

Supplementary Material

1. Supplementary Figures and Tables

Supplementary Table 1. Baseline characteristics of patients included and excluded.

| Variables | Included(n=638) | Excluded for missing glucose, HbA1c or brain imaging (n=1022) | P value |
|---|-----------------|---|---------|
| Age(year), median (IQR) | 70(61-79) | 70(59-79) | 0.69 |
| Male, n (%) | 378(59.2) | 575(56.3) | 0.231 |
| Hypertension, n (%) | 367(57.5) | 593(58.0) | 0.841 |
| Hyperlipidemia, n (%) | 46(7.2) | 116(11.4) | 0.006 |
| Diabetes mellitus, n (%) | 159(24.9) | 238(23.3) | 0.448 |
| Atrial fibrillation/Atrial flutter, n (%) | 219(34.3) | 380(37.2) | 0.239 |
| Previous ischemic stroke/TIA, n (%) | 63(9.9) | 145(14.2) | 0.01 |
| Valvular heart disease, n (%) | 59(9.2) | 130(12.7) | 0.03 |
| Smoking, n (%) | 225(35.3) | 351(34.3) | 0.701 |
| Alcohol consumption, n (%) | 177(27.7) | 272(26.6) | 0.615 |
| Baseline NIHSS, median (IQR) | 10(3-15) | 9(3-15) | 0.06 |

HbA1c, hemoglobin A1c; IQR, interquartile range; TIA, transient ischemic attack; NIHSS: National Institutes of Health Stroke Scale

Supplementary Table 2. Predictive factors for the development of moderate-to-severe CED on training dataset

| Variables | Univariate analysis OR (95%CI) | P value | Multivariate analysis OR (95%CI), model 1 | P value | Multivariate analysis OR (95%CI), model 2 | P value |
|------------------------------------|-----------------------------------|---------|--|---------|--|---------|
| Age | 1.01(0.99-1.03) | 0.274 | 1.00(0.98-1.03) | 0.697 | 1.01(0.98-1.03) | 0.653 |
| Male | 0.83(0.51-1.34) | 0.448 | | | | |
| Hypertension | 1.13(0.70-1.83) | 0.615 | | | | |
| Hyperlipidemia | 1.38(0.48-3.93) | 0.552 | | | | |
| Atrial fibrillation/Atrial flutter | 2.62(1.61-4.26) | <0.001 | | | | |
| Previous stroke/TIA | 0.84(0.39-1.80) | 0.644 | | | | |
| Valvular heart disease | 0.85(0.38-1.91) | 0.698 | | | | |
| Diabetes mellitus | 0.64(0.34-1.20) | 0.161 | 0.27(0.09-0.79) | 0.016 | 0.44(0.17-1.19) | 0.108 |
| Smoking | 0.79(0.47-1.34) | 0.381 | | | | |
| Alcohol consumption | 0.68(0.33-1.39) | 0.29 | | | | |
| Onset to admission time | 0.95(0.92-0.98) | 0.002 | | | | |
| Baseline NIHSS(≥ 15) | 4.32(2.62-7.13) | <0.001 | 1.93(1.05-3.57) | 0.036 | 2.23(1.21-4.11) | 0.01 |
| Occlusion site | | | | | | |
| No record or other | Reference | | Reference | | Reference | |
| Carotid occlusion | 27.71(11.96-64.20) | <0.001 | 18.44(7.28-46.70) | <0.001 | 16.45(6.57-41.20) | <0.001 |
| MCA occlusion (M1-M2) | 9.65(4.32-21.54) | <0.001 | 8.47(3.44-20.87) | <0.001 | 7.89(3.24-19.23) | <0.001 |
| Endovascular treatment | 2.10(1.18-3.73) | 0.012 | 0.43(0.20-0.92) | 0.03 | 0.58(0.28-1.21) | 0.149 |
| Intravenous thrombolysis | 1.43(0.75-2.72) | 0.279 | 1.27(0.57-2.79) | 0.559 | 1.33(0.61-2.89) | 0.479 |
| Antihypertensive therapy | 0.91(0.55-1.49) | 0.698 | | | | |
| Insulin | 1.24(0.63-2.43) | 0.536 | | | | |
| Oral hypoglycemic agents | 0.62(0.28-1.38) | 0.242 | | | | |
| Antiplatelet | 0.37(0.20-0.70) | 0.002 | | | | |
| Statin | 0.78(0.41-1.51) | 0.462 | | | | |
| mTICI (2b/3) | 0.24(0.05-1.10) | 0.067 | | | | |
| RPG | 1.00(0.93-1.08) | 0.964 | 1.03(0.90-1.18) | 0.633 | 1.04(0.92-1.18) | 0.535 |
| SHR (per 0.1-point increases) | 1.40(1.27-1.54) | <0.001 | 1.39(1.24-1.57) | <0.001 | | |
| SHR(≥ 1.25) | 8.97(5.14-15.65) | <0.001 | | | 6.87(3.46-13.63) | <0.001 |

