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# The incidence of diagnostic error in medicine

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## ABSTRACT

A wide variety of research studies suggest that breakdowns in the diagnostic process result in a staggering toll of harm and patient deaths. These include autopsy studies, case reviews, surveys of patient and physicians, voluntary reporting systems, using standardised patients, second reviews, diagnostic testing audits and closed claims reviews. Although these different approaches provide important information and unique insights regarding diagnostic errors, each has limitations and none is well suited to establishing the incidence of diagnostic error in actual practice, or the aggregate rate of error and harm. We argue that being able to measure the incidence of diagnostic error is essential to enable research studies on diagnostic error, and to initiate quality improvement projects aimed at reducing the risk of error and harm. Three approaches appear most promising in this regard: (1) using 'trigger tools' to identify from electronic health records cases at high risk for diagnostic error; (2) using standardised patients (secret shoppers) to study the rate of error in practice; (3) encouraging both patients and physicians to voluntarily report errors they encounter, and facilitating this process.

In God we trust, all others bring data<sup>1</sup>

The patient safety movement in the USA has entered its second decade. A wide range of important safety concerns have been studied, and to this point, including medication errors, hospital-acquired infections, wrong-site surgery and a host of other issues. Strangely lacking, however, is a concerted effort to find, understand and address diagnostic errors.<sup>2–4</sup> One factor that may contribute to its relative neglect is that the true incidence of diagnostic error is not widely appreciated. Measuring the rate of error and, in particular, error-related harm,<sup>5</sup> would provide the necessary motivation to begin addressing this large and silent problem. How likely is a diagnosis to be

wrong, missed, or egregiously delayed? How often do diagnostic errors cause harm? In this report, we briefly summarise the methods that have been used to estimate the rate of diagnostic error, and comment on their relative merits and limitations. A more comprehensive presentation of studies using each of these methodologies has been presented elsewhere.<sup>6</sup>

## THE INCIDENCE OF DIAGNOSTIC ERROR

Arthur Elstein, a cognitive psychologist interested in 'how doctors think', studied clinical decision making for his entire career and concluded the diagnosis is wrong 10–15% of the time.<sup>7</sup> A diverse range of research approaches that have focused on this issue over the past several decades suggest that this estimate is very much on target.<sup>6</sup>

The incidence of diagnostic error has been estimated using eight different research approaches (table 1).

*Autopsy studies* identify major diagnostic discrepancies in 10–20% of cases. Most cases in autopsy series derive from inpatient settings, but they also include deaths from the emergency department which, for many reasons, is considered to be the natural laboratory for studying diagnostic error. Although autopsies have virtually disappeared in the USA, autopsies are still common in many other countries, and despite the availability of modern imaging, continue to show diagnoses being missed that might have been lifesaving, particularly infections and cardiovascular conditions.

Although autopsy data is considered the 'gold standard' in terms of providing the most definitive data on the accuracy of diagnosis, only a subset of cases ever reach autopsy, and in many cases, the relationship between clinical diagnoses and autopsy findings remains unclear.



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**Table 1** Research approaches used to estimate the incidence of diagnostic error

Research approach	Findings—examples	Suitable for evaluating incidence	Suitable for evaluating aetiology
Autopsies	Major unexpected discrepancies that would have changed the management are found in 10–20% <sup>8 9</sup>	Yes	No
Patient and provider surveys	One-third of patients relate a diagnostic error that affected themselves, a family member, or close friend <sup>10</sup> ; Over half the surveyed paediatricians report making a diagnostic error at least once or twice a month <sup>11</sup>	Limited	Limited
Standardised patients	Internists misdiagnosed 13% of patients presenting with common conditions to clinic (COPD, RA, others) <sup>12</sup>	Yes	Yes
Second reviews	10–30% of breast cancers are missed on mammography <sup>13</sup> ; 1–2% of cancers are misread on biopsy samples <sup>14</sup>	Yes	No
Diagnostic testing audits	Errors related to laboratory testing are the most common reason for a diagnostic error. <sup>15 16</sup>	Very limited	Limited
Malpractice claims	Problems relating to diagnostic error are the leading cause for paid malpractice suits in every large system	Very limited	Limited
Case reviews (cross-sectional studies by symptom, disease, or condition); (may be enriched by trigger tools)	Patients with asthma—median delay in making the correct diagnosis was 3 years, or 7 visits <sup>17</sup> ; 12–51% of patients with subarachnoid haemorrhage are misdiagnosed in the emergency department. <sup>18</sup> Of 1000 hospital deaths, 5% were considered preventable, and the most frequent aetiology was diagnostic error. <sup>19</sup>	Yes	Limited
Voluntary reports	1674 reports of diagnostic error were submitted to the UK's National Reporting and Learning System over a 2-year period, 0.5% of all incidents reported <sup>20</sup>	Yes	Yes

