

Eurocytology - Immunocytochemistry

Objectives:

- Present the current landscape of immunocytochemistry in Europe
- Discuss application of QA/QC measures for reliable ICC
- Describe parameters to assess quality of ICC on cytology slides
- Practice quality assessment of ICC on a selected cases

Introduction

Immunocytochemistry (ICC) has become an indispensable ancillary method in a modern cytopathology for fine-tuning a diagnosis as well as for assessing prognostic and predictive markers. As shown by survey performed by European Federation of Cytology Societies (EFCS), ICC is extensively used in European cytology laboratories on a differently prepared and fixed cytology slides (1). Formalin-fixed paraffin-embedded (FFPE) cell blocks are regarded the optimal choice for ICC from QA/QC perspective, however low cellular cell blocks with insufficient cells are the main issue in a daily cytology practice as shown by the survey performed among European (data not published yet) and USA laboratories (2). Other cytology slides are therefore vital to perform ICC on a limited and low cellular cytology samples. The basic requirements to assure reliable ICC results on non-FFPE cytology preparations are control slides prepared as the patient sample as well as optimization and validation of ICC protocols.

Principles, basic steps, quality assurance and quality control (QA/QC) measures as well as platforms, reagents and antibodies used for ICC on cytology slides are the same as for immunohistochemistry (IHC) on a standardly prepared formalin-fixed paraffin-embedded (FFPE) tissue and are described in several articles (3-9). In this chapter, special consideration of ICC on cytology samples will be presented.

Chapters

- State of the Art and Science of Immunocytochemistry – webinar originally presented for International Society for IHC and Molecular Morphology (ISIMM) on Wednesday, March 8, 2023 and link to the article published in *Acta Cytologica*
- Assessing quality of ICC on a cytology slides
- Control slides
- Optimization and validation of ICC

References

1. Srebotnik Kirbis I. State of the Art and Science of Immunocytochemistry. *Acta Cytol.* 2025;69(1):51-59. doi: 10.1159/000539634. Epub 2024 Jun 10. PMID: 38857584.
2. Srebotnik Kirbiš I, Rodrigues Roque R, Bongiovanni M, Strojjan Fležar M, Cochand-Priollet B. Immunocytochemistry practices in European cytopathology laboratories—Review of European Federation of Cytology Societies (EFCS) online survey results with best practice recommendations. *Cancer Cytopathology.* 2020;128(10):757-66.

3. Crapanzano JP, Heymann JJ, Monaco S, Nassar A, Saqi A. The state of cell block variation and satisfaction in the era of molecular diagnostics and personalized medicine. *Cytojournal*. 2014;11:7.
4. O'Hurley G, Sjostedt E, Rahman A, Li B, Kampf C, Ponten F, et al. Garbage in, garbage out: a critical evaluation of strategies used for validation of immunohistochemical biomarkers. *Mol Oncol*. 2014;8(4):783-98.
5. Fitzgibbons PL, Bradley LA, Fatheree LA, Alsabeh R, Fulton RS, Goldsmith JD, et al. Principles of analytic validation of immunohistochemical assays: Guideline from the College of American Pathologists Pathology and Laboratory Quality Center. *Arch Pathol Lab Med*. 2014;138(11):1432-43.
6. Fitzgibbons PL, Goldsmith JD, Souers RJ, Fatheree LA, Volmar KE, Stuart LN, et al. Analytic Validation of Immunohistochemical Assays: A Comparison of Laboratory Practices Before and After Introduction of an Evidence-Based Guideline. *Arch Pathol Lab Med*. 2017;141(9):1247-54.
7. Hammond MEH, Hayes DF, Dowsett M, Allred DC, Haggerty KL, Badve S, et al. American Society of Clinical Oncology/College of American Pathologists Guideline Recommendations for Immunohistochemical Testing of Estrogen and Progesterone Receptors in Breast Cancer. *Journal of Clinical Oncology*. 2010;28(16):2784-95.
8. Canadian Association of Pathologists-Association canadienne des pathologistes National Standards C, Torlakovic EE, Riddell R, Banerjee D, El-Zimaity H, Pilavdzic D, et al. Canadian Association of Pathologists-Association canadienne des pathologistes National Standards Committee/Immunohistochemistry: best practice recommendations for standardization of immunohistochemistry tests. *Am J Clin Pathol*. 2010;133(3):354-65.
9. Torlakovic EE, Cheung CC, D'Arrigo C, Dietel M, Francis GD, Gilks CB, et al. Evolution of Quality Assurance for Clinical Immunohistochemistry in the Era of Precision Medicine. Part 3: Technical Validation of Immunohistochemistry (IHC) Assays in Clinical IHC Laboratories. *Applied Immunohistochemistry & Molecular Morphology*. 2017;25(3):151-
10. Torlakovic EE, Nielsen S, Vyberg M, Taylor CR. Getting controls under control: the time is now for immunohistochemistry. *J Clin Pathol*. 2015;68(11):879-82.