100,000 adults. (15) Because ITP is a chronic persistent disorder in some children and most adults, the prevalence is expected to exceed the annual incidence. However the prevalence of ITP has only been reported in four studies of four different patient groups. (14;16–18) None of these four studies identified patients with ITP of all ages regardless of their clinical characteristics, insurance status, and their source of health care. Therefore the objective of this study was to determine the prevalence of ITP in the State of Oklahoma regardless of age, clinical characteristics, insurance status, or source of health care.

METHODS

Identification of hematologists

Hematologists in Oklahoma were identified by searching the Oklahoma Board of Medical Licensure and the American Osteopathic Association websites for physicians who listed hematology, oncology, hematology-oncology, and pediatric hematology-oncology as a specialty. Hematologists were included if they had an active Oklahoma license and were currently treating hematology patients in Oklahoma. Current practice was determined by contacting the hematologists’ offices.

Identification of patients with ITP

In our previous study, we developed a definition for a “definite diagnosis of ITP” using administrative data and we identified children and adults who had a definite diagnosis of ITP by a hematologist as an outpatient at the Oklahoma University Medical Center (OUMC) between 1995 and 2004. (19) In the current study, all other hematologists in Oklahoma were surveyed. Data were requested from the hematologists’ office billing managers for all visits of all patients billed with the ICD-9-CM code of 287.3 in the first, second, or third position between 1995 and 2004. If the name of the office or clinic had ‘hospital’ as part of its name, the billing manager confirmed the patients were seen in an outpatient clinic and submitted only outpatient data. Data from the Oklahoma City and Muskogee Veterans Administration (VA) Hospitals were also restricted to billing for outpatient visits to a hematology clinic. Patients in the Oklahoma City VA Hospital hematology clinic were not included in our previous OUMC analysis (19) but were seen by OUMC hematologists; patients in the Muskogee VA Hospital were seen by their medical staff hematologists. No other VA clinics in Oklahoma have hematologists on their staff. The Indian Health Service was contacted to confirm that none of their clinics were practice sites for any of the state’s hematologists. The last four digits of the social security number, date of birth, and gender were requested to identify individual patients, to prevent duplicate counting of patients who had received care by multiple hematology practice groups. Due to confidentiality, some hematology offices would not release the last four digits of the social security number. As a result, exact matching to identify individual patients was performed using gender and date of birth. Patients’ city and state was also requested and patients were excluded if they were not residents of the state of Oklahoma at the time of service.

Definitions

Time period—Since ITP is an uncommon disorder, (15) a period of 10 years, 1995–2004, was initially selected to identify a sufficient number of patients, some of whom may see a
hematologist infrequently or only for their initial evaluation. The 10-year period was also used in case complete data for all years were not available for all hematologists' offices. Therefore it was assumed that there would be some years within this period when data would be available for all or most hematologists in Oklahoma. Period prevalence estimates were calculated for the 10-year period, the five-year period, and the two-year period. The decision for which time period may provide the best estimate of prevalence was based on [1] the precision of the estimates assessed by the width of the 95% confidence intervals and [2] the completeness of the data obtained assessed by the number of hematologists and number of patients included. Ultimately the two-year period (2003–2004) was reported.

**Age stratification**—Children and adults were analyzed separately because of their distinct clinical features that suggest a different pathogenesis. Most children have acute ITP with a sudden onset of severe, symptomatic thrombocytopenia which spontaneously resolves within six to 12 months. In contrast, most adults have chronic ITP with an insidious onset of moderate to severe thrombocytopenia which rarely resolves spontaneously. Throughout adolescence, chronic ITP becomes more common. Therefore the age defining childhood was set as less than 16 years, the age commonly used in studies of the incidence of ITP. There was no minimum age criterion.