COPD, chronic obstructive pulmonary disease; RA, rheumatoid arthritis.

Autopsies also discover a large number of incidental findings that were not suspected during life, but that were clinically irrelevant.

*Surveys* have found that diagnostic errors are a major concern of both patients and physicians. A survey of over 2000 patients found that 55% listed a diagnostic error as their chief concern when seeing a physician in an outpatient setting.<sup>21</sup> Similarly, physician surveys have consistently found that approximately half the respondents encounter diagnostic errors at least monthly.<sup>11 22 23</sup> Moreover, compared with the many different safety concerns encountered in practice, physicians perceive diagnostic errors to be more likely to cause serious harm or death compared with other safety concerns.<sup>24</sup>

*Standardised patients* studies using 'secret shoppers' have also been used to estimate the accuracy of diagnosis. In these studies, real or simulated patients with classical presentations of common diseases, like rheumatoid arthritis, asthma, or chronic obstructive pulmonary disease (COPD) are sent anonymously into real practice settings.

The diagnostic error rates reported (13–15%) are very much in line with estimates from the other types of research approaches, and have substantial 'face' validity in that the studies are being carried out prospectively in real-world settings. In addition to providing an estimate of diagnostic error rates, this approach offers the unique ability to probe the various factors that promote or detract from optimal diagnosis.<sup>25 26</sup>

The chief limitation of these studies in regard to estimating error rates is that they are performed under research-directed conditions, presenting a much

smaller subset of conditions than would be seen in usual practice. Moreover, in studies seeking to also study the factors relevant to accurate diagnosis, these patients may present with comorbid conditions or contextual complexities that are not representative of typical patients, and case complexity is a major factor in determining diagnostic accuracy.<sup>27</sup>

*Second reviews* refers to research protocols in the visual subspecialties (eg, radiology, pathology, dermatology) where a second radiologist examines the same films after a first radiologist, or a second pathologist reviews the same biopsy or cytology specimen as another pathologist. These second review studies may be performed under controlled conditions, involving the review of many or mostly abnormal cases. This approach has advantages from a research perspective, but substantially increases the possibility for diagnostic error, which can range from 10% to 50%.<sup>28–31</sup> Interestingly, the studies also show that a diagnostician will also disagree with his or her own prior interpretation in a small fraction of cases.

In 'real world' situations, the majority of examinations are normal. Under these conditions, a critical abnormality is detected by a second expert reviewer in the range of 2–5%.

*Diagnostic testing audits* are used to estimate the incidence of error in the clinical laboratory. Thanks to impressive advances in quality control procedures, diagnostic errors in the modern age are rarely the result of an error in the analytical test itself. Most laboratory-related errors now originate from the pre-analytical and postanalytical phases, namely issues related to the physician ordering and interpreting the

test result. Overall, it is estimated that laboratory results are misleadingly wrong in 2–4% of cases, and roughly the same error rates are found in diagnostic radiology. The major limitation of diagnostic testing audits is that, although they excel at finding errors in test performance per se, they underestimate the true clinical impact because of the related preanalytical and postanalytical errors occurring outside the walls of the laboratory or radiology department which, typically, escape detection. Lapses in the reliable communication of abnormal test results, for example, is a universal problem, even in systems with advanced electronic medical records.<sup>32 33</sup>