OR, odds ratio; CI, confidence interval; TIA, transient ischemic attack; NIHSS, the National Institutes of Health Stroke Scale; MCA, middle cerebral artery; mTICI, modified Thrombolysis in Cerebral Infarction; RPG: random plasma glucose; SHR, stress hyperglycemia ratio

Supplementary Table 3. Predictive factors for the development of 90-day death on training dataset

| Variables | Univariate analysis OR (95%CI) | P value | Multivariate analysis OR (95%CI), model 1 | P value | Multivariate analysis OR (95%CI), model 2 | P value |
|------------------------------------|-----------------------------------|---------|--|---------|--|---------|
| Age | 1.04(1.02-1.07) | 0.001 | 1.03(1.01-1.06) | 0.018 | 1.03(1.00-1.05) | 0.02 |
| Male | 1.15(0.67-1.96) | 0.62 | | | | |
| Hypertension | 1.16(0.68-1.98) | 0.585 | | | | |
| Hyperlipidemia | 1.94(0.67-5.65) | 0.223 | | | | |
| Atrial fibrillation/Atrial flutter | 1.30(0.76-2.23) | 0.337 | | | | |
| Previous ischemic stroke/TIA | 1.33(0.63-2.83) | 0.458 | 1.04(0.44-2.46) | 0.92 | 1.08(0.46-1.53) | 0.87 |
| Valvular heart disease | 1.20(0.53-2.74) | 0.658 | | | | |
| Smoking | 0.72(0.40-1.32) | 0.291 | | | | |
| Alcohol consumption | 0.78(0.35-1.75) | 0.552 | | | | |
| Diabetes mellitus | 1.27(0.69-2.34) | 0.449 | 0.85(0.37-1.99) | 0.71 | 0.97(0.43-2.22) | 0.95 |
| Onset to admission time | 0.99(0.96-1.02) | 0.387 | | | | |
| Baseline NIHSS | 1.09(1.05-1.13) | <0.001 | 1.07(1.02-1.11) | 0.0018 | 1.07(1.02-1.11) | 0.0017 |
| TOAST classification | | | | | | |
| Large artery atherosclerosis | 3.71(1.06-13.04) | 0.041 | | | | |
| Cardio embolism | 4.71(1.37-16.23) | 0.014 | | | | |
| Others | 3.57(0.97-13.07) | 0.055 | | | | |
| Small-artery occlusion | Reference | | Reference | | Reference | |
| Occlusion site | | | | | | |
| Carotid occlusion | 3.61(1.89-6.90) | <0.001 | 1.88(0.87-4.06) | 0.11 | 1.71(0.78-3.77) | 0.18 |
| MCA occlusion (M1-M2) | 1.04(0.53-2.03) | 0.908 | 0.62(0.27-1.40) | 0.25 | 0.59(0.26-1.35) | 0.21 |
| No record or other | Reference | | Reference | | Reference | |
| Endovascular treatment | 1.36(0.70-2.63) | 0.359 | 1.16(0.50-2.68) | 0.73 | 1.27(0.55-2.91) | 0.57 |
| Intravenous thrombolysis | 0.56(0.23-1.36) | 0.196 | 0.53(0.21-1.36) | 0.19 | 0.53(0.20-1.36) | 0.18 |
| Antihypertensive therapy | 0.82(0.47-1.43) | 0.475 | | | | |
| Insulin | 1.45(0.70-3.00) | 0.321 | | | | |
| Oral hypoglycemic agents | 0.87(0.39-1.94) | 0.731 | | | | |
| Antiplatelet | 0.32(0.16-0.63) | 0.001 | | | | |
| Statin | 0.48(0.25-0.94) | 0.032 | | | | |
| mTICI (2b/3) | 0.77(0.14-4.28) | 0.761 | | | | |
| RPG | 1.04(0.96-1.12) | 0.314 | 1.04(0.93-1.16) | 0.45 | 1.04(0.93-1.16) | 0.47 |
| SHR(per 0.1-point increases) | 1.21(1.11-1.31) | <0.001 | 1.13(1.03-1.25) | 0.01 | | |

