

Live Telemicroscopy is Substantially Equivalent to In-person Intraoperative Frozen Section Diagnosis

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Introduction

Intraoperative diagnosis by frozen section is a mainstay of surgical pathologic practice, since the use of frozen saline in the early nineteenth century to "Orth's fluid" in 1895, methylene-blue based stains in the early 1900s, and finally the mounting-medium, H&E-based modern method codified in 1931.¹ Despite good accuracy with modern methods,² access to intraoperative surgical pathology with an appropriate turn-around time (TAT)³ has been a limiting factor for small or remote surgical centers with negative impacts on cost and patient care.⁴

As a major regional referral center for pathology services, the University of Nebraska Medical Center utilizes remote telemicroscopy to provide intraoperative diagnoses to satellite hospitals. A pathology assistant or high-level technician cuts the tissue and loads the slides into the telemicroscope, and the pathologist at the main site provides the diagnoses. I sought to compare this mechanism to our standard in-person intraoperative diagnoses.

Design

All frozen section diagnoses in a four-year period were queried in our laboratory information system for surgical center, anatomic site of frozen section performed, intraoperative organ site, TAT, reading pathologist, and concordance with final (paraffin) diagnoses. Intraoperative diagnoses performed by telemicroscopy were also compared to intraoperative diagnoses on glass slides.

Telemicroscopy was performed using the SL-5 dual-slide scanner from Mikroscan®. Intraoperative reads not performed exclusively by telemicroscopy, and intraoperative reads for adequacy only, were excluded. Comparisons were performed by Kruskal-Wallis, Mann-Whitney, and Chisquare tests as appropriate. For cases with

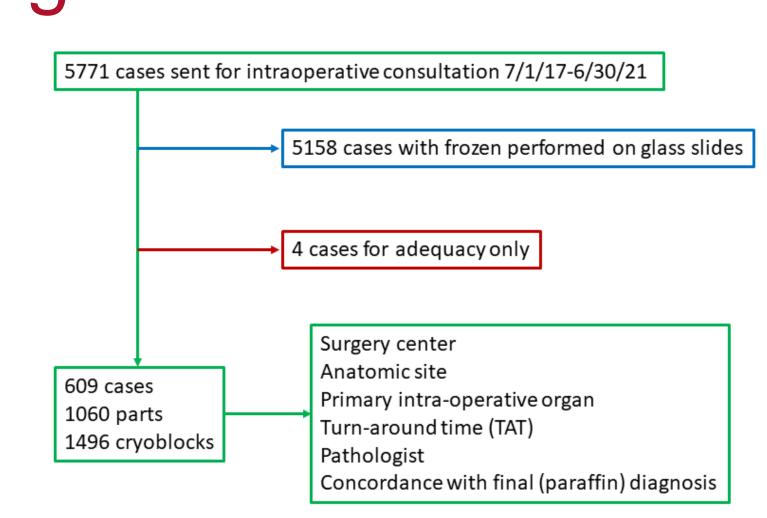


Figure 1: Analysis pipeline. Of 5771 total frozen section cases performed, 609 were performed by telemicroscopy at two regional hospitals. Of those, 576 (94.6%) had concordance data and 556 (91.3%) had TAT data available.

multiple frozen blocks, only the main specimen was included.

