

# **Background**



- Management of urothelial carcinoma (UC) is expensive
  - High recurrence rate, long disease course, intensive surveillance strategies
- Urine cytology is a noninvasive and cost-effective screening/ diagnostic test
  - Surveillance intervals and hematuria workups
- Urine cytology diagnostic classification: a long journey
  - None had gained wide acceptance: lack of standard terminology and diagnostic criteria
  - A variety of terminologies are currently utilized in different pathology labs
  - High percentage of indeterminate (atypical) diagnosis: may lead to unnecessary procedures

# The Paris System (TPS) for Reporting Urine Cytology

- 2013 International Cytology Congress in Paris: recognized the need to standardize terminology and reporting
- TPS was developed, presented in conferences and published in 2016
- Advantages:
  - More definite morphological and numerical diagnostic criteria
  - Aims to improve sensitivity and specificity in detecting high-grade UC (HGUC)
  - Decrease indeterminate/atypical diagnosis



# The Paris System (TPS) for Reporting Urine Cytology

- Unsatisfactory
- NHGUC (negative for high-grade UC)
- AUC (atypical urothelial cells): N/C ratio > 0.5 + one of the following-
  - Hyperchromasia/Irregular clumpy chromatin/Irregular nuclear contours
- SHGUC (suspicious for HGUC): N/C ratio > 0.7 <10 abnormal cells
  - · Hyperchromasia + one of the following
  - Irregular clumpy chromatin/Irregular nuclear membranes
- **HGUC** (high-grade UC):

N/C ratio > 0.7 >5-10 abnormal cells

- Similar to SHGUC
- LGUN (low-grade urothelial neoplasm): papilloma, PUNLMP, LGUC
- Other malignancies

Pre-TPS terminology

NUC (negative for UC)

AUC, included MAUC (Mildly atypical urothelial cells, favor reactive changes)



## **Aim**



 To study the impact of TPS on the urine cytology diagnostic patterns in our high-volume community practice.

 Introduce TPS to urologists and demonstrate its potential impact on urological practices.



## **Methods**



| Pre-TPS                                       | Learning Period   | Post-TPS   |
|---|---|--|
| Jan 2013 to Dec 2014 (two years, 7,658 cases) | <ul> <li>Jan 2015 - April 2016</li> <li>Initial grand rounds presentation</li> <li>Change terminology</li> <li>Daily pathology consensus conference</li> <li>Urine cytology case conference /2 wk.</li> </ul> | May 2016 to April 2018 (two years, 20,027 cases) |

- The comparison was made between Pre- and Post-TPS diagnostic categories and different collection methods
- Each diagnostic category was correlated with UroVysion results



# Results



Table 1. Urine cytology diagnostic categories Pre- and Post-TPS

| Diagnosis     | Pre-TPS          |                     | Post-TPS |       | P Value  |  |
|---------------|------------------|---------------------|----------|-------|----------|--|
| Diagnosis     | n                | %                   | n        | %     | P value  |  |
| NUC/NHGUC     | 5,293            | 69.2%               | 18,507   | 92.4% | <0.00001 |  |
| AUC<br>(MAUC) | 2,227<br>(1,437) | 29%<br>(65% of AUC) | 1,237    | 6.2%  | <0.00001 |  |
| SHGUC/HGUC    | 138              | 1.8%                | 282      | 1.4%  | 0.0057   |  |
| Total         | 7,658            | 100%                | 20,026   | 100%  |          |  |



Table 2. Diagnostic categories in voided vs instrumented urine in Pre- and Post-TPS

| Collection method | Voided Urine n (%) |          | Instrumented urine n (%) |          |  |
|-------------------|--------------------|----------|--------------------------|----------|--|
| Diagnosis         | Pre-TPS            | Post-TPS | Pre-TPS                  | Post-TPS |  |
| NUC/NHGUC         | 71.5%              | 92.7%    | 60.9%                    | 88.5%    |  |
| AUC               | 26.9%              | 6%       | 36.6%                    | 9.1%     |  |
| SHGUC/HGUC        | 1.6%               | 1.3%     | 2.5%                     | 2.4%     |  |
| Total %           | 100%               | 100%     | 100%                     | 100%     |  |
| <b>Total</b> n    | 5,951              | 18,877   | 1,707                    | 1,149    |  |

Table 3. AUC Rate Among Pathologists Pre- and Post-TPS

| Pathologists   | A   | В   | С   | D   | E   | Average |
|----------------|-----|-----|-----|-----|-----|---------|
| Pre-TPS AUC %  | 24  | 26  | 37  | 39  | 23  | 29      |
| Post-TPS AUC % | 5.8 | 5.3 | 7.2 | N/A | 6.4 | 6.2     |

All pathologists showed significant decreases in AUC (P<0.00001) with less interobserver variation (23-39% vs 5.3-7.2%)



# Figure 1. Urine Cytology Diagnosis Demonstrated Superior Correlation with UroVysion Results in Post-TPS

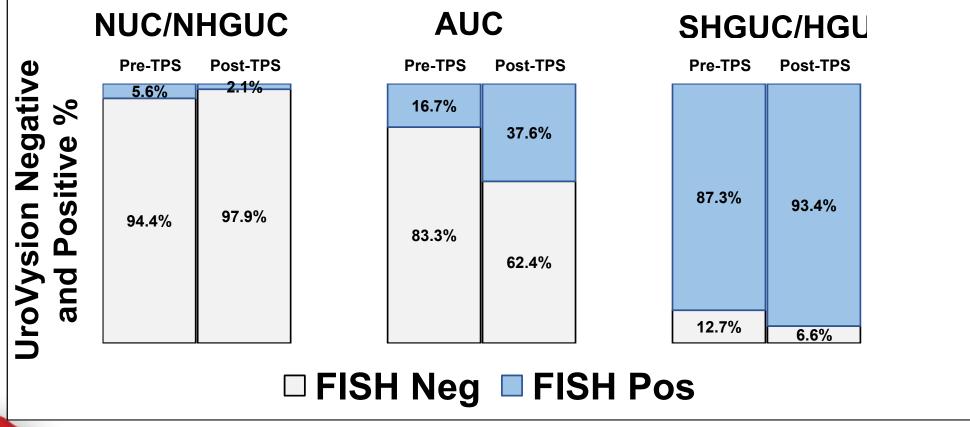




Table 4. Utilization of UroVysion test was significantly decreased Post-TPS for AUC category

| AUC with UroVysion    | Pre-TPS     | Post-TPS     |
|-----------------------|-------------|--------------|
| Total urine cytology  | 7,658       | 20,026       |
| AUC n (atypical rate) | 2,227 (29%) | 1,237 (6.2%) |
| AUC with UroVysion n  | 980         | 550          |

Presumptive: UroVysion requests in AUC if we didn't implement TPS: 20,026 cases x 29% atypical rate x 44% requests in AUC= 2,555 UroVysion tests Compared to 550 → a significant decrease in UroVysion requests and medical cost



## Conclusion



- Implementation of TPS resulted in a significant decrease in atypical diagnoses with less interobserver variation.
- AUC was significantly better correlated with UroVysion results
  - Decreased UroVysion requests
  - Saved medical cost
- AUC should be considered a clinically relevant group, and requires more serious clinical workup in the TPS era.



## References



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- 3. Miki Y, Neat M, Chandra A. Application of The Paris System to atypical urine cytology samples: correlation with histology and UroVysion<sup>®</sup> FISH. *Cytopathology*. 2017 Apr;28(2):88-95.
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