

4

5



COLLEGE of AMERICAN
PATHOLOGISTS

Supplemental Digital Content* | Methodology | March 2015

Interpretive Diagnostic Error Reduction in Surgical Pathology and Cytology:

Joint Guideline from the College of American
Pathologists Pathology and Laboratory Quality
Center and the Association of Directors of
Anatomic and Surgical Pathology

Corresponding Author:
Raouf E. Nakhleh, MD

Authors :

Vania Nosé, MD, PhD
Lisa Fatheree, SCT(ASCP)
Christina Ventura, MT(ASCP)
Douglas McCrory, MD
Carol Colasacco SCT(ASCP), MLIS, AHIP

<http://www.archivesofpathology.org/doi/abs/10.5858/arpa.2014-0511-SA>

* The Supplemental Digital Content was not copyedited by *Archives of Pathology*

College of American Pathologists | 325 Waukegan Rd. | Northfield, IL 60093 | 800-323-4040 | cap.org

METHODS USED TO PRODUCE THE GUIDELINE

Panel Composition

The College of American Pathologists (CAP) Pathology and Laboratory Quality Center (the Center) and Association of Directors of Anatomic and Surgical Pathology (ADASP) convened an expert and advisory panel consisting of practicing pathologists with expertise in surgical pathology and cytology. CAP and ADASP approved the appointment of the project co-chairs (RN and VN) and panel members. These panel members and the methodologist served as the Expert Panel (EP) for the systematic evidence review.

Conflict of Interest (COI) Policy

Prior to acceptance on the expert or advisory panel, potential members completed the CAP conflict of interest (COI) disclosure process, whose policy and form (in effect April 2010) require disclosure of material financial interest in, or potential for benefit of significant value from, the guideline's development or its recommendations 12 months prior through the time of publication. The potential members completed the COI disclosure form, listing any relationship that could be interpreted as constituting an actual, potential, or apparent conflict. The CAP Center uses the following criteria:

Nominees who have the following conflicts may be excused from the panel:

- a. Stock or equity interest in a commercial entity that would likely be affected by the guideline or white paper
- b. Royalties or licensing fees from products that would likely be affected by the guideline or white paper
- c. Employee of a commercial entity that would likely be affected by the guideline or white paper

Nominees who have the following potentially manageable direct conflicts may be appointed to the panel:

- a. Patents for products covered by the guideline or white paper
- b. Member of an advisory board of a commercial entity that would be affected by the guideline or white paper
- c. Payments to cover costs of clinical trials, including travel expenses associated directly with the trial
- d. Reimbursement from commercial entity for travel to scientific or educational meetings

Everyone was required to disclose conflicts prior to beginning and continuously throughout the project's timeline. Expert panel members' disclosed conflicts are listed in the appendix of the manuscript. The CAP and ADASP provided funding for the administration of the project; no industry funds were used in the development of the guideline. All panel members volunteered their time and were not compensated for their involvement.



51 CAP/ADASP Expert Panel Literature Review and Analysis

52 The expert panel met 26 times through teleconference webinars from December 2011 through June 12,
53 2014. Additional work was completed via electronic mail and the panel met in person October 12, 2013 to
54 review evidence to date and draft recommendations.

55
56 All expert panelists participated in the systematic evidence review (SER) level of title-abstract, full-text
57 review, and data extraction. The co-chairs (RN and VN) and methodologist (DM) performed the audit of
58 data extraction. All articles were available as discussion or background references. All members of the
59 expert panel participated in developing draft recommendations, reviewing open comment feedback,
60 finalizing and approving recommendations and writing/editing of the manuscript.

61 62 Peer Review

63 An open comment period was held from December 2, 2013 through January 21, 2014. Five draft
64 recommendations and three methodology questions were posted online on the CAP Web site
65 www.cap.org. An announcement was sent to the following societies deemed to have interest:

66
67 CAP Board of Governors, Councils, Committees and Membership
68 Association of Directors of Anatomic and Surgical Pathology (ADASP)
69 American Society for Clinical Pathology (ASCP)
70 American Society of Cytopathology (ASC)
71 Papanicolaou Society of Cytology (PSC)
72 Arthur Purdy Stout Society (APSS)
73 Association of Pathology Chairs (APC)
74 Canadian Association of Pathologists (CAP-APC)
75 United States & Canadian Academy of Pathology (USCAP)
76 Quality Initiative in Interpretive Pathology (QIIP) Canadian Partnership Against Cancer
77 Society to Improve Diagnoses in Medicine (SIDM)
78 American Society of Clinical Oncologists (ASCO)
79 Veteran's Affairs (VA) and Department of Defense (DOD)
80 Centers for Disease Control and Prevention (CDC)
81 Centers for Medicare and Medicaid Services (CMS)

82
83 The website received 303 comments in total (Agree and Disagree responses were also captured). All the
84 recommendations achieved between 87% to 93% agreement. Pairs of expert panel members were
85 assigned 1 draft recommendation for which to review all comments received and provide an overall
86 summary to the rest of the panel. Following panel discussion, and the final quality of evidence
87 assessment, the panel members determined whether to maintain the original draft recommendation as is,
88 revise it with minor language change, or consider it as a major recommendation change. Resolution of all
89 changes was obtained by majority consensus of the panel using nominal group technique (rounds of
90 email discussion and multiple edited recommendations) amongst the panel members. The final
91 recommendations were approved by the expert panel with a formal vote. The panel considered the risks
92 and benefits throughout the whole process in their considered judgment process.¹ Formal cost analysis or
93 cost effectiveness was not performed.

94
95 An independent review panel (IRP) was assembled to review the guideline and recommend approval to
96 the CAP. The IRP was masked to the expert panel and vetted through the COI process. Final approval
97 was done by CAP Council on Scientific Affairs and ADASP Executive Board.

98 99 Assessing the Strength of Recommendations

100 Strength of recommendation is determined by the balance between desirable and undesirable
101 consequences of alternative management strategies, quality of evidence, variability in values and
102 preferences, resource use.²

103



- 104 The central question that the panel addressed in developing the guideline was “*What are the most*
105 *effective ways to reduce interpretive diagnostic errors in Anatomic Pathology?*”
- 106 Development of recommendations requires that the panel review the identified evidence and make a
107 series of key judgments:
- 108 1) What are the significant findings related to each KQ or outcome?
- 109 2) What is the overall quality of evidence supporting each KQ or outcome? Quality of evidence was
110 assessed according to the GRADE framework as described below.² Summary of Findings tables or when
111 sufficient information was not available, an alternative short evidence table format, were prepared for
112 each question. To maintain consistency with previous CAP guideline language, quality of evidence is
113 described as Convincing, Adequate or Inadequate as shown in Table 1.
- 114 3) What is the net balance of benefits and harms? The consideration of net balance of benefits and harms
115 will focus on the laboratory redundancy, efficiency and feasibility in comparison to the reduction of errors
116 or potential errors and their impact on patient outcomes.
- 117 4) What is the strength of each recommendation? The implications of a strong recommendation for
118 clinicians is that most patients should receive the recommended course of action; while the implications of
119 a weak recommendation are that different choices may be appropriate for different patients or that
120 different management options may be preferred by different stakeholders. We used the current CAP
121 designations for strength of recommendations of *strong recommendation, recommendation and expert*
122 *consensus opinion* (Table 2), as determined by a considered judgment process in which the guideline
123 panel weighs the quality of evidence, balance of benefits and harms, variability in values preference and
124 data on costs or resource use.¹

125 Dissemination Plans

126 CAP will host an Interpretive Diagnostic Error Reduction Through Targeted Case Reviews In Surgical
127 Pathology And Cytology Resource web page which will include a link to the [manuscript](#) and supplemental
128 digital content; summary of recommendations, teaching PowerPoint, and a frequently asked question
129 (FAQ) document. The ADASP webpage will include a link to the CAP guideline resource page. The
130 guideline will be promoted and presented at various professional society meetings including the College
131 of American Pathologists, the United States and Canadian Academy of Pathology (USCAP), and the
132 American Society of Clinical Pathology (ASCP).

134 SYSTEMATIC EVIDENCE REVIEW (SER)

136 The objectives of the SER were to investigate the most effective ways to reduce interpretive diagnostic
137 errors in surgical pathology and cytology. If of sufficient quality, findings from this review could provide an
138 evidence base to support development of the laboratory practice guideline. The scope of the SER and the
139 key questions (KQs) were established by the EP in consultation with a methodologist.

