#### RESEARCH ARTICLE

# Gamma gap thresholds and HIV, hepatitis C, and monoclonal gammopathy

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Table 2. Diagnostic performance of gamma gap for HIV using different gamma gap thresholds, N = 25,680.

| Gamma gap (g/dL) | Sn, % | Sp, % | LR +  | LR - | Overall AUC (95% CI)      |  |  |
|------------------|-------|-------|-------|------|---------------------------|--|--|
| ≥ 2.5            | 94.8  | 10.5  | 1.1   | 0.5  | 0.80 (95% CI: 0.75, 0.85) |  |  |
| ≥ 3.0            | 84.4  | 51.9  | 1.8   | 0.3  |                           |  |  |
| ≥ 3.5            | 59.3  | 88.1  | 5.0   | 0.5  |                           |  |  |
| ≥ 4.0            | 39.3  | 98.4  | 24.2  | 0.6  |                           |  |  |
| ≥ 4.5            | 19.3  | 99.6  | 54.1  | 0.8  |                           |  |  |
| ≥ 5.0            | 10.4  | 99.9  | 139.4 | 0.9  |                           |  |  |
| ≥ 5.5            | 4.4   | 100.0 | 283.9 | 1.0  |                           |  |  |

Table 3. Diagnostic performance of gamma gap for HCV using different gamma gap threshold, N = 45,134.

| Gamma gap (g/dL) | Sn, % | Sp, % | LR + | LR - | Overall AUC (95% CI)      |  |  |
|------------------|-------|-------|------|------|---------------------------|--|--|
| ≥ 2.5            | 97.8  | 9.9   | 1.1  | 0.2  | 0.74 (95% CI: 0.72, 0.76) |  |  |
| ≥ 3.0            | 80.8  | 50.2  | 1.6  | 0.4  |                           |  |  |
| ≥ 3.5            | 44.5  | 86.9  | 3.4  | 0.6  |                           |  |  |
| ≥ 4.0            | 19.0  | 97.8  | 8.5  | 0.8  |                           |  |  |
| ≥ 4.5            | 7.0   | 99.5  | 13.7 | 0.9  |                           |  |  |
| ≥ 5.0            | 2.6   | 99.8  | 16.3 | 1.0  |                           |  |  |
| ≥ 5.5            | 1.1   | 99.9  | 13.9 | 1.0  |                           |  |  |

Table 4. Diagnostic performance of gamma gap for MGUS using different gamma gap thresholds, N = 6,118.

| Gamma gap (g/dL) | Sn, % | Sp, % | LR + | LR - | Overall AUC (95% CI)      |  |  |
|------------------|-------|-------|------|------|---------------------------|--|--|
| ≥ 2.5            | 97.6  | 6.4   | 1.0  | 0.4  | 0.64 (95% CI: 0.60, 0.69) |  |  |
| ≥ 3.0            | 75.7  | 39.9  | 1.3  | 0.6  |                           |  |  |
| ≥ 3.5            | 39.1  | 80.1  | 2.0  | 0.8  |                           |  |  |
| ≥ 4.0            | 15.4  | 95.4  | 3.4  | 0.9  |                           |  |  |
| ≥ 4.5            | 7.7   | 98.9  | 6.9  | 0.9  |                           |  |  |
| ≥ 5.0            | 5.3   | 99.7  | 16.7 | 0.9  |                           |  |  |
| ≥ 5.5            | 3.0   | 99.8  | 17.6 | 1.0  |                           |  |  |

#### Conclusions:

- Gamma gap is not a sufficient screening tool (you'd want high sensitivity for a screen)
- An elevated gamma gap warrants further testing for HCV and HIV but not MGUS
- Many patients with HIV and HCV did not have an elevated gamma gap
- HIV and HCV stage of treatment influence the gamma gap

#### Limitations:

- This study was performed in defined patient populations, so patients with comorbid conditions were not assessed and there was no assessment of a mixed population (HIV, HCV, and MGUS)
- These were outpatients and none had autoimmune conditions, chronic inflammatory conditions, or chronic infections
- When the gamma gap was test further in this study

HIV: age > 18HCV: age > 6MGUS: age > 50

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## A Clinical Decision Rule to Aid Ordering of Serum and Urine Protein Electrophoresis for Case-Finding of Paraproteins in Hospitalized Inpatients

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### For MGUS use the following equation:

$$Logit \left[ \frac{SPE/UPE^+}{SPE/UPE^-} \right] = -4.4 + 1.5 x globulin + 1.1 x age + 0.6 x hgb + 0.4 x sex + .2 x gfr$$

| Variables    | OR (95% CI)            | Scores |   |
|--------------|------------------------|--------|---|
| Globulin, g/ | I                      |        |   |
| >41          | 4.5 (3.8, 5.4)         | 5      |   |
| ≤41          |                        | 0      |   |
| Age, years   |                        |        |   |
| ≥60          | 2.9 (2.4, 3.7)         | 3      | 1.00 -                                      |
| <60          |                        | 0      | 0.90 -                                      |
| Hemoglobin   | , g/1                  |        | 0.80 -                                      |
| <121         | 1.9 (1.5, 2.3)         | 2      | 0.70 -                                      |
| ≥121         |                        | 0      | 0.70 - 0.60 - >=8                           |
| Sex          |                        |        | Sensitivity 0.50 - S=10                     |
| Male         | 1.6 (1.4, 1.9)         | 2      | 0.40  |
| Female       |                        | 0      | 0.30 -                                      |
| eGFR, m1/m   | in/1.73 m <sup>2</sup> |        | 0.20 - >=12                                 |
| <60          | 1.3 (1.1, 1.5)         | 1      | 0.10  |
| ≥60          |                        | 0      | 0.00 -                                      |
| Total scores |                        | 0-13   | 0.00 0.25 0.50 0.75 1.00<br>1 - Specificity |

| Score (prob. of para-protein) | Derivation set |          |                          |         | Validation set |          |                          |        |  |
|-------------------------------|----------------|----------|--------------------------|---------|----------------|----------|--------------------------|--------|--|
|                               | SPE/UPE        |          | LR <sup>+</sup> (95% CI) | PPV (%) | SPE/UPE        |          | LR <sup>+</sup> (95% CI) | PPV(%) |  |
|                               | Positive       | Negative |                          |         | Positive       | Negative |                          |        |  |
| 0-5(Low)                      | 207            | 6,537    | 1.0                      | 3.1     | 91             | 2,797    | 1.0                      | 3.1    |  |
| 6-10(Med)                     | 303            | 2,205    | 2.5(2.4-2.7)             | 12.1    | 129            | 941      | 2.5(2.3-2.7)             | 12.1   |  |
| ≥11(High)                     | 294            | 506      | 6.7(5.9-7.6)             | 36.8    | 127            | 221      | 6.6(5.4-79)              | 36.5   |  |