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| Variables | Univariate analysis OR (95%CI) | P value | Multivariate analysis OR (95%CI), model 1 | P value | Multivariate analysis OR (95%CI), model 2 | P value |
|---------------------|-----------------------------------|---------|--|---------|--|---------|
| SHR (≥ 1.25) | 4.64(2.59-8.31) | <0.001 | | | 2.79(1.42-5.49) | 0.003 |

OR, odds ratio; CI, confidence interval; TIA, transient ischemic attack; NIHSS, the National Institutes of Health Stroke Scale; TOAST, Trial of Org 10172 in Acute Stroke Treatment; MCA, middle cerebral artery; mTICI, modified Thrombolysis in Cerebral Infarction; RPG: random plasma glucose; SHR, stress hyperglycemia ratio

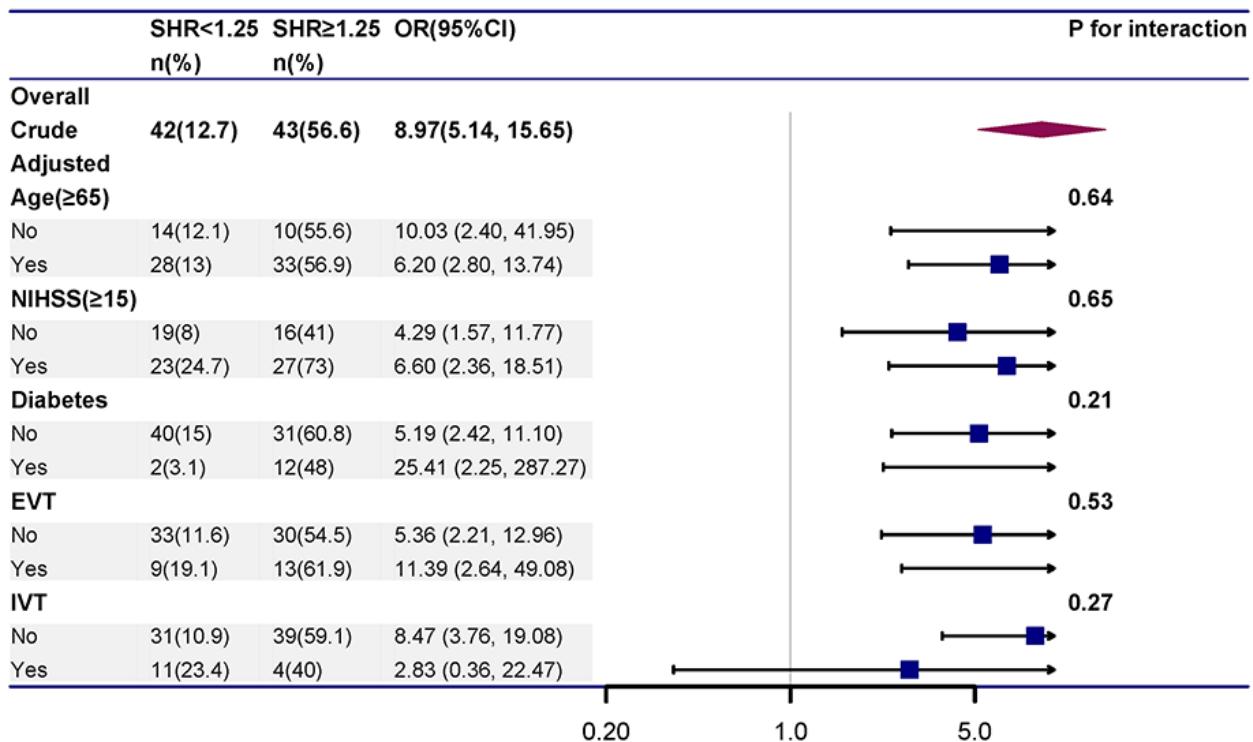
Supplementary Table 4. Predictive performance of dichotomized SHR (≥ 1.25 versus < 1.25) on training dataset and validation dataset