141 Key Questions:

- 142 1. Does targeted review (either done at analytic or post-analytic phase) of surgical pathology or cytology
143 cases (slides and/or reports) reduce the error rate (often measured as amended reports) or increase the
144 rate of interpretive error detection compared to no review, random review or usual review procedures?
- 145 2. What methods of selecting cases for review have been shown to increase the rate of interpretive error
146 detection compared to no review, random review or usual review procedures?

148 Detailed Scope Questions:

- 149 • Can a targeted review of cases lead to increased detection of errors?
- 150 • Is there a particular method (eg, prospective vs. retrospective, random reviews, etc.) that results in
151 lower error rates? Or amended report rates? If yes, what is the method?
- 152 • Is there a particular type of diagnosis(es) (eg, malignant, benign, borderline) more prone to error? If
153 yes, what?



- 154 • Is there a particular organ/system associated with a higher rate of error/disagreements revealed as
 155 problematic in the literature (eg, thyroid FNA, lymphoma, brain biopsy)? If yes, what?
 156 • How does the error rate for multi-organ reviews compare with single organ reviews? (eg, internal
 157 and/or external)
 158 • Do blinded reviews find more or less errors than non-blinded reviews?
 159 • Can we learn anything from external review studies that can be applied to internal reviews (or vice-
 160 versa)?
 161 • Can we establish an external error rate (benchmark) for surgical pathology or cytology?
 162 • Can we establish an internal error rate (benchmark) for surgical pathology or cytology?
 163 • What are the costs of conducting internal or external reviews?
 164 • What is the effect on turn-around times (efficacy) for internal or external reviews?
 165 • What is the effect on patient care when conducting internal or external reviews?
 166 • Has there been improvement in reducing error reduction over time?
 167 • What other factors (eg, standard criteria, clinical correlation, ancillary testing) are emphasized most
 168 frequently and with which organ system or diagnosis?

169 Search and Selection

170 The systematic literature search for relevant studies included a search of MEDLINE using the Ovid SP
 171 interface on November 12, 2013, with the date parameters of January 1992 through October 2012.
 172 Medical subject headings (MeSH) for the concept *pathology* (eg, Pathology, Surgical/ Pathology, Clinical;
 173 Pathology) were combined with MeSH terms for the concept *quality* (eg, Quality Assurance, Health Care;
 174 Quality Control; Quality Improvement; Reproducibility of Results; Diagnostic Errors). MeSH terms were
 175 supplemented with keywords (eg, histopathology, cytopathology, histology, or cytology; and second
 176 opinion, misinterpretation, or interpretation errors). A targeted concept of slide/case review included
 177 keywords such as targeted review, peer review, or random review and the keywords slide, case, or report.
 178 Limits were set for human studies published in English. The search was not limited by study design in
 179 order to capture editorials, letters, or commentaries that might be relevant and useful for discussion
 180 purposes. A literature refresh of the OvidSP search strategy was run on November 21, 2013, to identify
 181 relevant studies published since October 2012. The full Ovid search strategy is included in the appendix.
 182

183 The Ovid search strategy was modified for PubMed (1/1/92-12/31/12), and Google Scholar (1/1/12 –
 184 1/26/13). In addition, a handsearch of relevant journals (*American Journal of Clinical Pathology, American*
 185 *Journal of Surgical Pathology, Archives in Pathology and Laboratory Medicine, Cancer, Cancer*
 186 *Cytopathology, Diagnostic Cytopathology, Histopathology, Modern Pathology*) was completed for issues
 187 published from January 2008, through December 2012. A search for meeting abstracts was completed
 188 utilizing Biosis Previews (Web of Science) (1/1/2008-12/31/2012) and by handsearching published
 189 abstracts from relevant meetings (American Society of Cytopathology, American Society for Clinical
 190 Pathology, British Society for Clinical Cytology, College of American Pathologists, European Congress of
 191 Cytopathology, International Academy of Pathology, United States and Canadian Academy of Pathology)
 192 held from January 2008 through December 2012. Reference lists from included articles were scanned for
 193 additional relevant studies.
 194

195 Two reviewers were used at all levels of review (eg, title/abstract, full article) and for data/information
 196 extraction. Conflicts were resolved by discussion or referral to the panel co-chairs for a decision. When
 197 article abstracts or document summaries were not available or a conflict was not resolved, full articles
 198 were reviewed.
 199

200 Selection at all levels of the review was based on predetermined inclusion/exclusion criteria.
 201

202 Inclusion criteria:

- 203 • Surgical pathology or cytology studies
 204 • Original research addressing targeted review
 205 • English language articles
 206 • All study types were initially included
 207



208
 209
 210
 212
 213
 215
 216
 217
 218
 219
 220
 221
 222
 223
 224
 225
 226
 227
 228
 229
 230
 231
 232
 233
 234
 235
 236
 237
 238
 239
 240
 241
 242
 243
 244
 245
 246
 247
 248
 249
 250
 251
 252
 253
 254
 255
 256
 257
 258
 259
 260
 261

Exclusion criteria:

- Clinical pathology studies
- Additional diagnostic techniques, issues related to competency, use of checklists, standardized language, taxonomy or formatting
- Studies focused on pre-analytic specimen processes
- Post-analytic typographic errors
- Non-English studies
- Animal studies

Outcomes of Interest

We are interested in identifying discrepancies in interpretation between a primary pathologist review and a second pathologist review as a way of estimating the error rate. To the extent that erroneous readings can be identified in excess of an expected degree of disagreement, then a method of targeted review would be said to be effective. Thus, studies with a control group are desirable; as a practical matter, however, it is necessary to examine uncontrolled series, too. Studies had to report numbers of discrepant diagnoses among a defined population of specimens examined to allow calculation of a discrepancy rate.

We are not interested in discrepancies from the pre-analytic specimen process (ie, related to tissue collection and processing) or post-analytic errors (eg, typographic or transcription errors, amended reports), additional diagnostic techniques (eg, immunomarkers), issues related to competency, or the use of checklists, standardized language, taxonomy or formatting.

Various studies classify errors in different ways (eg, major versus minor). Recognizing that all errors are not alike, we assessed the severity of interpretive errors according to the clinical impact on a patient.³ We considered the clinical impact of errors as follows: 1) diagnostic thinking (error results in a change in diagnosis or diagnostic category); 2) therapeutic efficacy (error results in a change in therapeutic choice); or 3) patient outcome efficacy (error results in a change in outcome (eg, procedure avoided); demonstrating this unequivocally may require long-term follow-up). We also considered the efficiency or cost (in terms of effort or dollars) that a targeted review strategy entails.

Data Extraction and Management

The data elements from an included article/document were extracted by one reviewer into standard data formats and tables developed using systematic review database software (DistillerSR, Evidence Partners Inc., Ottawa, Canada); a second reviewer confirmed accuracy and completeness. In all cases, the methodologist acted as either the primary or secondary reviewer. Any discrepancies in data extraction were resolved by discussion with the methodologist. A bibliographic database was established in EndNote (Thomson Reuters, Carlsbad, CA) to track all literature identified and reviewed during the study.

Environmental Scan

An environmental scan for established guidelines was performed including a targeted search of pathology organizations' web sites and a search of guideline clearinghouse websites (TRIP Database, Guidelines International Network, Agency for Healthcare Research and Quality) using the search terms "pathology or laboratory" and "guidelines or regulations."

Quality Assessment

Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach provides a system for rating quality of evidence and strength of recommendations that is explicit, comprehensive, transparent, and pragmatic and is increasingly being adopted by organizations worldwide.² The GRADE approach examines the quality of evidence as the level of individual studies and also at the review level. GRADE was used for rating the quality of evidence.