**Prevalence estimation**

Census year 2000 population data, the midpoint of this analysis period, were used as the denominator for all prevalence calculations. The total population of Oklahoma in 2000 was 3,450,654; for children age less than 16 years, 784,688; for adults 16 years and older, 2,665,966. Some hematology offices could not retrieve billing information for the entire 10-year period. Due to differences in completeness of the data received, prevalence was estimated for three different time periods: 10-year (initial service dates 1995–2004), five-year (initial service dates 2000–2004), and two-year (initial service dates 2003–2004). Prevalence estimates were calculated for all patients and also separately for children and adults. Age was calculated for each patient at each service date. If a patient was less than 16 years old at the time of the initial service date, he/she was included in the prevalence estimate for children. If during the time period the patient’s age was initially less than 16 years and then greater than or equal to 16 years, two service dates were documented, the initial service date at age less than 16 years and the initial service date after age greater than or equal to 16 years. Therefore some patients were included in analyses of both children and adults if they were seen before and after their sixteenth birthday. However, in the combined prevalence estimates of children and adults, each patient was counted only once. The Poisson asymptotic 95% confidence interval (CI) was calculated and reported.

The 95% confidence intervals were calculated based on a Poisson distribution and the standard error of a Poisson distribution is based on the square root of the numerator. Since more patients were observed over the longer time periods of five and 10 years and since the denominator per year remained the same with each calculation, confidence interval were wider with the longer periods of observation. After the best period prevalence estimate was determined, one-year prevalence estimates were used to calculate an annual average prevalence. Sensitivity analyses to adjust the observed prevalence estimates for the effects of potential limitations of the methods used to identify patients with ITP were performed. Sensitivity analyses adjusted for the following assumptions: [1] that the administrative code ICD-9-CM 287.3 was sufficiently accurate (as determined by calculation of the positive predictive value [PPV] of the code) to identify both children and adults with ITP, [2] that most patients with ITP were diagnosed and/or managed by hematologists, and [3]...
that most patients with ITP were diagnosed and/or managed as outpatients. Data were stored
in a password protected Microsoft Access® database. Analyses were performed using SAS

**Ethics approvals**

This study was approved by the Institutional Review Board (IRB) of the University of
Oklahoma Health Sciences Center (OUHSC). Additional approval from the Oklahoma City VA
Research and Development Committee was obtained to review data for patients at both
the Oklahoma City and Muskogee VA hospitals. Site-specific IRB approval was obtained
from two large hematology-oncology offices. For all other hematology offices, OUHSC IRB
approval was sufficient.

**RESULTS**

**Identification of hematologists**

Ninety-eight hematologists were identified in Oklahoma; five were excluded (one whose
office was outside of Oklahoma, three who only treated general internal medicine patients,
and one who only treated oncology patients). Therefore 93 hematologists were eligible to
participate in this study. Of the 93 eligible hematologists, data were obtained from 88
(95%); five hematologists had no retrievable data from 1995–2004 because of new billing
systems established after 2004. Table 1 presents the data from the 15 practice groups for the
88 hematologists who contributed to this analysis.

In addition to the 93 eligible hematologists, 41 additional hematologists were identified who
had practiced in Oklahoma at some time during 1995–2004 but who were no longer active
during the time this study was performed, 2007–2008. Nine of these 41 hematologists did
not see patients during 1995–2004; 22 were associated with one of the groups participating
in this study and it was assumed that their patients would have continued their care within
this group and those records were obtained. The remaining 10 hematologists may have had
patients who were not included in the prevalence estimates.

**Prevalence estimates**

The 10-year period prevalence estimate included 342 children and 817 adults. The
numerator for the combined 10-year period prevalence for all patients was 1,147 after
excluding 12 children who aged into the adult category during these 10 years. The five-year
period prevalence estimate included 215 children and 716 adults. The numerator for the
combined five-year period prevalence for all patients was 923 after excluding eight children
who aged into the adult category during these five years. The two-year period prevalence
estimate included 107 children and 515 adults. The numerator for the combined two-year
period prevalence for all patients was 620 after excluding two children who aged into the
adult category during these two years.