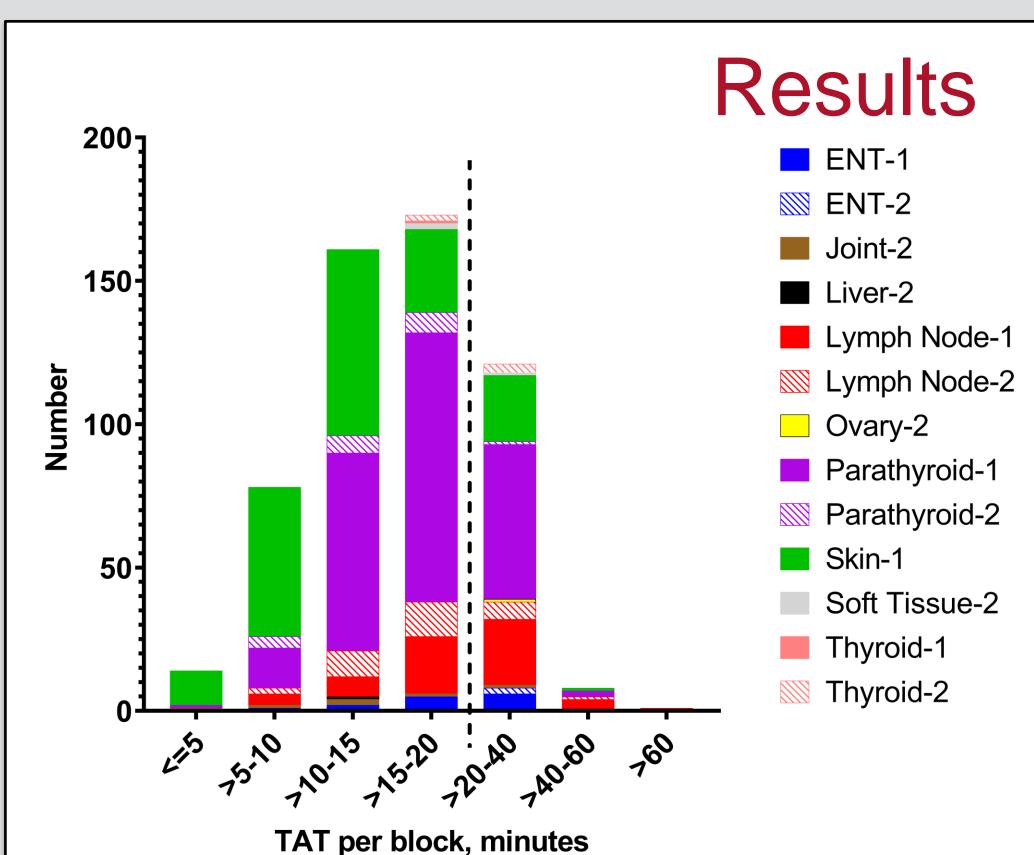


Figure 2: Turn-around time (TAT) per block for anatomic site and hospital. The TAT per block for each case (n=556) is included. The numbers in the legends indicate the surgical center. The dashed line at 20 minutes indicates the quality metric from the College of American Pathologists. Four-hundred twenty-seven (76.80%) of the 556 cases met this criterion. In two cases (0.33% of total), computer technical issues did not allow a diagnosis to be rendered. In the same timeframe, 4812/5432 in-person cases with fully available data (88.59%) met the 20-minute cutoff (p<0.0001).

Results, continued

| | Hospital 1 | | | Hospital 2 | | | Combined | | |
|---------------|------------|-------------------|-------------|------------|-------------------|-------------|----------|-------------------|-------------|
| Anatomic site | Number | <i>n</i> with TAT | mean TAT | Number | <i>n</i> with TAT | mean TAT | Number | <i>n</i> with TAT | mean TAT |
| Parathyroid | 243 | 234 | 18:01 | 18 | 18 | 15:07 | 261 | 252 | 17:49 |
| Skin | 213 | 182 | 13:08 | 1 | 0 | N/A | 214 | 182 | 13:08 |
| Lymph node | 66 | 59 | 23:14 | 31 | 30 | 19:15 | 97 | 89 | 21:53 |
| ENT | 15 | 14 | 20:04 | 2 | 2 | 24:30 | 17 | 16 | 20:37 |
| Thyroid | 1 | 1 | 18:00 | 6 | 5 | 22:00 | 7 | 6 | 21:20 |
| Joint | 0 | 0 | N/A | 6 | 6 | 13:02 | 6 | 6 | 13:02 |
| Soft tissue | 0 | 0 | N/A | 4 | 3 | 27:20 | 4 | 3 | 27:20 |
| Liver | 0 | 0 | N/A | 1 | 1 | 13:00 | 1 | 1 | 13:00 |
| Brain | 0 | 0 | N/A | 1 | 0 | N/A | 1 | 0 | N/A |
| Ovary | 0 | 0 | N/A | 1 | 1 | 23:00 | 1 | 1 | 23:00 |
| sum/mean | 538 | 490 | 16:53 | 71 | 66 | 18:15 | 609 | 556 | 17:03 |

Table 1: Tissue site and turnaround time (TAT) for telemicroscopic cases by hospital. For each hospital where telemicroscopic intraoperative diagnosis was performed, the site of frozen tissue with average block turnaround time was calculated for the 556 cases available. Tissue site was determined by the frozen tissue, not primary surgery; e.g., incidental parathyroids for ear, nose and throat (ENT) surgeries without another frozen diagnosis were included in "Parathyroid." For cases with multiple frozen tissues, only the main specimen (e.g. cancer resection) was included in calculations. Mean TAT was $17m03s\pm8m03s$. In the same timeframe, glass diagnoses in our center had a mean TAT of $14m17s\pm7m07s$ (p<0.0001). No difference in mean TAT was identified between the two external hospitals (p=0.131).