At the individual study level, we assessed studies according to three criteria: 1) study design rating; 2) risk of bias rating; and 3) applicability concerns. Study design was assessed according to the following hierarchy:



- 262 1) noncomparative studies (where a single method of targeted review is considered, in determining a
263 single error rate)
264 2) comparative studies (where two or more methods of targeted review are independently applied and
265 compared in one group of patients)
266 3) comparative studies (where two or more methods of targeted review are compared in different groups
267 of patients
268 a. historical (eg, a before-after series)
269 b. concurrent (eg, comparison of two different institution's targeted review programs)
270 c. quasi-randomized (concurrent, similar or identical sites, allocated with attempt to reduce bias
271 beyond that provided by 3.a. or 3.b.)
272 d. randomized

273
274 Risk of bias ratings were based on the following three criteria, rated as yes, no, or unclear:

- 275 1) Were review diagnoses independent, blinded from the primary diagnosis?
276 2) Was case selection done using explicit, objective and reproducible criteria?
277 3) Was interpretive error ascertained without confounding by pre- or post-analytic error?
278

279 Applicability concerns were assessed for three areas, rated as yes, no, or unclear:

- 280 1) Is study limited to particular organ systems, specimen types or diagnoses?
281 2) Is the inter-rater performance examined and acceptable?
282 3) Is there a distinction between major and minor errors based on impact on patient treatment or
283 outcome?
284

285 At the review level, we examined the collection of studies according to the following domains:

- 286 1) Consistency
287 2) Directness
288 3) Precision
289 4) Magnitude of effect
290 5) Number of studies/patients

291 We developed a GRADE evidence table, and mapped the quality of evidence ratings to the CAP rating
292 scheme according to table 1.
293

294 Data Analysis

295 Rates of discrepancy and major discrepancy were described for all studies, and subgroups based on type
296 of specimen (surgical pathology, cytopathology or both), focusing on one organ or organ system (single-
297 organ) versus multi-organ studies, and whether conducted within a single institution (internal) or reviews
298 of cases diagnosed at a different institution (external). Studies were tested for homogeneity using
299 Comprehensive Meta Analysis version 2.2.064. Nonparametric descriptive statistics including median,
300 and 1st and 3rd quartiles were calculated using Excel.
301

302 Results

303 Among the 823 citations identified by electronic and hand searches, 141 were selected for inclusion.
304 These included 130 published peer-reviewed articles, and 11 grey literature documents (Appendix 1).
305 Among the extracted documents, 4 articles/documents did not meet minimum quality standards,
306 presented incomplete data or data that were not in useable formats, or included only information based
307 on expert opinion. These articles were not included in analyses or narrative summaries.
308

309 Of 137 studies included, 128 (91.4%) were single arm clinical case series. Of 12 comparative studies, 2
310 reported a comparison of 2 or more methods of targeted review in one group of patients; 6 compared
311 targeted review methods in different historical cohorts (eg a before-after study), and 4 compare 2 or more
312 targeted review methods in concurrent groups of patients. No studies used random allocation or other
313 robust measures to control for potential bias.



314
315 Risk of bias assessments of included studies showed that review diagnoses were made independently
316 (blinded from the primary diagnosis) in 30 (21.4%) studies, unclear in 2 (1.4%) and without blinding in 108
317 (77.1%). Case selection was done using explicit, objective and reproducible criteria in 56 (40%), and with
318 unclear, subjective, non-reproducible or without criteria in 84 (59.3%). Interpretive error was ascertained
319 without confounding by pre- or post-analytic error in 120 (85.7%), unclear in 2 (1.4%) and not in 18
320 (12.9%) studies.

321
322 Applicability concerns were as follows: Studies were limited to particular organ systems, specimen types
323 or diagnoses in 111 (79.3%) cases, and not limited in 29 (20.7%) cases. Inter-rater performance was
324 examined and acceptable in 29 (20.7%), unclear in 6 (4.3%), and not in 105 (75%) of studies. There was
325 a distinction between major and minor errors based on impact on patient treatment or outcome in 78
326 (55.7%), unclear in 2 (1.4%), and not in 60 (42.9%) of studies.

327
328 Discrepancy rates and major discrepancy rates by study characteristics are provided in the manuscript.

329
330 The distribution of discrepancy rates shows that the variability was greater for studies of smaller sample
331 size. The distribution of discrepancy rates by sample size is shown in figures 1 and 2, both of which show
332 high variability in discrepancy rates for studies with smaller sample sizes, with larger studies tending to
333 have lower discrepancy rates.

334
335 Evidence on each question was summarized in terms of study quality and effects in Evidence Profile
336 Tables (Tables 3-5) which were used as the basis for quality of evidence determinations with the panel.

337
338
339
340
341



Table 1. Quality of Evidence Ratings: mapping from GRADE categories to CAP

GRADE	Definition	CAP	Definition
High	Further research is very unlikely to change our confidence in the estimate of effect.	High (Convincing)	High confidence that available evidence reflects true effect
Moderate	Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.	Intermediate (Adequate)	Moderate confidence that available evidence reflects true effect
Low	Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.	Low (Inadequate)	Little confidence that available evidence reflects true effect
Very low	Any estimate of effect is very uncertain.	Insufficient	Evidence is insufficient to discern net effect

Adapted from Guyatt et al² with permission from BMJ Publishing Group Ltd..

342
343

Table 2. Strength of Recommendations

CAP Designation	Recommendation	Rationale
Strong Recommendation	Recommend For or Against a particular pathology review practice (Can include must or should)	Supported by high (convincing) or intermediate (adequate) quality of evidence and clear benefit that outweighs any harms
Recommendation	Recommend For or Against a particular pathology review practice (Can include should or may)	Some limitations in quality of evidence (intermediate [adequate] or low [inadequate]), balance of benefits and harms, values, or costs but panel concludes that there is sufficient evidence to inform a recommendation.
Expert Consensus Opinion	Recommend For or Against a particular pathology review practice (Can include should or may)	Serious limitations in quality of evidence (low [inadequate] or insufficient), balance of benefits and harms, values or costs, but panel consensus is that a guideline is necessary.
No Recommendation	No recommendation for or against a particular pathology review practice	Insufficient evidence, confidence, or agreement to provide a recommendation.

Adapted from Teutsch et al⁴ with permission from Macmillian Publishers Ltd. Modified by the CAP Pathology and Laboratory Quality Center.

344
345

Table 3: Evidence profile for Recommendation 1: Should case review be used for diagnostic evaluation of pathology materials?*

Quality assessment							No. of patients		Effect		Quality	Importance
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Case review	Control	Relative (95% Confidence Interval)	Absolute		
Discrepancy Rate (follow-up 1 years; assessed with case review)												
122	observational studies ^a	no serious risk of bias	no serious inconsistency ^b	no serious indirectness ^c	no serious imprecision	none	27178/513268 (5.3%)	-	-	-	⊕⊕O LOW	IMPORTANT
Major Discrepancy Rate (assessed with: case review)												
79	observational studies ^a	no serious risk of bias	no serious inconsistency ^b	no serious indirectness ^d	no serious imprecision	none	4416/179130 (2.5%)	-	-	-	⊕⊕O LOW	CRITICAL
Inter-observer agreement												
27	observational studies	no serious risk of bias	Serious ^d	Serious ^e	no serious imprecision	reporting bias	-	-	kappa ranged from 0.16 to 0.97	-	⊕OO VERY LOW	IMPORTANT

*Settings: Pathology laboratories and practices

^a Studies include single- and multi-institution case series that quantify diagnostic discrepancies or errors according to various definitions

^b Although the magnitude of discrepancy rates vary considerably from study to study based on sample size, specimen type, single versus multi-organ, review type (internal versus external), and definition of discrepancy, studies are remarkably consistent in finding non-trivial discrepancy rates.

^c Few studies reported the impact of discrepant diagnoses on treatment choice, and even fewer on patient outcome.

^d Inter-rater reliability studies had high variability. Experts agreed better than non-experts.

^e Inter-rater agreement studies usually used highly selected samples and small numbers of observations

346
347
348
349
350
351
352
353
354
355
356

357

Table 4: Evidence profile for Recommendation 2: Should prospective review versus retrospective review be used for diagnostic evaluation of pathology materials?

Quality assessment							No. of patients		Effect		Quality	Importance	
No of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Prospective review	Retrospective review	Relative (95% CI)	Absolute			
Discrepancy Rate													
4	observational studies	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	510/15515 (3.3%)	0.13%	RD ranged from -2.4 to 1.1	-	⊕⊕○○	IMPORTANT	
								3.4%		-			LOW
								13%		-			
Major discrepancy													
2	observational studies	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	32/6129 (0.52%)	0.1%	RD ranged from -0.1 to -0.5	-	⊕⊕○○	IMPORTANT	
								1.7%		-			LOW

358 CI – confidence interval; RD – risk difference

359

360

Downloaded from <http://meridian.allenpress.com/doi/pdf/10.5858/arpa.2014-0511-SA> by guest on 22 October 2020



Table 5: Evidence Profile for Recommendation 3: Should targeted case review versus random case review be used for diagnostic evaluation of pathology materials?