The two-year period prevalence estimates were determined to be better estimates than
estimates from 10-year or five-year data, based on better precision assessed by the 95%
confidence intervals (Table 2) and because all but one of the 88 participating hematologists
could provide data for 2003–2004 (Table 1). For adults, hematology groups 6 and 7 were
only able to provide data for these two years and they contributed a large number of patients.
The two-year period prevalence estimates were used to calculate annual prevalence
estimates. The numerators for annual prevalence estimates for children were 69 for 2003 and
58 for 2004; for adults were 309 for 2003 and 335 for 2004. The average annual prevalence
estimates for children, adults and all patients are presented in Table 3.
Sensitivity analyses

Table 4 presents the prevalence estimates independently adjusted for PPV of the ICD-9-CM code, exclusion of inpatients, exclusion of patients treated by a non-hematologist, and for all three variables combined, using data previously determined by medical record review at OUMC.(19) When the observed prevalence estimates were adjusted for the previously determined PPV for the ICD-9-CM code of 287.3 of 0.72 for children and 0.69 for adults, the prevalence estimates decreased. When the observed prevalence estimates were adjusted for the 2% of children and 8% of adults who may have been diagnosed and managed for their ITP only by physicians other than hematologists and for the 5% of children and 10% of adults who may have been diagnosed and managed for their ITP only as inpatients, the adjusted prevalence estimates increased. When all three factors were considered together, the adjusted prevalence estimates were lower than the observed prevalence estimates, although they were still contained within the 95% confidence interval of the observed estimates.

Age and gender-specific prevalence estimates

The age and gender distributions of patients in 2003 and 2004 are illustrated in Figure 1 and descriptively compared. Although the absolute prevalence estimates were slightly different between these two years, the relative estimates among different age groups and between genders were similar. Prevalence was slightly greater among children less than 10 years old than among children and adolescents 10–19 years old. In both of these groups the prevalence among girls slightly exceeded the prevalence among boys. The prevalence of ITP peaked among women aged 20–29 years relative to women less than age 50 years, while the prevalence of ITP decreased among men aged 20–29 years relative to men less than age 50 years. Among adults aged 20–39 years, the prevalence among women was substantially greater than among men. The prevalence of ITP increased every decade after age 50 years, with the peak occurring among persons 80 years and older. Among persons greater than or equal to 70 years, the prevalence among men was greater than among women.

DISCUSSION

Estimating prevalence is essential to understand the population burden of a disorder. However estimating the prevalence of ITP is challenging because there are no specific diagnostic criteria that document the presence of ITP; the diagnosis can only be established by excluding other causes of thrombocytopenia.(1–3) In addition, ITP is uncommon(15) and therefore potentially unfamiliar to both physicians and medical record coders.

To determine the prevalence of ITP regardless of age, clinical characteristics, insurance status, or source of health care, we surveyed hematologists in the State of Oklahoma to determine the number of patients identified as outpatients utilizing the administrative code ICD-9-CM 287.3. This method was based on three assumptions: [1] that the administrative code ICD-9-CM 287.3 was sufficiently accurate to identify both children and adults with ITP, [2] that most patients with ITP were diagnosed and/or managed by hematologists, and [3] that most patients with ITP were diagnosed and/or managed as outpatients. The validity of these three assumptions was established by our previous review of OUMC medical records in which we established a definition for a definite diagnosis of ITP.(19) Data from this prior study documented that the PPV of the administrative code ICD-9-CM 287.3 for identifying patients with a definite diagnosis of ITP by a hematologist was 0.72 for children and 0.69 for adults, similar to a previous report on the accuracy of this administrative code for identification of patients with ITP which determined a PPV of 0.71.(23) We also documented that in 95% of children and 92% of adults with a definite diagnosis of ITP, the diagnosis was established by a hematologist; this supported the validity of including only
hematologists and not all physicians for our determination of prevalence. Finally, we
determined that in 95% of children and 83% of adults with a definite diagnosis of ITP
established by a hematologist, the diagnosis was established as an outpatient; this supported
the validity of surveying only the offices of hematologists and not records of all hospitals.
Therefore these data documented that a survey of hematologists’ offices for patients with
ITP identified by the administrative code ICD-9-CM 287.3 was an appropriate method to
determine the prevalence of patients with ITP. Because we were successful in obtaining data
from 88 (95%) of the state’s 93 hematologists, this was also a feasible method to determine
the prevalence of ITP in Oklahoma.