| Pathologist | Specialty | Total counts | Category A | Category B | Category C | totals |
|-------------|-----------|--------------|------------|------------|------------|--------|
| 1 | Derm | 131 | | | | 0 |
| 2 | Derm | 73 | | | | 0 |
| 3 | Surg | 47 | 3 | | 1 | 4 |
| 4 | GI | 42 | 2 | 1 | | 3 |
| 5 | B/G | 41 | 2 | 2 | | 4 |
| 6 | GI | 40 | 1 | | | 1 |
| 7 | GI | 40 | | | | 0 |
| 8 | GI | 39 | | 2 | | 2 |
| 9 | Surg | 37 | | 1 | | 1 |
| 10 | Neuro | 23 | | | | 0 |
| 11 | GI | 22 | | 1 | 2 | 3 |
| 12 | B/G | 22 | | | | 0 |
| 13 | Surg | 14 | | | | 0 |
| 14 | Neuro | 8 | | | | 0 |
| 15 | Surg | 6 | | 1 | | 1 |
| 16 | Surg | 6 | | | | 0 |
| 17 | Surg | 5 | | | | 0 |
| 18 | B/G | 4 | | | | 0 |
| 19 | Surg | 2 | | | | 0 |
| 20 | Surg | 2 | | | | 0 |
| 21 | Surg | 2 | | | | 0 |
| 22 | GI | 2 | | | | 0 |
| 23 | Surg | 1 | | | | 0 |
| | Sum | 609 | 8 | 8 | 3 | 19 |

Table 2: Telemicroscopy by reading pathologist and discordance. Twenty-three different pathologists performed telemicroscopic diagnoses an average of 26.5 times (range: 1-131). Of 609 cases, 576 (94.6%) had concordance data. Of those, 557 (96.7%) were concordant. Nineteen discordant cases (3.30% of available) were identified among eight pathologists. Categories A; does not change management; B: may change management; C: requires additional intervention. Derm: dermatopathology; Surg: multiple fields of surgical pathology; GI: gastrointestinal pathology; B/G: breast/gynecologic pathology; Neuro: neuropathology.

Results, continued

| Category | Anatomic site | Number | Narrative | Specialty |
|----------|---------------|--------|--|------------|
| A | Lymph node | 2 | Additional node positive for breast carcinoma | B/G x2 |
| | | | One case already known positive | |
| | | | One case negative on frozen with isolated carcinoma cells on levels | |
| | | 1 | Additional node negative for breast carcinoma | GI |
| | Skin | 1 | Focal positive margin > negative margin | Surg |
| | SKIII | 1 | Possible tumor > negative | GI |
| | Parathyroid | 1 | Misreport weight/more parathyroid in fat | Surg |
| | | 1 | No parathyroid > small parathyroid | GI |
| | | 1 | Parathyroid > thyroid | GI |
| В | Skin | 3 | Focal positive margin (BCC or SCC) > negative margin | GI x2, B/G |
| | | 1 | Actinic keratosis > in situ squamous cell carcinoma | GI |
| | | 1 | Negative margin > focal positive margin | GI |
| | Lymph node | 2 | Negative for breast carcinoma > positive | Surg, B/G |
| | ENT 1 | | Negative for squamous cell carcinoma > positive | Surg |
| С | Thyroid | 1 | Lymphocytic thyroiditis with focal papillary change > follicular thyroid carcinoma | Surg |
| | Parathyroid | 1 | Endocrine tissue, favor parathyroid > thyroid | GI |
| | Skin | 1 | Negative margin > positive margin | GI |

Table 3: Discordance by anatomic site. The specific discordant scenarios for the nineteen cases are outlined. Eight of 23 reading pathologists had any discordant read. The subspecialty of the reading pathologist for these nineteen cases is included. B/G: breast/gynecologic pathology; GI: gastrointestinal pathology; Surg: multiple fields of surgical pathology.

Table 4: Discordance by method of intraoperative diagnosis. There is not difference in concordance between inperson and telemicroscopic diagnosis at either site (p=0.375).

| | | Glass | Hospital 1 | Hospital 2 |
|----------|------------|-------|------------|------------|
| of no | Concordant | 6539 | 494 | 63 |
| | Discordant | 262 | 16 | 3 |
| n- | А | 90 | 7 | 1 |
| at | В | 140 | 7 | 1 |
| | С | 32 | 2 | 1 |

Discussion

Diagnoses by telemicroscopy did not meaningfully differ from on-glass on TAT and concordance; although in-person diagnoses were statistically faster, the great majority of telemicroscopic diagnoses were returned in <20 minutes. This remained true through numerous pathologists, pathology assistants and/or technicians, two different hospitals, and over four years. The concentration of discordant diagnoses among relatively few pathologists suggests a level of individual comfort with telemicroscopy and/or frozen section. Rare cases of technological issues prevented telemicroscopic diagnosis. Overall, this provides further justification for continued use and expansion of telemicroscopic services in primary intraoperative diagnoses.

References

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