Quality assessment							No. of patients		Effect		Quality	Importance
No. of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Targeted case review	Random case review	Relative (95% CI)	Absolute		
Discrepancy rate												
1	observational studies	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	50/380 (13.2%)	195/7444 (2.6%)	OR 5.6 (4.1 to 7.8)	105 more per 1000 (from 73 more to 147 more)	⊕⊕○○ LOW	
Major discrepancy												
1	observational studies	no serious risk of bias	no serious inconsistency	no serious indirectness	no serious imprecision	none	12/380 (3.2%)	27/7444 (0.36%)	OR 9.0 (4.5 to 18)	28 more per 1000 (from 12 more to 58 more)	⊕⊕○○ LOW	

361 CI – confidence interval; OR – Odds ratio

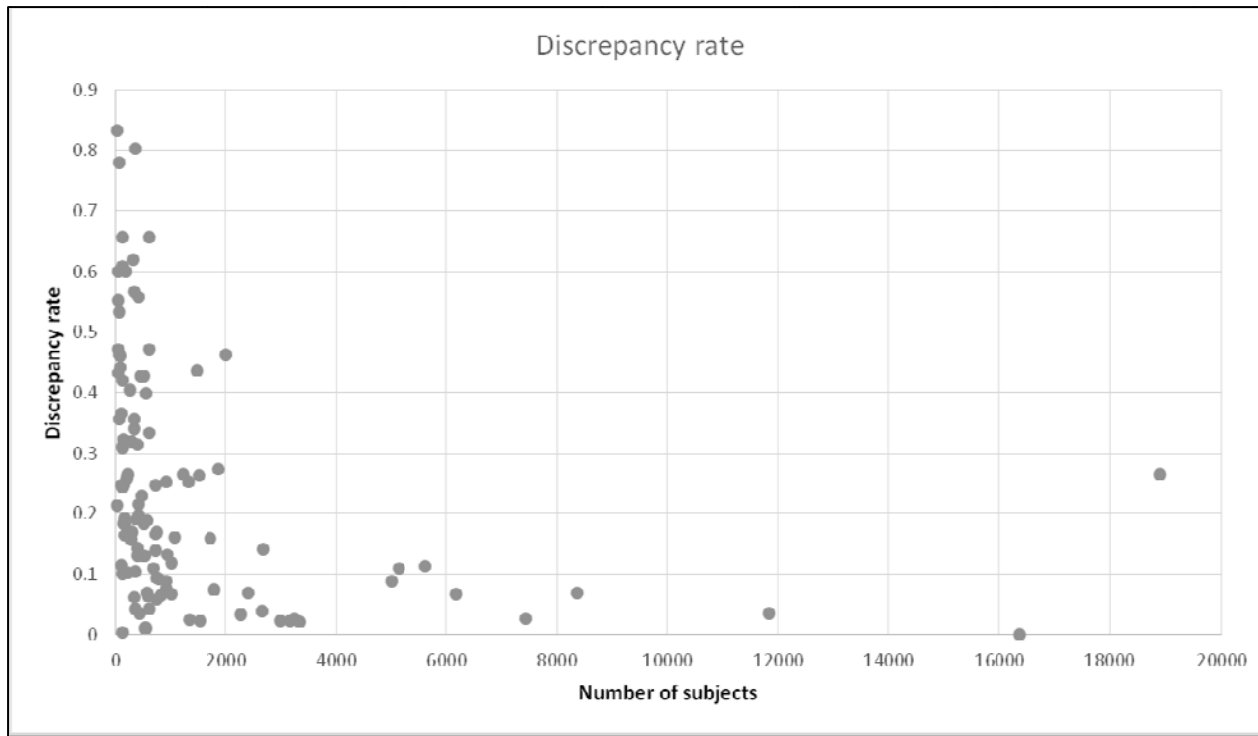
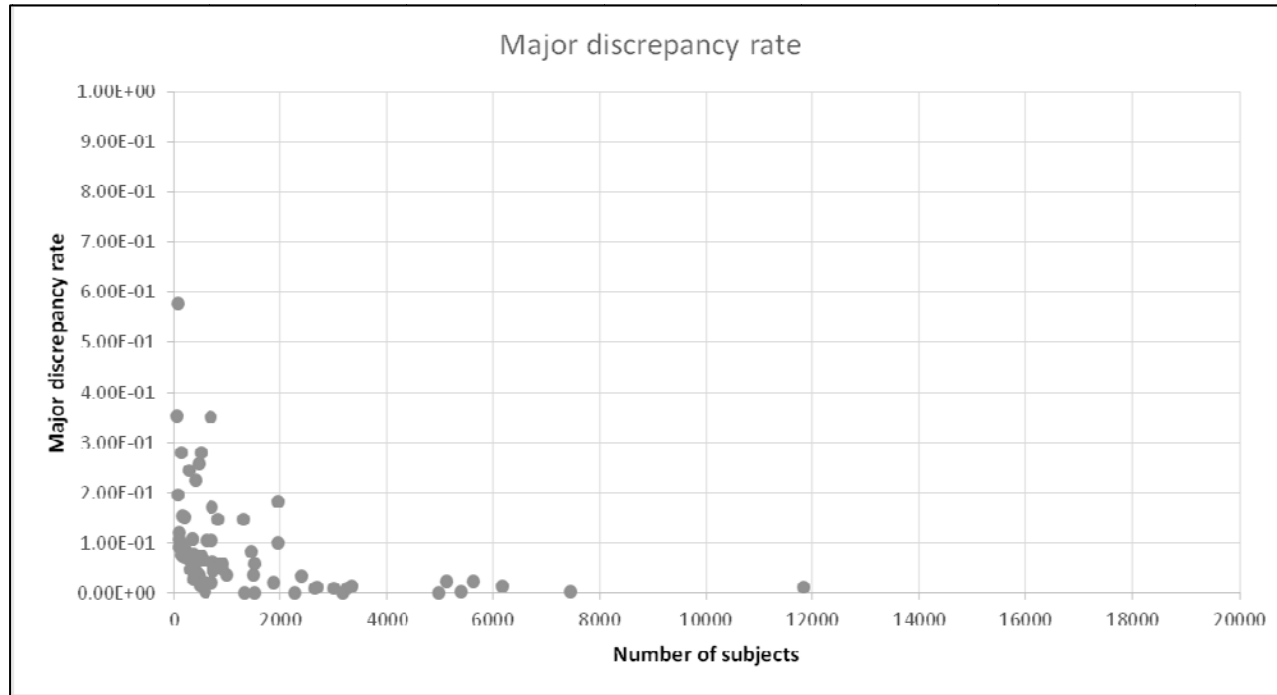