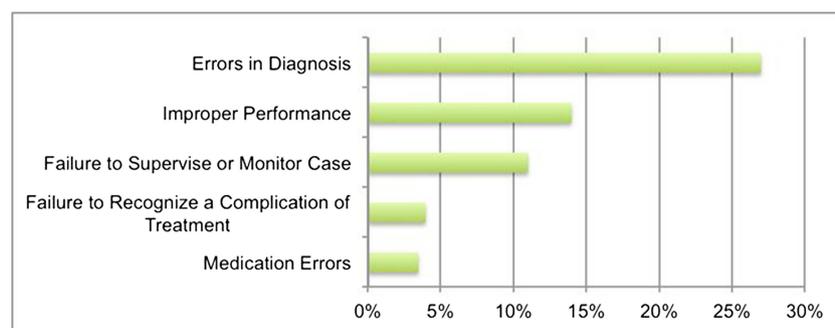
*Malpractice claim databases* are easily studied and have provided interesting data on diagnostic errors. Typical data (eg, from the Physicians Insurers Institute of America, shown below) reveals that problems related to diagnostic error are the leading cause of paid claims. Essentially identical data has been reported from the Department of Veterans Affairs, Kaiser-Permanente Healthcare System, and CRICO-RMF (CRICO Risk Management Foundation), the self-insurance entity covering the Harvard teaching hospitals, and from studies in the UK.<sup>34</sup> A recent analysis of malpractice cases extracted from the National Practitioner Data Bank over a 25-year period identified 100 249 cases of diagnostic error.<sup>35</sup> Diagnostic error was the most common reason for a claim (29%) and the most costly, averaging \$386 849 per claim.

Studying closed claims provides a convenient approach to analysing large numbers of cases, already preclassified as representing diagnostic errors. Limitations of this approach include the fact that only a small subset of errors results in claims, and these, typically, span just a narrow range of clinical conditions, predominated by cases involving cancer or cardiovascular disease in young or middle-aged patients. Moreover, these cases provide only limited opportunities to study the aetiology of diagnostic errors, as the proceedings by their nature are focused more on consequences than causes.

*Case Reviews.* Many specific symptoms and disease states have been studied using retrospective case reviews, and in each of these the incidence of

diagnostic error is unacceptably high. For example, a systematic review of more than 8000 ER patients found a delayed diagnosis of stroke in 9%.<sup>36</sup> Kostopoulou and colleagues reported a systematic review of this literature and identified 21 studies meeting their quality criteria.<sup>37</sup> In one such study, children with asthma experienced a median delay in making the correct diagnosis of more than 3 years, spanning 7 clinic/ER visits. Delayed or wrong diagnosis rates of 10–50% have been identified in studies of coronary artery disease, HIV-associated complications, tuberculosis and a wide range of malignancies. Very similar data has been reported by Gandhi and colleagues, who studied 181 cases of diagnostic error in ambulatory settings (figure 1).<sup>16</sup> Delayed diagnosis of cancer and coronary artery disease were the most commonly identified problems in the study by Gandhi *et al*, and a growing number of studies have confirmed the frequency of missed opportunities for earlier cancer diagnosis.<sup>31 38 39</sup> The point to be made is that whereas diagnostic errors may sometimes reflect encounters with extremely rare diseases or very unusual presentations of common diseases, in many cases, it is a relatively common disease that is mislabelled or missed entirely. A convenience sampling of studies that measured diagnostic error rates in 40 different symptom complexes or diseases is presented in online supplementary appendix table 1.

The chief limitation of case reviews is that they typically rely just on data contained in the medical record, and many diagnostic errors are missed as a result. As an example, Schwartz and colleagues used standardised patients to estimate costs resulting from diagnostic errors in ambulatory practices. Only their knowledge of the actual diagnosis allowed accurate estimates of these costs; virtually none could have been surmised from record reviews alone.<sup>26</sup> An additional limitation is that the medical record, typically, is lacking documentation on what the clinician was thinking at the time the diagnosis was being considered. Finally, random chart reviews are not well suited to measuring incidence rates of diagnostic error because the rate of error is low, thus requiring a large number of reviews.



**Figure 1** Top alleged medical error named in claims where the patient expired (Physician Insurers Association of America (PIAA) Data Sharing Project Data 1985–2009, Physician Insurer, Vol 55, 2010).