| | moderate-to-severe CED | 90-day poor functional outcome (mRS >2) | 90-day death |
|--------------------|---------------------------|--|--------------|
| Training dataset | | | |
| Sensitivity (%) | 50.59 | 29.21 | 42.42 |
| Specificity (%) | 89.75 | 92.97 | 86.29 |
| PPV (%) | 56.58 | 81.94 | 38.89 |
| NPV (%) | 87.31 | 54.6 | 87.94 |
| Accuracy (%) | 81.57 | 59.69 | 78.81 |
| Validation dataset | | | |
| Sensitivity (%) | 49.09 | 41.41 | 44.44 |
| Specificity (%) | 76.70 | 82.88 | 73.77 |
| PPV (%) | 39.71 | 68.33 | 20 |
| NPV (%) | 82.82 | 61.33 | 90 |
| Accuracy (%) | 70.13 | 63.33 | 70 |

CED: Cerebral edema; PPV: Positive Predictive Value; NPV: Negative Predictive Value

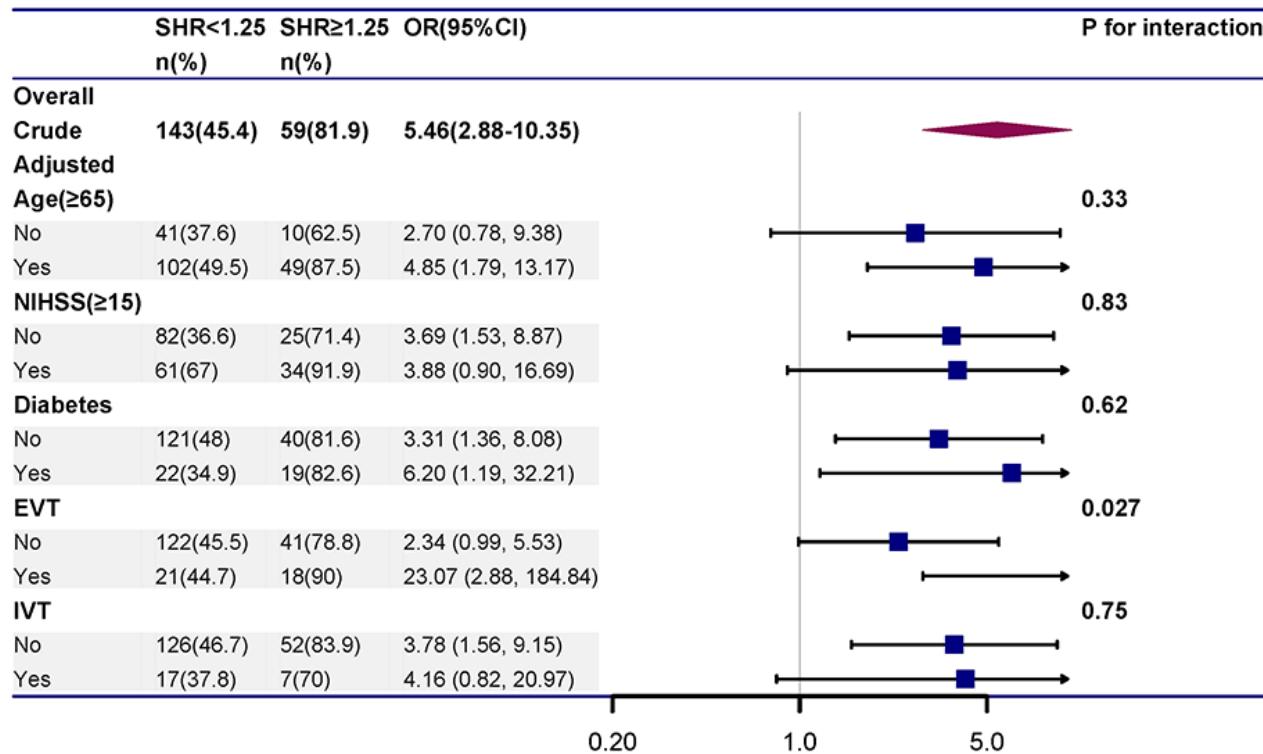
1.1 Supplementary Figure

Supplementary Figure 1. Association between the stress hyperglycemia ratio (SHR) and moderate-to-severe cerebral edema (training dataset).