Figure 1: Discrepancy rate distribution by sample size⁵⁻¹¹²

362
363
364
365
366
367
368
369
370

371
372



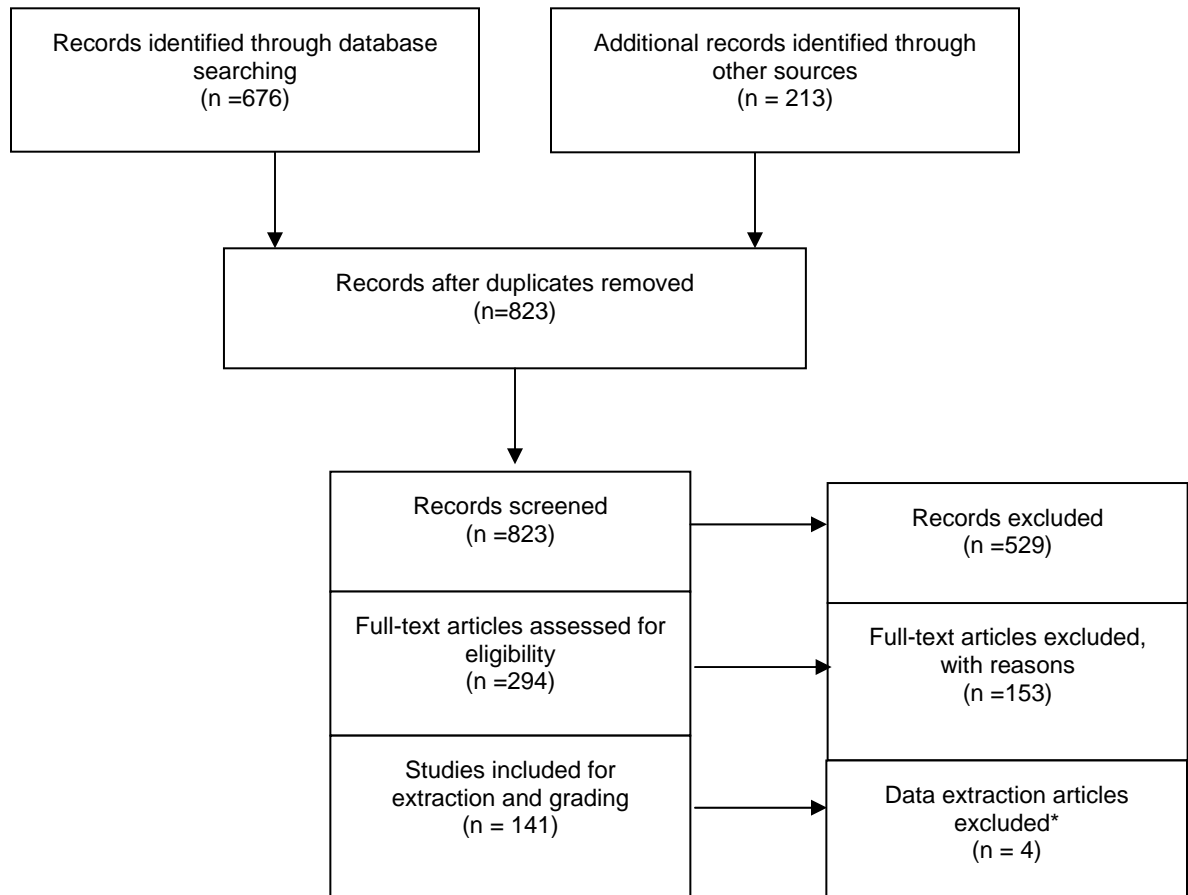
373
374
375
376
377
378

Figure 2: Major discrepancy rate distribution by sample size^{5, 6, 8, 10, 11, 13-15, 17-20, 22-24, 31, 33, 36, 40, 42-44, 46, 50, 51, 53-55, 58-61, 64, 67-69, 74, 76, 78, 83, 84, 86-88, 91, 93, 94, 96, 97, 99-101, 106, 108, 110, 113-129}



379 **APPENDIX**

380
381 Appendix A: Literature Review Results
382 Adapted with permission from Moher et al.¹³⁰
383



417 *Excluded based on expert opinion, did not meet minimum quality standards, presented incomplete data or
418 data that were not in useable formats
419

420 Appendix B: Ovid MEDLINE Search Strategy

- 421
422 1. *Pathology/
423 2. *Pathology, Surgical/
424 3. Pathology, Clinical/
425 4. Pathology department, Hospital/
426 5. Cytodiagnosis/
427 6. Biopsy/
428 7. *"Diagnostic Techniques and Procedures"/
429 8. (pathology\$ or cytolog\$ or histolog\$ or histopatholog\$ or cytopatholog\$).tw.
430 9. or/1-8
431

- 432 AND
 433
 434 1. *Medical Errors/
 435 2. *"Referral and Consultation"/
 436 3. *Quality Assurance, Health Care/
 437 4. *Quality Control/
 438 5. exp *Diagnostic Errors/
 439 6. *Quality Improvement/
 440 7. exp *"Peer Review"/
 441 8. *"Root Cause Analysis"/
 442 9. *Total Quality Management/
 443 10. *"Reproducibility of Results"/
 444 11. *"Sensitivity and Specificity"/
 445 12. *Medical Audit/
 446 13. *"Insurance Claim Review"/
 447 14. *Malpractice/lj [Legislation & Jurisprudence]
 448 15. performance improvement.tw.
 449 16. ((corrected or amended) adj2 report\$.tw.
 450 17. (misinterpretation or misdiagnosis or medicolegal or "patient safety" or "second opinion").tw.
 451 18. ((interpret\$ or diagnostic) adj2 error\$.tw.
 452 19. (error adj2 (reduction or prevention or rate)).tw.
 453 20. "diagnostic disagreement\$.tw.
 454 21. (quality adj3 (improvement or control or assurance or practice\$ or measure\$ or process\$)).tw.
 455 22. or/1-21
 456
 457 AND
 458
 459 1. ((routine or target\$ or random\$ or blind\$ or peer\$) adj4 review\$.tw.
 460 2. ((case\$ or slide\$ or report\$) adj4 review\$.tw.
 461 3. or/1-2
 462
 463 Limits: Humans/English/1/1/1992-10/31/2012. Rerun on 11/21/2013 to include 10/31/2012-11/21/2013.
 464
 465
 466
 467



REFERENCES

1. Scottish Intercollegiate Guidelines Network (SIGN). SIGN 50: a guideline developer's handbook. Edinburgh: SIGN; 2014. (SIGN publication no. 50). [October 2014]. Available from URL: <http://www.sign.ac.uk>.
2. Guyatt GH, Oxman AD, Vist GE, et al. GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. *BMJ*. 2008;336(7650):924-926.
3. Fryback DG, Thornbury JR. The efficacy of diagnostic imaging. *Med Decis Making*. 1991;11(2):88-94.
4. Teutsch SM, Bradley LA, Palomaki GE, et al. The evaluation of genomic applications in practice and prevention (EGAPP) initiative: methods of the EGAPP Working Group. *Genet Med*. 2009;11(1):3-14.
5. Abt AB, Abt LG, Olt GJ. The effect of interinstitution anatomic pathology consultation on patient care. *Arch Pathol Lab Med*. 1995;119(6):514-517.
6. Agarwal S, Wadhwa N. Revisiting old slides--how worthwhile is it? *Pathol Res Pract*. 2010;206(6):368-371.
7. Ahmed Z, Yaqoob N, Muzaffar S, Kayani N, Pervez S, Hasan SH. Diagnostic surgical pathology: the importance of second opinion in a developing country. *J Pak Med Assoc*. 2004;54(6):306-311.
8. Aldape K, Simmons ML, Davis RL, et al. Discrepancies in diagnoses of neuroepithelial neoplasms: the San Francisco Bay Area Adult Glioma Study. *Cancer*. 2000;88(10):2342-2349.
9. Allsbrook WC Jr, Mangold KA, Johnson MH, Lane RB, Lane CG, Epstein JI. Interobserver reproducibility of Gleason grading of prostatic carcinoma: general pathologist. *Hum Pathol*. 2001;32(1):81-88.
10. Al-Maghrabi JA, Sayadi HH. The importance of second opinion in surgical pathology referral material of lymphoma. *Saudi Med J*. 2012;33(4):399-405.
11. Arbiser ZK, Folpe AL, Weiss SW. Consultative (expert) second opinions in soft tissue pathology. Analysis of problem-prone diagnostic situations. *Am J Clin Pathol*. 2001;116(4):473-476.
12. Ascoli V, Bosco D, Carnovale Scalzo C. Cytologic re-evaluation of negative effusions from patients with malignant mesothelioma. *Pathologica*. 2011;103(6):318-324.
13. Bajaj J, Morgenstern N, Sugrue C, Wasserman J, Wasserman P. Clinical impact of second opinion in thyroid fine needle aspiration cytology (FNAC): a study of 922 interinstitutional consultations. *Diagn Cytopathol*. 2012;40(5):422-429.
14. Baloch ZW, Hendreen S, Gupta PK, et al. Interinstitutional review of thyroid fine-needle aspirations: impact on clinical management of thyroid nodules. *Diagn Cytopathol*. 2001;25(4):231-234.
15. Bejarano PA, Koehler A, Sherman KE. Second opinion pathology in liver biopsy interpretation. *Am J Gastroenterol*. 2001;96(11):3158-3164.
16. Berney DM, Fisher G, Kattan MW, et al. Pitfalls in the diagnosis of prostatic cancer: retrospective review of 1791 cases with clinical outcome. *Histopathology*. 2007;51(4):452-457.
17. Boiko PE, Piepkorn MW. Reliability of skin biopsy pathology. *J Am Board Fam Pract*. 1994;7(5):371-374.
18. Bomeisl PE Jr, Alam S, Wakely PE Jr. Interinstitutional consultation in fine-needle aspiration cytopathology: a study of 742 cases. *Cancer*. 2009;117(4):237-246.
19. Bruner JM, Inouye L, Fuller GN, Langford LA. Diagnostic discrepancies and their clinical impact in a neuropathology referral practice. *Cancer*. 1997;79(4):796-803.
20. Butler ST, Youker SR, Mandrell J, Flanagan KH, Fosko SW. The importance of reviewing pathology specimens before Mohs surgery. *Dermatol Surg*. 2009;35(3):407-412.
21. Castanon A, Ferryman S, Patnick J, Sasieni P. Review of cytology and histopathology as part of the NHS Cervical Screening Programme audit of invasive cervical cancers. *Cytopathology*. 2012;23(1):13-22.
22. Chafe S, Honore L, Pearcey R, Capstick V. An analysis of the impact of pathology review in gynecologic cancer. *Int J Radiat Oncol Biol Phys*. 2000;48(5):1433-1438.
23. Chan YM, Cheung AN, Cheng DK, Ng TY, Ngan HY, Wong LC. Pathology slide review in gynecologic oncology: routine or selective? *Gynecol Oncol*. 1999;75(2):267-271.
24. Clary KM, Silverman JF, Liu Y, et al. Cytohistologic discrepancies: a means to improve pathology practice and patient outcomes. *Am J Clin Pathol*. 2002;117(4):567-573.