**Voluntary reports.** Voluntary error-reporting systems are now in place in most healthcare organisations in the USA, and were expected to provide a reliable way of identifying both minor and major adverse events. When they are used, and especially if follow-up interviews with providers can be obtained, voluntary reporting offers the unique potential to explore both the system-related and cognitive aetiologies of diagnostic error.<sup>40 41</sup>

Unfortunately, these programmes capture only a small fraction of diagnostic errors. Factors that detract from reporting include the time required to submit cases, a natural reluctance to call attention to one's own mistakes, and the ever-present fear of provoking a malpractice suit despite the reassurance that reports can be submitted anonymously. Another barrier to voluntary reporting of diagnostic errors is the lack of a 'common format' for reporting. The Agency for Healthcare Research and Quality has taken the lead in developing 'common formats' which healthcare organisations can use to report other types of patient safety incidents, but at the present time there is no specific common format tool to report diagnostic errors.

## METHODS CURRENTLY USED TO IDENTIFY DIAGNOSTIC ERRORS IN PRACTICE

Diagnostic error rates are being measured in very few, if any, healthcare organisation in the USA. With regard to ambulatory settings, Tsang and colleagues recently reviewed the methods available for measuring adverse events and found that none were helpful in identifying diagnostic errors.<sup>42</sup> The situation is no better for capturing diagnostic errors involving inpatients: A recent Inspector General study of 785 hospitalised Medicare beneficiaries using five different approaches identified a 13% incidence of serious adverse events, but not a single episode categorised as diagnostic error.<sup>43</sup>

## SUGGESTIONS ON IMPROVING MEASUREMENT

Given the limitations of the error-detection approaches currently in use, how can healthcare organisations begin to identify diagnostic errors? We suggest three novel approaches:

### Consider using trigger tools

Various types of trigger tools are now being used in quality programmes,<sup>44</sup> the most widely known of which is the 'Global Trigger Tool' developed by the Institute for Healthcare Improvement.<sup>45</sup> A growing fraction of US healthcare organisations are using this instrument, but it was designed specifically to identify treatment errors, specifically errors of commission, and is poorly suited to detecting diagnostic errors, many of which are errors of omission.<sup>46</sup> A more promising approach was recently reported by Singh and colleagues, who designed an electronic trigger to identify patients with an unscheduled hospitalisation within 2 weeks of a primary care visit. The frequency

of diagnostic errors in patients meeting this criterion was about 20%, compared with just 2% for unselected patients.<sup>47-49</sup> A similar process identifying hospitalisations after an ER 'treat and release' visit has recently been described by Newman-Toker *et al.*<sup>50</sup> Electronic surveillance is also effective in identifying diagnostic errors through discrepancies between laboratory and pharmacy records,<sup>51</sup> and discovering diagnostic errors through data mining is probably just around the corner. Although the use of trigger tools will not capture all diagnostic errors, their use will substantially enrich the yield compared with random chart reviews and, thus, bring more errors to attention.<sup>52</sup>

### Encourage and facilitate voluntary or prompted reports from patients

There is substantial preliminary evidence that patients are acutely aware of the diagnostic error problem.<sup>10 53</sup> Several authors have called for patients to take a more active role in ensuring the reliability of the diagnostic process<sup>54-56</sup> and enlisting their help to identify breakdowns is a logical, practical and simple approach to explore. Weingart and colleagues, for example, surveyed 228 inpatients who reported 20 adverse events, of which 11 were verified in the medical record but none were captured in the hospitals SE-detection programmes.<sup>57</sup> Reports from Canada,<sup>58</sup> Japan<sup>59</sup> and Sweden<sup>60</sup> have similarly found that patients are both willing and capable of participating effectively in identifying errors in their care.

The chief limitation of soliciting error reports from patients is that they will require verification by healthcare providers. As evident from the Weingart study and others,<sup>61</sup> not all their safety concerns accurately reflect true lapses in care.

### Encourage and facilitate error reporting from physicians

At least two different approaches have successfully improved on the power of voluntary error reporting systems to capture diagnostic errors. Phillips and colleagues encouraged physicians, their staff and patients to report all safety concerns over just a constrained 10-week period, and to focus on specific 'intensive reporting' days. The approach yielded 935 reports from 10 practices, including 12 instances of suspected diagnostic error.<sup>62</sup> Trowbridge was also successful in obtaining diagnostic error reports by acting as a clinical champion to encourage participation from colleagues.<sup>63</sup> In pilot work with this process at Maine Medical Center, 36 diagnostic errors were reported over 6 months that otherwise would have escaped detection. The severity of errors uncovered was high; 73% involving moderate or serious harm to the patient. By comparison, the facility identified only six diagnostic errors from their standard, existing systems for identifying medical errors during the same time period.