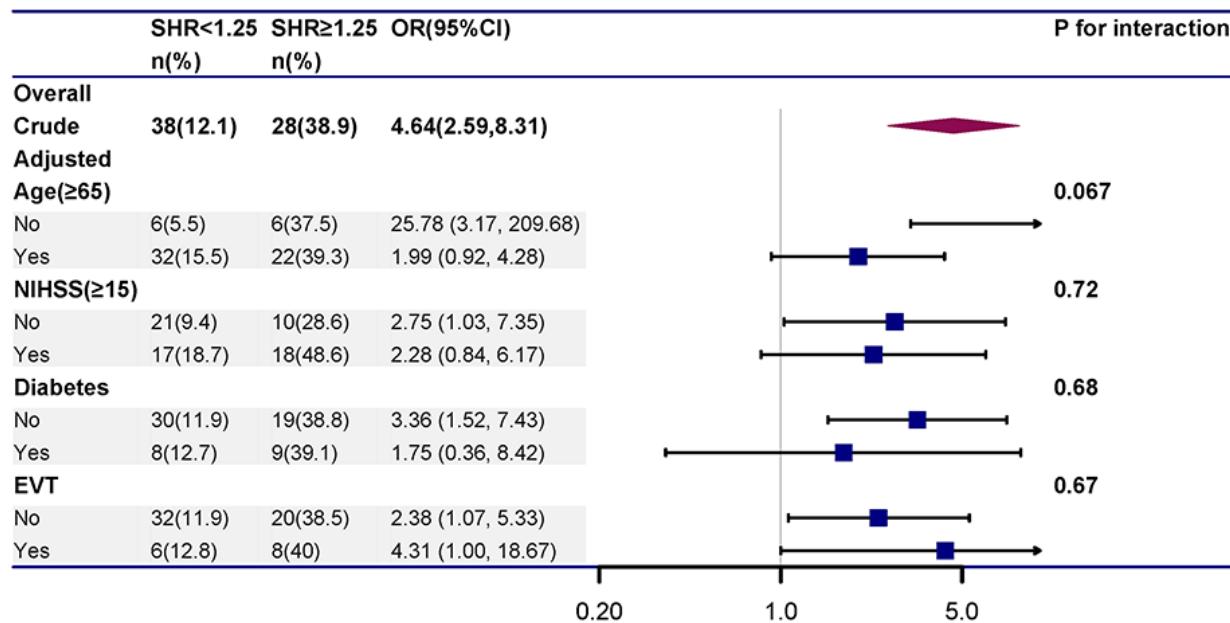
The following covariates expect for the stratified variables were adjusted as in model 2(Supplementary table 2): age, the National Institutes of Health Stroke Scale (NIHSS) score, diabetes, endovascular treatment, intravenous thrombolysis, occlusion site, and admission plasma glucose. SHR: stress hyperglycemia ratio, OR: odds ratio, CI, confidence interval; EVT: endovascular treatment, IVT: intravenous thrombolysis

Supplementary Figure 2. Association between the stress hyperglycemia ratio (SHR) and 90-day poor functional outcome (training dataset)



The following covariates expect for the stratified variables were adjusted as in model 2(table 3): age, the National Institutes of Health Stroke Scale (NIHSS) score, diabetes, endovascular treatment, intravenous thrombolysis, occlusion site, previous stroke/transient ischemic attack, and admission plasma glucose. SHR: stress hyperglycemia ratio, OR: odds ratio, CI, confidence interval; EVT: endovascular treatment, IVT: intravenous thrombolysis

Supplementary Figure 3. Association between stress hyperglycemia ratio (SHR) and 90-day death (training dataset)



The following covariates expect for the stratified variables were adjusted as in model 2(Supplementary table 3): age, the National Institutes of Health Stroke Scale (NIHSS) score, diabetes, endovascular treatment, intravenous thrombolysis, occlusion site, previous stroke/transient ischemic attack, and admission plasma glucose. SHR: stress hyperglycemia ratio, OR: odds ratio, CI, confidence interval; EVT: endovascular treatment