- 522 25. Coblentz TR, Mills SE, Theodorescu D. Impact of second opinion pathology in the definitive
523 management of patients with bladder carcinoma. *Cancer*. 2001;91(7):1284-1290.
- 524 26. Coffin CS, Burak KW, Hart J, Gao ZH. The impact of pathologist experience on liver transplant biopsy
525 interpretation. *Mod Pathol*. 2006;19(6):832-838.
- 526 27. Cook IS, McCormick D, Poller DN. Referrals for second opinion in surgical pathology: implications for
527 management of cancer patients in the UK. *Eur J Surg Oncol*. 2001;27(6):589-594.
- 528 28. Corley DA, Kubo A, DeBoer J, Rumore GJ. Diagnosing Barrett's esophagus: reliability of clinical and
529 pathologic diagnoses. *Gastrointest Endosc*. 2009;69(6):1004-1010.
- 530 29. Davidov T, Trooskin SZ, Shanker BA, et al. Routine second-opinion cytopathology review of thyroid
531 fine needle aspiration biopsies reduces diagnostic thyroidectomy. *Surgery*. 2010;148(6):1294-1301.
- 532 30. Epstein JI, Walsh PC, Sanfilippo F. Clinical and cost impact of second-opinion pathology. Review of
533 prostate biopsies prior to radical prostatectomy. *Am J Surg Pathol*. 1996;20(7):851-857.
- 534 31. Eskander RN, Baruah J, Nayak R, et al. Outside slide review in gynecologic oncology: impact on
535 patient care and treatment. *Int J Gynecol Pathol*. 2013;32(3):293-298.
- 536 32. Fraser S, Lanaspri E, Pinto T, Goderya R, Chandra A. Thyroid FNA diagnosis: correlation between
537 referral and review diagnosis in a network MDM. *Cytopathology*. 2011;22(4):ii.
- 538 33. Gaudi S, Zarandona JM, Raab SS, English JCI, Jukic DM. Discrepancies in dermatopathology
539 diagnoses: the role of second review policies and dermatopathology fellowship training. *J Am Acad*
540 *Dermatol*. 2013;68(1):119-128.
- 541 34. Gerhard R, da Cunha Santos G. Inter- and intraobserver reproducibility of thyroid fine needle
542 aspiration cytology: an analysis of discrepant cases. *Cytopathology*. 2007;18(2):105-111.
- 543 35. Golfier F, Clerc J, Hajri T, et al. Contribution of referent pathologists to the quality of trophoblastic
544 diseases diagnosis. *Hum Reprod*. 2011;26(10):2651-2657.
- 545 36. Hahm GK, Niemann TH, Lucas JG, Frankel WL. The value of second opinion in gastrointestinal and
546 liver pathology. *Arch Pathol Lab Med*. 2001;125(6):736-739.
- 547 37. Haws B, St Romain P, Mammen J, Fraga GR. Secondary review of external histopathology on
548 cutaneous oncology patients referred for sentinel lymph node biopsy: how often does it happen and is
549 it worth it? *J Cutan Pathol*. 2012;39(9):844-849.
- 550 38. Idowu MO, Jain R, Pedigo MA, Powers CN. Is a second pathologist's review of ASC-H useful in
551 reducing false negative diagnosis. *Mod Pathol*. 2008;21(suppl 1s):74A.
- 552 39. Jacques SM, Qureshi F, Munkarah A, Lawrence WD. Interinstitutional surgical pathology review in
553 gynecologic oncology: II. Endometrial cancer in hysterectomy specimens. *Int J Gynecol Pathol*.
554 1998;17(1):42-45.
- 555 40. Jara-Lazaro AR, Thike AA, Tan PH. Diagnostic issues in second opinion consultations in prostate
556 pathology. *Pathology*. 2010;42(1):6-14.
- 557 41. Jing X, Knoepp SM, Roh MH, et al. Group consensus review minimizes the diagnosis of "follicular
558 lesion of undetermined significance" and improves cytohistologic concordance. *Diagn Cytopathol*.
559 2012;40(12):1037-1042.
- 560 42. Jones K, Jordan RC. Patterns of second-opinion diagnosis in oral and maxillofacial pathology. *Oral*
561 *Surg Oral Med Oral Pathol Oral Radiol Endod*. 2010;109(6):865-869.
- 562 43. Kamat S, Parwani AV, Khalbuss WE, et al. Use of a laboratory information system driven tool for pre-
563 signout quality assurance of random cytopathology reports. *J Pathol Inform*. 2011;2:42.
- 564 44. Kennecke HF, Speers CH, Ennis CA, Gelmon K, Olivotto IA, Hayes M. Impact of routine pathology
565 review on treatment for node-negative breast cancer. *J Clin Oncol*. 2012;30(18):2227-2231.
- 566 45. Kishimoto R, Saika T, Bekku K, et al. The clinical impact of pathological review on selection the
567 treatment modality for localized prostate cancer in candidates for brachytherapy monotherapy. *World*
568 *J Urol*. 2012;30(3):375-378.
- 569 46. Kronz JD, Milord R, Wilentz R, Weir EG, Schreiner SR, Epstein JI. Lesions missed on prostate
570 biopsies in cases sent in for consultation. *Prostate*. 2003;54(4):310-314.
- 571 47. Kukreti V, Patterson B, Callum J, Etchells E, Crump M. Pathology the gold standard - a retrospective
572 analysis of discordant "second-opinion" lymphoma pathology and its impact on patient care. *Blood*.
573 2006;108(11):348.
- 574 48. Kuroiwa K, Shiraishi T, Naito S. Discrepancy between local and central pathological review for radical
575 prostatectomy specimens. *J Urol*. 2009;181(4, suppl):58.