## DISCUSSION

We have proposed that measuring the incidence of diagnostic error in everyday practice is an essential requirement of a comprehensive quality management programme.<sup>2</sup> Of eight methods used to study diagnostic errors, some are more suitable than others in terms of their potential for providing meaningful data on diagnostic error rates (table 1). Analysing closed malpractice claims can provide only relative data on error rates; estimating absolute rates is not possible because so few true errors actually result in claims, and not all claims reflect true errors. Diagnostic testing audits are meaningful only to the extent that audits examine true clinical impact, because many diagnostic testing errors are ultimately detected and corrected, or discarded as meaningless. Of the remaining methods, each has its own advantages and limitations. Moreover, it is probably safe to assume that each of these approaches will underestimate the actual rate of error, and may identify entirely separate cohorts of errors. This problem is compounded by the different definitions and classification systems of diagnostic error that have been applied in these various studies. The goal of knowing the diagnostic error rate in practice may, therefore, require further research to standardise definitions and other methodologic issues, and combining results from several different approaches.

Similarly, the eight different approaches offer differing capabilities in their ability to provide insight regarding the aetiology of diagnostic error. Autopsies and second reviews reveal that an error was made, but not why. Using standardised patients is a particularly powerful way to study these factors, because at least some of the variables (case presentation and complexity, for example) can be controlled. Voluntary reports from physicians also provide unique opportunities to gather insights on the cognitive and system-related factors that might have contributed to the error. With the exception of using standardised patients, none of the approaches are well suited to study the human factor issues, such as distractions, fatigue and workload stress, thought to play dominant roles influencing clinical decision making.

Knowing the incidence of diagnostic error may be less important than being able to measure the likelihood of harm that results.<sup>5</sup> Extrapolating from the Class 1 errors (a major discrepancy that likely leads to the patient's death) identified at autopsy, Leape, Berwick and Bates estimated that 80 000 deaths per year might be caused by diagnostic error, including both ambulatory and inpatient errors.<sup>64</sup> A recent systematic review of autopsy data concluded that 36 000 deaths a year were due to diagnostic errors in just ICUs alone.<sup>65</sup> These estimates, of course, do not include the many instances of non-fatal injury related to misdiagnosis, events that will be far more numerous, and the many instances where the harm is psychological or financial more than physical.

Preventability, however, is a difficult parameter to judge, and these estimates may exaggerate the impact of diagnostic error to the extent that it is overestimated.<sup>66</sup> Both case review studies and closed claims studies<sup>67 35</sup> find that diagnostic errors are more likely to cause harm than other patient safety problems.

*In summary*, a wide range of different research approaches have been used to estimate diagnostic error rates, all suggesting that the incidence is unacceptably high. Although true incidence data are lacking, a wide variety of research studies suggest breakdowns in the diagnostic process result in a staggering toll of harm and patient deaths. A recent, authoritative review by the AMA of ambulatory patient safety concerns reached the same conclusion.<sup>68</sup>

What's missing from these estimates are true incidence data, as typically the denominators are not available or provided. Also missing are the aggregate rates of injury and harm. There is a clear need for additional research to identify better ways to bring diagnostic errors to light. Promising approaches in this regard include the use of trigger tools focusing on diagnostic error, the use of standardised patients, and encouraging both patients and physicians to voluntarily report errors they encounter.

The most fundamental principle of performance improvement is that 'You can't fix what you don't measure'. Efforts to begin addressing diagnostic error must begin with measurement. In no area of patient safety is this need more acute than in trying to identify the true incidence of diagnostic errors, and the harm associated with these events.