Adjustment of the average annual prevalence estimates for all three of the assumptions in
our method resulted in lower average annual prevalence estimates for both children (7.4 per
100,000 children) and adults (11.1 per 100,000 adults) but both adjusted estimates were
within the 95% confidence intervals of our observed estimates. Therefore adjustment for
these three assumptions did not significantly impact the magnitude of the observed
prevalence estimates.

In children, ITP is typically an acute and spontaneously resolving disorder. ITP resolves
within six months of initial diagnosis in 70% of children(20) and within 12 months in nearly
80% of children.(11) Consistent with the uncommon persistence of ITP, the annual
incidence of chronic ITP in children, defined as persistence for more than 6 months after
diagnosis, has been reported to be 0.46 per 100,000 children,(24) while the annual incidence
of all (acute and chronic) children with ITP is between 1.9 and 6.4 per 100,000 children.(15)
Since the clinical course of ITP in most children is less than one year, it may be expected
that the annual prevalence of ITP in children would be similar to the annual incidence.
However the average annual prevalence for ITP in children, estimated from our data, 8.1 per
100,000 children, is higher than the reported annual incidence.(15) One possible explanation
for the higher prevalence may be continued follow-up of children after the ITP resolves. The
prevalence of chronic ITP in children has been reported to be 4.6 per 100,000 children,(16)
approximately half of our observed prevalence for all (acute and chronic) children with ITP.
Our average annual prevalence estimate for children of 8.1 per 100,000 children (95% CI
6.7, 9.5) is not appreciably different from the previously reported annual prevalence estimate
for children of 7.2 per 100,000 children aged 1–14 years.(17)

In adults, ITP is typically a chronic disorder, commonly persistent for many months or years
unless effectively treated by splenectomy or immunosuppressive agents.(1–3) Therefore it
would be expected that the prevalence would be greater than the incidence. Our average
annual prevalence estimate for adults of 12.1 per 100,000 adults (95% CI 11.1, 13.0), is
nearly four-fold greater than the incidence of ITP in adults, 3.3 per 100,000 adults per year.
Segal and Powe(17) previously reported the annual prevalence of ITP for privately insured
patients ages 1–64 years to be 9.6 (95% CI, 8.6, 11.0) per 100,000 persons (Table 4).
Feudjo-Tepie, et al.(14) previously reported the annual prevalence of chronic ITP in
privately insured adults ages 18–64 years to be 7.1 for 2002 and 9.5 for 2004 per 100,000
adults. Using data from the United Kingdom General Practice Research Database, Bennett,
et al.(18) previously reported the annual prevalence for adults 18 years old and older for
2003 to be 4.0 per 100,000 adults and for 2004 to be 4.6 per 100,000 adults. The annual
prevalence of Segal and Powe(17) is lower than our estimate of the prevalence for children
and adults and the annual prevalence of Feudjo-Tepie, et al.(14) is lower than our estimate
of the prevalence for adults. Because of differences in inclusion criteria these estimates are
not directly comparable. However, our estimate of prevalence was higher probably because
we included patients over 64 years old and included patients without private insurance. Our
annual prevalence estimates are higher than the adult estimates from Bennett, et al. (18) possibly due to the differences between cases identified from administrative data compared to electronic medical records or differences between the USA and UK.

Our prevalence data for children are consistent with previous reports that the annual incidence of ITP in children is greater among children younger than age 10 years than among children 10 years and older. (20;25) The predominance of women among patients age 20–39 years is similar to the increased relative frequency of other autoimmune disorders such as systemic lupus erythematosus in young women. (26) The progressively increasing prevalence among older people and a greater prevalence of men relative to women among people 60 years and older is notable. Segal and Powe (17) and Bennett, et al. (18) also described increased prevalence among older adults. The increased prevalence of ITP among older people is consistent with previous reports that the annual incidence of ITP is greatest among the oldest age cohorts; (5;13;27) in one of these reports the increased incidence among older males was also described. (13)