- 576 49. Kwon JS, Francis JA, Qiu F, Weir MM, Ettler HC. When is a pathology review indicated in
577 endometrial cancer? *Obstet Gynecol.* 2007;110(6):1224-1230.
- 578 50. LaCasce AS, Kho ME, Friedberg JW, et al. Comparison of referring and final pathology for patients
579 with non-Hodgkin's lymphoma in the National Comprehensive Cancer Network. *J Clin Oncol.*
580 2008;26(31):5107-5112.
- 581 51. Layfield LJ, Jones C, Rowe L, Gopez EV. Institutional review of outside cytology materials: a
582 retrospective analysis of two institutions' experiences. *Diagn Cytopathol.* 2002;26(1):45-48.
- 583 52. Lehnhardt M, Daigeler A, Hauser J, et al. The value of expert second opinion in diagnosis of soft
584 tissue sarcomas. *J Surg Oncol.* 2008;97(1):40-43.
- 585 53. Lester JF, Dojcinov SD, Attanoos RL, et al. The clinical impact of expert pathological review on
586 lymphoma management: a regional experience. *Br J Haematol.* 2003;123(3):463-468.
- 587 54. Lind AC, Bewtra C, Healy JC, Sims KL. Prospective peer review in surgical pathology. *Am J Clin*
588 *Pathol.* 1995;104(5):560-566.
- 589 55. Lueck N, Jensen C, Cohen MB, Weydert JA. Mandatory second opinion in cytopathology. *Cancer.*
590 2009;117(2):82-91.
- 591 56. Lurkin A, Ducimetiere F, Vince DR, et al. Epidemiological evaluation of concordance between initial
592 diagnosis and central pathology review in a comprehensive and prospective series of sarcoma
593 patients in the Rhone-Alpes region. *BMC Cancer.* 2010;10:150.
- 594 57. Lytwyn A, Salit IE, Raboud J, et al. Interobserver agreement in the interpretation of anal intraepithelial
595 neoplasia. *Cancer.* 2005;103(7):1447-1456.
- 596 58. Manion E, Cohen MB, Weydert J. Mandatory second opinion in surgical pathology referral material:
597 clinical consequences of major disagreements. *Am J Surg Pathol.* 2008;32(5):732-737.
- 598 59. Matasar MJ, Shi W, Silberstien J, et al. Expert second-opinion pathology review of lymphoma in the
599 era of the World Health Organization classification. *Ann Oncol.* 2012;23(1):159-166.
- 600 60. McBroom HM, Ramsay AD. The clinicopathological meeting. A means of auditing diagnostic
601 performance. *Am J Surg Pathol.* 1993;17(1):75-80.
- 602 61. McGinnis KS, Lessin SR, Elder DE, et al. Pathology review of cases presenting to a multidisciplinary
603 pigmented lesion clinic. *Arch Dermatol.* 2002;138(5):617-621.
- 604 62. Mellink WA, Henzen-Logmans SC, Bongaerts AH, Ooijen BV, Rodenburg CJ, Wiggers TH.
605 Discrepancy between second and first opinion in surgical oncological patients. *Eur J Surg Oncol.*
606 2006;32(1):108-112.
- 607 63. Murali R, Hughes MT, Fitzgerald P, Thompson JF, Scolyer RA. Interobserver variation in the
608 histopathologic reporting of key prognostic parameters, particularly Clark level, affects pathologic
609 staging of primary cutaneous melanoma. *Ann Surg.* 2009;249(4):641-647.
- 610 64. Murphy WM, Rivera-Ramirez I, Luciani LG, Wajzman Z. Second opinion of anatomical pathology: a
611 complex issue not easily reduced to matters of right and wrong. *J Urol.* 2001;165(6 Pt 1):1957-1959.
- 612 65. Nguyen PL, Schultz D, Renshaw AA, et al. The impact of pathology review on treatment
613 recommendations for patients with adenocarcinoma of the prostate. *Urol Oncol.* 2004;22(4):295-299.
- 614 66. Novis D. Routine review of surgical pathology cases as a method by which to reduce diagnostic
615 errors in a community hospital. *Pathol Case Rev.* 2005;10(2):63-67.
- 616 67. Owens SR, Dhir R, Yousem SA, et al. The development and testing of a laboratory information
617 system-driven tool for pre-sign-out quality assurance of random surgical pathology reports. *Am J Clin*
618 *Pathol.* 2010;133(6):836-841.
- 619 68. Owens SR, Wiehagen LT, Kelly SM, et al. Initial experience with a novel pre-sign-out quality
620 assurance tool for review of random surgical pathology diagnoses in a subspecialty-based university
621 practice. *Am J Surg Pathol.* 2010;34(9):1319-1323.
- 622 69. Park JH, Kim HK, Kang SW, et al. Second opinion in thyroid fine-needle aspiration biopsy by the
623 Bethesda system. *Endocr J.* 2012;59(3):205-212.
- 624 70. Pinto Sanchez MI, Smecuol E, Vazquez H, Mazure R, Maurino E, Bai JC. Very high rate of
625 misdiagnosis of celiac disease in clinical practice. *Acta Gastroenterol Latinoam.* 2009;39(4):250-253.
- 626 71. Pomianowska E, Grzyb K, Westgaard A, Clausen OP, Gladhaug IP. Reclassification of tumour origin
627 in resected periampullary adenocarcinomas reveals underestimation of distal bile duct cancer. *Eur J*
628 *Surg Oncol.* 2012;38(11):1043-1050.
- 629 72. Pongpruttipan T, Sitthinamsuwan P, Rungkaew P, Ruangchira-urai R, Vongjirad A, Sukpanichnant S.
630 Pitfalls in classifying lymphomas. *J Med Assoc Thai.* 2007;90(6):1129-1136.

- 631 73. Prayson RA, Agamanolis DP, Cohen ML, et al. Interobserver reproducibility among neuropathologists
632 and surgical pathologists in fibrillary astrocytoma grading. *J Neurol Sci.* 2000;175(1):33-39.
- 633 74. Proctor IE, McNamara C, Rodriguez-Justo M, Isaacson PG, Ramsay A. Importance of expert central
634 review in the diagnosis of lymphoid malignancies in a regional cancer network. *J Clin Oncol.*
635 2011;29(11):1431-1435.
- 636 75. Qureshi A, Loya A, Azam M, Hussain M, Mushtaq S, Mahmood T. Study of parameters to ensure
637 quality control in histopathology reporting: a meta-analysis at a tertiary care center. *Indian J Pathol*
638 *Microbiol.* 2012;55(2):180-182.
- 639 76. Raab SS, Geisinger EM, Parwani AV, Jensen C, Vrbin CM, Grzybicki DM. Effect of double viewing
640 needle core prostate biopsy tissues on error reduction. *Mod Pathol.* 2008;21 (suppl 1s):358A.
- 641 77. Raab SS, Grzybicki DM, Janosky JE, et al. Clinical impact and frequency of anatomic pathology
642 errors in cancer diagnoses. *Cancer.* 2005;104(10):2205-2213.
- 643 78. Raab SS, Grzybicki DM, Mahood LK, Parwani AV, Kuan S-F, Rao UN. Effectiveness of random and
644 focused review in detecting surgical pathology error. *Am J Clin Pathol.* 2008;130(6):905-912.
- 645 79. Raab SS, Nakhleh RE, Ruby SG. Patient safety in anatomic pathology: measuring discrepancy
646 frequencies and causes. *Arch Pathol Lab Med.* 2005;129(4):459-466.
- 647 80. Rakovitch E, Mihai A, Pignol JP, et al. Is expert breast pathology assessment necessary for the
648 management of ductal carcinoma in situ? *Breast Cancer Res Treat.* 2004;87(3):265-272.
- 649 81. Ramsay AD, Gallagher PJ. Local audit of surgical pathology. 18 month's experience of peer review-
650 based quality assessment in an English teaching hospital. *Am J Surg Pathol.* 1992;16(5):476-482.
- 651 82. Randall RL, Bruckner JD, Papenhausen MD, Thurman T, Conrad EUI. Errors in diagnosis and margin
652 determination of soft-tissue sarcomas initially treated at non-tertiary centers. *Orthopedics.*
653 2004;27(2):209-212.
- 654 83. Ray-Coquard I, Montesco MC, Coindre JM, et al. Sarcoma: concordance between initial diagnosis
655 and centralized expert review in a population-based study within three European regions. *Ann Oncol.*
656 2012;23(9):2442-2449.
- 657 84. Renshaw AA, Cartagena N, Granter SR, Gould EW. Agreement and error rates using blinded review
658 to evaluate surgical pathology of biopsy material. *Am J Clin Pathol.* 2003;119(6):797-800.
- 659 85. Renshaw AA, Gould EW. Measuring the value of review of pathology material by a second
660 pathologist. *Am J Clin Pathol.* 2006;125(5):737-739.
- 661 86. Renshaw AA, Gould EW. Correlation of workload with disagreement and amendment rates in surgical
662 pathology and nongynecologic cytology. *Am J Clin Pathol.* 2006;125(6):820-822.
- 663 87. Renshaw AA, Pinnar NE, Jiroutek MR, Young ML. Blinded review as a method for quality
664 improvement in surgical pathology. *Arch Pathol Lab Med.* 2002;126(8):961-963.
- 665 88. Renshaw AA, Pinnar NE, Jiroutek MR, Young ML. Quantifying the value of in-house consultation in
666 surgical pathology. *Am J Clin Pathol.* 2002;117(5):751-754.
- 667 89. Saglam O, Pederson A, Zhang Z, Stone CH, Kini S. Retrospective review and analysis of
668 pancreaticobiliary specimens with discordant cytohistologic correlation. *Cancer Cytopathology.*
669 2008;114(s5):421-422.
- 670 90. Santillan AA, Messina JL, Marzban SS, Crespo G, Sondak VK, Zager JS. Pathology review of thin
671 melanoma and melanoma in situ in a multidisciplinary melanoma clinic: impact on treatment
672 decisions. *J Clin Oncol.* 2010;28(3):481-486.
- 673 91. Santoso JT, Coleman RL, Voet RL, Bernstein SG, Lifshitz S, Miller D. Pathology slide review in
674 gynecologic oncology. *Obstet Gynecol.* 1998;91(5 Pt 1):730-734.
- 675 92. Scott CB, Nelson JS, Farnan NC, et al. Central pathology review in clinical trials for patients with
676 malignant glioma. A Report of Radiation Therapy Oncology Group 83-02. *Cancer.* 1995;76(2):307-
677 313.
- 678 93. Selman AE, Niemann TH, Fowler JM, Copeland LJ. Quality assurance of second opinion pathology in
679 gynecologic oncology. *Obstet Gynecol.* 1999;94(2):302-306.
- 680 94. Sharif MA, Hamdani SN. Second opinion and discrepancy in the diagnosis of soft tissue lesions at
681 surgical pathology. *Indian J Pathol Microbiol.* 2010;53(3):460-464.
- 682 95. Shoo BA, Sagebiel RW, Kashani-Sabet M. Discordance in the histopathologic diagnosis of melanoma
683 at a melanoma referral center. *J Am Acad Dermatol.* 2010;62(5):751-756.
- 684 96. Staradub VL, Messenger KA, Hao N, Wiley EL, Morrow M. Changes in breast cancer therapy
685 because of pathology second opinions. *Ann Surg Oncol.* 2002;9(10):982-987.