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**Appendix – The Incidence of Diagnostic Error in Medicine**

**Table 1: Diagnostic Error Rates in Specific Conditions**

<i>Mikulin and Hardcastle (1987)(1)</i>	<b>Gastric cancer</b>	Of 83 patients with gastric cancer, the median delay in diagnosis was 7 weeks
<i>Lee et al (1987)(2)</i>	<b>Myocardial infarction</b>	In a prospective multicenter study of 3077 patients seen in the ER with chest pain, 477 had acute MI's, and of these 18 (4%) had been discharged from the ER without making the diagnosis
<i>Jones et al (1990)(3)</i>	<b>Asthma</b>	Of 187 children with asthma, diagnosis delayed in many and the length of delay was longer in older children
<i>Charlton et al (1991)(4)</i>	<b>Asthma</b>	<i>Of 212 children with asthma, the median delay in making the correct diagnosis was 3 years (or 7 office visits)</i>
<i>Schnetler (1992)(5)</i>	<b>Oral Cancer</b>	<i>Of 96 cases seen in 3 oral surgery departments, the referring general practitioner had made the correct diagnosis in only 52%.</i>
<i>Sorensen et al (1992)(6)</i>	<b>Meningitis</b>	<i>Of 177 patients with meningococcal meningitis, the diagnosis was delayed in 52%</i>
<i>McCarthy et al (1993)(7)</i>	<b>Myocardial infarction</b>	<i>In a multicenter study of 5773 patients seen in the ER with suspicious chest pain, the diagnosis of MI was made correctly in 1030 and missed in 20 (1.9%)</i>
<i>Lederle et al (1994)(8)</i>	<b>Ruptured aortic aneurysm</b>	Review of all cases at a single medical center over a 7 yr period. Of 23 abdominal aortic dissections, the diagnosis was initially missed in 14 (61%)
<i>Bleeker et al (1995)(9)</i>	<b>Myocardial infarction</b>	Of 300 patients admitted with myocardial infarction, the median delay from symptoms to admission was 84 minutes
<i>Paauw et al (1995)(10)</i>	<b>HIV complications</b>	Patients with classical complications were sent to 134 primary care physicians: Kaposi's sarcoma was noticed by 47% and misdiagnosed by 45%; Oral hairy leukoplakia was noticed by 56% and misdiagnosed by 59%; Lymphadenopathy was noticed by only 17%
<i>Beam et al (1996)(11)</i>	<b>Breast cancer</b>	50 accredited centers reviewed blinded mammograms of 79 women, 45 of whom had breast cancer. The diagnosis would have been missed in 21%
<i>Burton et al (1998)(12)</i>	<b>Cancer detection</b>	Autopsy study at a single hospital: Of 250 malignancies, 111 were either missed or misdiagnosed, and in 57 cases the cause of death was cancer-related
<i>Burgess et al (1998)(13)</i>	<b>Breast cancer</b>	Of 132 patients with breast cancer, referral for definitive management was delayed in 32 (17%)
<i>Bouma et al (1999)(14)</i>	<b>Myocardial infarction</b>	Of 400 patients admitted with myocardial infarction, the diagnosis was delayed in female compared to male patients
<i>Meera et al (1999)(15)</i>	<b>Parkinson's disease</b>	Of 402 cases with presumed Parkinson's disease and on medications, only 53% had the disease on definitive evaluation
<i>Goddard et al (1999)(16)</i>	<b>Retinoblastoma</b>	Of 100 children with retinoblastoma, the diagnosis was delayed in 23%
<i>Van Schayak et al (2000)(17)</i>	<b>Asthma</b>	Of 1155 randomly-screened patients, spirometry suggested asthma in 86. Only 29 of these had ever complained of relevant symptoms to their physician, and of the diagnosis was missed in 6 (21%)
<i>Pope et al(2000)(18)</i>	<b>Myocardial infarction</b>	Of 10,689 patients seen for suspicious chest pain in 10 different ER's, 889 had acute myocardial infarction, of which 22 (2.3%) were mistakenly discharged from the ER
<i>Graff et al (2000)(19)</i>	<b>Appendicitis</b>	Retrospective study at 12 hospitals. Of 1,026 patients who had surgery for suspected appendicitis. 110 patients had no appendicitis at surgery; Of 916 patients with a diagnosis of appendicitis, the diagnosis was missed or wrong in 170 (18.6%)
<i>von Kodolitsch</i>	<b>Ruptured aortic</b>	In patients presenting with chest pain due to dissections of the proximal

