Supplementary Material

**Supplementary Table S1.** Additional analytical parameters in stroke patients at hospital admission.

SD: standard deviation; IQR: interquartile range.

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| --- | --- | --- | --- | --- |
| **Variable** | **Mean** | **SD** | **Median** | **IQR** |
| Hemoglobin (g/dL) | 13.4 | 1.9 | 13.6 | 11.9-14.6 |
| Erythrocytes (106/mL) | 4.5 | 0.6 | 4.58 | 4.1-5 |
| Platelets (103/mL)  | 224.5 | 81.4 | 216 | 164.7-260 |
| Leukocytes (103/mL)  | 9.1 | 3.4 | 8.4 | 6.9-10.6 |
| LDH (IU/mL) | 258.6 | 67.5 | 256 | 220.7-299.5 |
| Ferritin (ng/mL) | 177.3 | 173.8 | 128.1 | 76.8-208.4 |
| C-reactive protein (mg/dL) | 1.4 | 2.3 | 0.51 | 0.2-1.2 |
| Total cholesterol (mg/dL) | 168.8 | 42.2 | 165 | 138.8-195.8 |
| Non-HDL Cholesterol (mg/dL) | 121.5 | 40 | 113.2 | 92.9-143.8 |
| Alanine aminotransferase (IU/mL) | 18 | 11.8 | 15.4 | 11-21.9 |
| Aspartate aminotransferase (IU/mL) | 21.7 | 10.5 | 19 | 15.5-25 |
| Creatinine (mg/dL) | 0.93 | 0.46 | 0.83 | 0.7-1 |

**Supplementary Table S2.** Prevalence of antiphospholipid antibodies in the control group of blood donors and ischemic stroke patients.

aB2GPI: anti-β2-glycoprotein-I antibodies; aCL: anti-cardiolipin antibodies; aPL: antiphospholipid antibodies; aPS/PT: anti-phosphatidylserine/prothrombin antibodies.

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| **Antibodies** | **BLOOD DONORS**(N=501) | **STROKE PATIENTS**(N=245) | **p-value** |
| Number of positive | % | Number of positive | % |
| IgG aB2GPI | 1 | 0.2 | 5 | 2.0 | 0.008 |
| IgM aB2GPI | 1 | 0.2 | 10 | 4.1 | <0.001 |
| IgA aB2GPI | 5 | 1.0 | 49 | 20.0 | <0.001 |
| IgG aCL | 2 | 0.4 | 4 | 1.6 | 0.076 |
| IgM aCL | 0 | 0.0 | 10 | 4.1 | <0.001 |
| IgG/IgM aPSPT | 6 | 1.2 | 19 | 7.8 | <0.001 |
| Classic aPL(IgG/IgM aCL/aB2GPI) | 3 | 0.6 | 15 | 6.1 | <0.001 |
| Any aPL  | 39 | 7.8 | 70 | 28.6 | <0.001 |

**Supplementary Table S3**. Mean and median levels of antiphospholipid antibodies in the reference population, blood donors and stroke patients.

aB2GPI: anti-β2-glycoprotein-I antibodies; aCL: anti-cardiolipin antibodies; aPS/PT: anti-phosphatidylserine/prothrombin antibodies; SD: standard deviation; IQR: interquartile range.

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| **Antibodies** | **BLOOD DONORS**(n=501) | **REFERENCE POPULATION**(n=121) | **STROKE PATIENTS**(n=245) |
| Mean | SD | Median | IQR | Mean | SD | Median | IQR | Mean | SD | Median | IQR |
| IgA aB2GP1 | 4,3 | 4,36 | 3,3 | 2,4-4,7 | 7,8 | 14,51 | 4,3 | 2,9-7,1 | 19,9 | 37,63 | 5,1 | 3,5-12,9 |
| IgG aB2GP1 | 1,6 | 1,66 | 1,4 | 1,4-1,4 | 1,8 | 1,73 | 1,4 | 1,4-1,4 | 3,4 | 15,72 | 1,4 | 1,4-1,4 |
| IgM aB2GP1 | 1,6 | 2,03 | 1,0 | 0,4-1,9 | 3,5 | 7,14 | 1,6 | 0,7-3,8 | 4,7 | 15,28 | 1,2 | 0,5-3,3 |
| IgG aCL | 1,9 | 1,80 | 1,6 | 1,6-1,6 | 2,3 | 2,31 | 1,6 | 1,6-1,6 | 3,6 | 15,31 | 1,6 | 1,6-1,6 |
| IgM aCL | 1,3 | 1,57 | 0,9 | 0,3-1,9 | 4,0 | 7,54 | 1,9 | 0,7-4,7 | 4,5 | 15,22 | 1,1 | 0,5-3,0 |
| IgG aPS/PT | 8,8 | 5,89 | 7,34 | 5,7-9,6 | 10,7 | 9,70 | 8,2 | 6,1-11,2 | 11,6 | 13,16 | 7,9 | 6,4-12,1 |
| IgM aPS/PT | 14,2 | 8,49 | 11,8 | 9,2-16,8 | 10,6 | 8,55 | 8,0 | 6,1-11,3 | 16,0 | 21,46 | 10,4 | 8,5-15,9 |
|  |  |  |  |  |  |  |  |  |  |