- 686 97. Swapp RE, Aubry MC, Salomao DR, Cheville JC. Outside case review of surgical pathology for
687 referred patients: the impact on patient care. *Arch Pathol Lab Med.* 2013;137(2):233-240.
- 688 98. Tan YY, Kebebew E, Reiff E, et al. Does routine consultation of thyroid fine-needle aspiration
689 cytology change surgical management? *J Am Coll Surg.* 2007;205(1):8-12.
- 690 99. Tatsas A, Herman J, Hruban R, et al. Second opinion in pancreatic cytopathology. *CytoJournal.*
691 2011;8(suppl 1):S89.
- 692 100. Thomas CW, Bainbridge TC, Thomson TA, McGahan CE, Morris WJ. Clinical impact of second
693 pathology opinion: a longitudinal study of central genitourinary pathology review before prostate
694 brachytherapy. *Brachytherapy.* 2007;6(2):135-141.
- 695 101. Trotter MJ, Bruecks AK. Interpretation of skin biopsies by general pathologists: diagnostic
696 discrepancy rate measured by blinded review. *Arch Pathol Lab Med.* 2003;127(11):1489-1492.
- 697 102. Tsuda H, Akiyama F, Kurosumi M, Sakamoto G, Watanabe T. Monitoring of interobserver
698 agreement in nuclear atypia scoring of node-negative breast carcinomas judged at individual
699 collaborating hospitals in the National Surgical Adjuvant Study of Breast Cancer (NSAS-BC)
700 protocol. *Jpn J Clin Oncol.* 1999;29(9):413-420.
- 701 103. van Dijk MC, Aben KK, van Hees F, et al. Expert review remains important in the histopathological
702 diagnosis of cutaneous melanocytic lesions. *Histopathology.* 2008;52(2):139-146.
- 703 104. van Rhijn BW, van der Kwast TH, Kakiashvili DM, et al. Pathological stage review is indicated in
704 primary pT1 bladder cancer. *BJU Int.* 2010;106(2):206-211.
- 705 105. Vivino FB, Gala I, Hermann GA. Change in final diagnosis on second evaluation of labial minor
706 salivary gland biopsies. *J Rheumatol.* 2002;29(5):938-944.
- 707 106. Wayment RO, Bourne A, Kay P, Tarter TH. Second opinion pathology in tertiary care of patients with
708 urologic malignancies. *Urol Oncol.* 2011;29(2):194-198.
- 709 107. Wechsler J, Bastuji-Garin S, Spatz A, et al. Reliability of the histopathologic diagnosis of malignant
710 melanoma in childhood. *Arch Dermatol.* 2002;138(5):625-628.
- 711 108. Weir MM, Jan E, Colgan TJ. Interinstitutional pathology consultations. A reassessment. *Am J Clin*
712 *Pathol.* 2003;120(3):405-412.
- 713 109. Westra WH, Kronz JD, Eisele DW. The impact of second opinion surgical pathology on the practice
714 of head and neck surgery: a decade experience at a large referral hospital. *Head Neck.*
715 2002;24(7):684-693.
- 716 110. Whitehead ME, Fitzwater JE, Lindley SK, Kern SB, Ulirsch RC, Winecoff WFI. Quality assurance of
717 histopathologic diagnoses: a prospective audit of three thousand cases. *Am J Clin Pathol.*
718 1984;81(4):487-491.
- 719 111. Wurzer JC, Al-Saleem TI, Hanlon AL, Freedman GM, Patchefsky A, Hanks GE. Histopathologic
720 review of prostate biopsies from patients referred to a comprehensive cancer center: correlation of
721 pathologic findings, analysis of cost, and impact on treatment. *Cancer.* 1998;83(4):753-759.
- 722 112. Zaino RJ, Kauderer J, Trimble CL, et al. Reproducibility of the diagnosis of atypical endometrial
723 hyperplasia: a Gynecologic Oncology Group study. *Cancer.* 2006;106(4):804-811.
- 724 113. Brimo F, Schultz L, Epstein JI. The value of mandatory second opinion pathology review of prostate
725 needle biopsy interpretation before radical prostatectomy. *J Urol.* 2010;184(1):126-130.
- 726 114. Brochez L, Verhaeghe E, Grosshans E, et al. Inter-observer variation in the histopathological
727 diagnosis of clinically suspicious pigmented skin lesions. *J Pathol.* 2002;196(4):459-466.
- 728 115. Chan TY, Epstein JI. Patient and urologist driven second opinion of prostate needle biopsies. *J Urol.*
729 2005;174(4 Pt 1):1390-1394.
- 730 116. Dhir R, Parwani AV, Zynger DL. Impact of bladder biopsy second review on pathological stage and
731 subsequent patient management. *Lab Invest.* 2009;89(suppl 1):165A.
- 732 117. Fajardo DA, Miyamoto H, Miller JS, Lee TK, Epstein JI. Identification of Gleason pattern 5 on
733 prostatic needle core biopsy: frequency of underdiagnosis and relation to morphology. *Am J Surg*
734 *Pathol.* 2011;35(11):1706-1711.
- 735 118. Khazai L, Middleton LP, Goktepe N, Liu BT, Sahin AA. Breast pathology second review identifies
736 clinically significant discrepancies in 10% of cases. *Lab Invest.* 2012;92(suppl 1):46A.
- 737 119. Kommos S, Pfisterer J, Reuss A, et al. Specialized pathology review in patients with ovarian
738 cancer: highly recommended to assure adequate treatment. Results from a prospective study. *Lab*
739 *Invest.* 2012;92(suppl 1):281A.

- 740
741
742
743
744
745
746
747
748
749
750
751
752
753
754
755
756
757
758
759
760
761
762
763
764
120. Kronz JD, Westra WH, Epstein JI. Mandatory second opinion surgical pathology at a large referral hospital. *Cancer*. 1999;86(11):2426-2435.
 121. Li X, Heller K, Cangiarella J, Simsir A. Interinstitutional second opinion in thyroid cytology: should second opinion be mandated prior to definitive surgery if fine needle aspiration was performed elsewhere? *CytoJournal*. 2011;8(3):S63.
 122. Price JA, Grunfeld E, Barnes PJ, Rheaume DE, Rayson D. Inter-institutional pathology consultations for breast cancer: impact on clinical oncology therapy recommendations. *Curr Oncol*. 2010;17(1):25-32.
 123. Raab SS, Stone CH, Jensen CS, et al. Double slide viewing as a cytology quality improvement initiative. *Am J Clin Pathol*. 2006;125(4):526-533.
 124. Renshaw AA, Gould EW. Comparison of disagreement and error rates for three types of interdepartmental consultations. *Am J Clin Pathol*. 2005;124(6):878-882.
 125. Renshaw AA, Gould EW. Comparison of disagreement and amendment rates by tissue type and diagnosis: identifying cases for directed blinded review. *Am J Clin Pathol*. 2006;126(5):736-739.
 126. Renshaw AA, Gould EW. Increasing agreement over time in interlaboratory anatomic pathology consultation material. *Am J Clin Pathol*. 2013;140(2):215-218.
 127. Safrin RE, Bark CJ. Surgical pathology sign-out. Routine review of every case by a second pathologist. *Am J Surg Pathol*. 1993;17(11):1190-1192.
 128. Tavora F, Fajardo DA, Lee TK, et al. Small endoscopic biopsies of the ureter and renal pelvis: pathologic pitfalls. *Am J Surg Pathol*. 2009;33(10):1540-1546.
 129. Tsung JS. Institutional pathology consultation. *Am J Surg Pathol*. 2004;28(3):399-402.
 130. Moher D, Liberati A, Tetzlaff J, Altman D. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *PLoS Med*. 2009;6(7):e1000097.