A limitation of this study was that not all hematology offices across the state were able to provide data for the entire time 10-year time period. However estimates derived for different time periods maximized the use of the available data. The two-year prevalence estimates represented 94% of the state’s hematologists. However, the two-year estimate is a conservative estimate of prevalence because some ITP patients may not visit a hematologist within a two-year period. This study also required diagnosis of ITP by a hematologist, a requirement that could limit the comparability of our results to previous studies. However the requirement for diagnosis by a hematologist reflects the current referral practice in Oklahoma. A survey of primary care providers in the Oklahoma Practice-based Research Network documented that 75% of respondents were ‘likely’ to send a patient with moderate thrombocytopenia (platelet count 30,000/μL) to a hematologist for further evaluation and management. The likelihood of referral increased to 85% when the moderate thrombocytopenia was associated with mild bleeding symptoms (petechiae) and to 92% when the patient presented with severe thrombocytopenia (platelet count 10,000/μL). (28) Some hematology offices may focus more on oncology rather than benign hematology which could raise concern in potential differences in diagnostic coding between groups.

In summary, our prevalence estimates are generalizable beyond privately insured persons and not limited by age. The generalizability of our data is supported by the similarity of our estimates to estimates previously reported from Maryland. (17) This study also provides the best current estimates of the age and gender distribution of ITP throughout the population. These data will be important to assess the population burden of ITP.

Acknowledgments

This project supported in part by the Utay Family Blood Research Fund. Dr. Terrell is partially supported by NIH 1U01HL72283-09S1.

Reference List


Figure 1.
Figure 1a. ITP Gender and Age-Specific Prevalence, Oklahoma
Figure 1b.
Table 1

Oklahoma hematology practice groups and their patients with ITP

<table>
<thead>
<tr>
<th>Hematologist groups</th>
<th>Individual hematologists</th>
<th>Patients eligible for analysis</th>
<th>Years of Data</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>Children</td>
<td>Adults</td>
</tr>
<tr>
<td>1 (OUMC)</td>
<td>23</td>
<td>220</td>
<td>120</td>
</tr>
<tr>
<td>2 (OKC VA *)</td>
<td>0</td>
<td>44</td>
<td>44</td>
</tr>
<tr>
<td>3</td>
<td>32</td>
<td>2</td>
<td>348</td>
</tr>
<tr>
<td>4</td>
<td>9</td>
<td>1</td>
<td>120</td>
</tr>
<tr>
<td>5</td>
<td>6</td>
<td>0</td>
<td>6</td>
</tr>
<tr>
<td>6</td>
<td>4</td>
<td>0</td>
<td>40</td>
</tr>
<tr>
<td>7</td>
<td>3</td>
<td>0</td>
<td>71</td>
</tr>
<tr>
<td>8</td>
<td>2</td>
<td>122</td>
<td>16</td>
</tr>
<tr>
<td>9</td>
<td>2</td>
<td>0</td>
<td>8</td>
</tr>
<tr>
<td>10</td>
<td>2</td>
<td>0</td>
<td>11</td>
</tr>
<tr>
<td>11</td>
<td>1</td>
<td>0</td>
<td>3</td>
</tr>
<tr>
<td>12</td>
<td>1</td>
<td>1</td>
<td>0</td>
</tr>
<tr>
<td>13</td>
<td>1</td>
<td>0</td>
<td>10</td>
</tr>
<tr>
<td>14</td>
<td>1</td>
<td>2</td>
<td>35</td>
</tr>
<tr>
<td>15</td>
<td>1</td>
<td>0</td>
<td>12</td>
</tr>
<tr>
<td>Total</td>
<td>88</td>
<td>348</td>
<td>844</td>
</tr>
</tbody>
</table>

Total individual patients **

342  817  1147

* Oklahoma City VA Hospital is staffed by OUMC hematologists

** Total individual patients after excluding duplicate patients reported by more than one site. Some patients were included in both the children and adults estimates because they were seen both before and after their sixteenth birthday; these patients were only counted once in the overall prevalence estimates. Therefore the total number is 1,147 rather than 1,159.
### Table 2

Comparison of ITP period prevalence estimates by age categories and years of data