et al (2000)(20)	<b>aneurysm</b>	aorta, the diagnosis was missed in 35%
Kantola et al (2001)(21)	<b>Tongue cancer</b>	Of 75 cases, referral to specialty care was delayed in 35%
Pidenda et al (2001)(22)	<b>Pulmonary embolism</b>	Review of fatal pulmonary embolism over a 5 year period at a single institution. Of 67 patients who died of pulmonary embolism, the diagnosis was not suspected clinically in 37 (55%)
Levack et al (2002)(23)	<b>Cancer-related spinal cord compression</b>	Of 319 patients, the median delay in diagnosis was 18 days
Edelman (2002)(24)	<b>Diabetes mellitus</b>	Retrospective review of 1,426 patients with laboratory evidence of diabetes mellitus (glucose > 200 mg/dl or HgbA1c > 7%): There was no mention of diabetes in the clinical record in 18% of these patients
Shojania (2002)(25)	<b>Pulmonary tuberculosis</b>	Review of autopsy studies that specifically focused on pulmonary TB: Approximately half of these diagnoses were not suspected antemortem
McGinnis et al (2002)(26)	<b>Melanoma</b>	Second review of 5,146 biopsies: The diagnosis changed in 11% (1.1% benign to malignant; 1.2% malignant to benign; 8% change in tumor grade)
Gorter et al (2002)(27)	<b>Psoriatic arthritis</b>	Patients with classical psoriatic arthritis visited 23 rheumatologists: The diagnosis was missed or wrong in 9 visits (39%)
Lopponen et al (2003)(28)	<b>Dementia</b>	All 1260 patients in a small town were screened for dementia. 9% screened positive, but only 4% had been previously diagnosed
Goyal et al (2004)(29)	<b>Bone cancer</b>	Of 103 patients with osteosarcoma or Ewing's sarcoma at two hospitals, delayed diagnosis was associated with being seen by a general practitioner (vs ER physician) and in patients under 12 years of age
Jiwa et al (2004)(30)	<b>Cancer</b>	Of 56 patients with cancer, significant delays in diagnosis were encountered in 8 cases
Vasudev et al (2004)(31)	<b>Testicular cancer</b>	Of 180 men with testicular cancer, referral to specialty care was delayed in 60%
Kowalski et al (2004)(32)	<b>Subarachnoid hemorrhage</b>	Of 482 patients with subarachnoid hemorrhage at a single hospital, the diagnosis was initially wrong in 56 (12%) and 22 of these patients suffered neurological complications before the diagnosis was confirmed
Yates et al (2004)(33)	<b>Iron deficiency anemia</b>	Of 431 patients with microcytic anemia, 57% had no GI investigations, and of 53 patients who ultimately died of GI cancer, 41 had had no investigation. Female sex, age < 65, and Hgb > 9 gm/dl were associated with non-investigation
Bogun et al (2004)(34)	<b>Atrial fibrillation</b>	Review of automated ECG interpretations: 35% of readings interpreted at atrial fibrillation were wrong and the error was detected by the reviewing cardiologist only 76% of the time
Buchewitz et al (2005)(35)	<b>Endometriosis</b>	Digital videotapes of laparoscopies were shown to 108 gynecologic surgeons: The interobserver agreement regarding the number of lesions was only 18%
Raab et al (2005)(36)	<b>Cancer pathology</b>	Reviewed pathology material from 4 hospitals over 1 year: The error rate was 2-9% for gynecologic cancer and 5-12% for non-gynecologic cancer
Perlis (2005)(37)	<b>Bipolar disorder</b>	The initial diagnosis was wrong in 69% of patients and delays in reaching the correct diagnosis were common
Edlow (2005)(38)	<b>Subarachnoid hemorrhage</b>	Review of published studies: Approximately 30% are misdiagnosed at presentation
Arnon et al (2006)(39)	<b>Infant botulism</b>	Of 129 infants with botulism over a 5 year period, only half the cases were suspected at the time of admission
Campos (2012)(40)	Alpha-1 Antitrypsin Deficiency	200,000 patients in the US have severe deficiency; Less than 10% have been properly diagnosed

Adapted in part from Schiff et al, 2005 (41) and Kostopolou et al, 2008 (42). Note: This list is a convenience sample of published articles, not a comprehensive listing.

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