**Supplementary Table S4.** Score values and characteristics of the ROC curve. A) Score values with sensitivity, specificity and predictive values. B) Youden index.

+LR: positive likelihood ratio; -LR: negative likelihood ratio; PPV: positive predictive value; NPV: negative predictive value; CI: confidence interval.

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| **A) Score values and statistical characteristics of the ROC curve** |  |  |
| Score | Sensitivity | 95% CI | Specificity | 95% CI | +LR | 95% CI | -LR | 95% CI | PPV | NPV |
| 0 | 100 | 98.5 - 100 | 0.00 | 0.0 - 3.0 | 1.00 | 1 - 1 |  |  | 66.9 |  |
| 1 | 92.65 | 88.6 - 95.6 | 35.54 | 27.0 - 44.8 | 1.44 | 1.3 - 1.6 | 0.21 | 0.1 - 0.3 | 74.4 | 70.5 |
| **2** | 76.73 | 70.9 - 81.9 | 59.50 | 50.2 - 68.3 | 1.89 | 1.5 - 2.4 | 0.39 | 0.3 - 0.5 | 79.3 | 55.8 |
| 3 | 49.80 | 43.4 - 56.2 | 80.99 | 72.9 - 87.6 | 2.62 | 1.8 - 3.9 | 0.62 | 0.5 - 0.7 | 84.1 | 44.3 |
| 4 | 38.37 | 32.2 - 44.8 | 90.91 | 84.3 - 95.4 | 4.22 | 2.4 - 7.6 | 0.68 | 0.6 - 0.8 | 89.5 | 42.1 |
| 5 | 21.22 | 16.3 - 26.9 | 96.69 | 91.8 - 99.1 | 6.42 | 2.4 - 17.3 | 0.81 | 0.8 - 0.9 | 92.9 | 37.7 |
| 6 | 9.80 | 6.4 - 14.2 | 98.35 | 94.2 - 99.8 | 5.93 | 1.4 - 24.7 | 0.92 | 0.9 - 1 | 92.3 | 35.0 |
| 7 | 4.08 | 2.0 - 7.4 | 100 | 97.0 - 100 |  |  | 0.96 | 0.9 - 1 | 100.0 | 34.0 |
| >7 | 0.00 | 0- 1.5 | 100 | 97.0 - 100 |  |  | 1 | 1 - 1 |  | 33.1 |
| **B) Youden Index** |
| Statistic | Value | 95% CI | Associated  criterion | 95% CI |
| Area under the ROC curve | 0.746 | 0.698 - 0.790 |  |  |
| Youden index J | 0.362 | 0.264 - 0.461 | 2 | 1 - 3 |

**Supplementary Figure S1.** Study disposition. The cohort included 245 patients from the Stroke Unit with a confirmed ischemic stroke who were followed-up for 2 years.

aB2GPI: anti-β2-glycoprotein-I antibodies; aCL: anti-cardiolipin antibodies; aPS/PT: anti-phosphatidylserine/prothrombin antibodies.

**Supplementary Figure S2.** Distribution of aPL levels in blood donors (grey) and stroke patients (black). A) Classification criteria aPL. B) Extra-criteria aPL.

aB2GPI: anti-β2-glycoprotein-I antibodies; aCL: anti-cardiolipin antibodies; aPS/PT: anti-phosphatidylserine/prothrombin antibodies.

**Excel file**. Data on age, sex, presence of risk factors and score value for stroke patients.