<table>
<thead>
<tr>
<th></th>
<th></th>
<th></th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Children &lt;16 years</td>
<td>43.6 (39.0, 48.2)</td>
<td>27.4 (23.7, 31.1)</td>
<td>13.6 (11.1, 16.2)</td>
<td>8.1 (6.7, 9.5)</td>
</tr>
<tr>
<td>Adults ≥16 years</td>
<td>30.6 (28.5, 32.7)</td>
<td>26.9 (24.9, 28.8)</td>
<td>19.3 (17.6, 21.0)</td>
<td>12.1 (11.1, 13.0)</td>
</tr>
<tr>
<td>All Patients</td>
<td>33.2 (31.3, 35.2)</td>
<td>26.7 (25.0, 28.5)</td>
<td>18.0 (16.6, 19.4)</td>
<td>11.2 (10.4, 12.0)</td>
</tr>
</tbody>
</table>

*Prevalence estimates, with 95% confidence intervals in parentheses, are per 100,000 children for children, per 100,000 adults for adults, and per 100,000 population for all patients.

**Average annual prevalence was calculated from the 2003 and 2004 one-year ITP prevalence estimates.
Table 3
Sensitivity analyses to adjust average annual prevalence estimates for methodology assumptions

<table>
<thead>
<tr>
<th>Methodology assumptions</th>
<th>Prevalence estimates</th>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Average annual prevalence estimate per 100,000 children</td>
<td>Average annual prevalence estimate per 100,000 adults</td>
<td></td>
</tr>
<tr>
<td>Observed prevalence estimate</td>
<td>8.1 (95% CI 6.7, 9.5)</td>
<td>12.1 (95% CI 11.1, 13.0)</td>
<td></td>
</tr>
<tr>
<td>Independently adjusted for PPV</td>
<td>6.5</td>
<td>8.9</td>
<td></td>
</tr>
<tr>
<td>Independently adjusted for exclusion of patients diagnosed and managed only by non-hematologists</td>
<td>8.3</td>
<td>13.1</td>
<td></td>
</tr>
<tr>
<td>Independently adjusted for exclusion of patients diagnosed and managed only as inpatients</td>
<td>8.5</td>
<td>13.3</td>
<td></td>
</tr>
<tr>
<td>Adjusted for all 3 variables</td>
<td>7.4</td>
<td>11.1</td>
<td></td>
</tr>
</tbody>
</table>

The observed average annual 2003 and 2004 prevalence estimates for children and adults are presented with the 95% confidence intervals per 100,000 children and adults, respectively. These estimates were adjusted for the independent effects of positive predictive value (PPV), exclusion of patients managed only by non-hematologists, exclusion of inpatients, and the overall effect of all three variables on the annual average prevalence estimates.
Table 4

Previously reported estimates of the prevalence of ITP*

<table>
<thead>
<tr>
<th>Study</th>
<th>Country</th>
<th>Reported Results/10^5/year</th>
<th>Age of subjects</th>
<th>Source of patients</th>
<th>Identification of patients</th>
</tr>
</thead>
<tbody>
<tr>
<td>1(17)</td>
<td>USA</td>
<td>9.6 (2002)</td>
<td>1–64 years</td>
<td>Medical Care Database of the Maryland Health Care Commission</td>
<td>Two diagnoses with the ICD-9-CM code 287.3 separated by a minimum of 30 days</td>
</tr>
<tr>
<td>2 (14)</td>
<td>USA</td>
<td>7.1 (2002) 9.5 (2004)</td>
<td>18–64 years</td>
<td>US Integrated Healthcare Information Database</td>
<td>Two diagnoses with the ICD-9-CM code 287.3 separated by a minimum of six months</td>
</tr>
<tr>
<td>3 (18)</td>
<td>United Kingdom</td>
<td>4.0 (2003) 4.6 (2004)</td>
<td>18 years and older</td>
<td>United Kingdom General Practice Research Database (GPRD)</td>
<td>At least one diagnostic Read code 42P2.11, D313.12, D313000, D313012 for ITP in the patient’s electronic medical record in the GPRD</td>
</tr>
</tbody>
</table>

*In addition to these three reports, one additional report on the prevalence of chronic ITP in children(16) was not included in this Table because it was not comparable to our data.