**S1 Table 6.** Characteristics of idiopathic lactic acidosis cases (lactate ≥2.0 mmol/l and pH <7.35) and matched controls

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| **Idiopathic LA,**  **Lactate ≥2.0**  **mmol/l** | **Cases** | **Controls** |
| **All** | (n=28) | (n=622) |
| **Age**, median (IQR) | 73 (66 - 81) | 72 (65 - 81) |
| **Gender** |  |  |
| Male | 15 (53.6%) | 334 (53.7%) |
| Female | 13 (46.4%) | 288 (46.3%) |
| **Use of metformin** |  |  |
| Non-usea | 11 (39.3%) | 274 (44.1%) |
| Recent useb | 0 (0.0%) | 41 (6.6%) |
| Current usec | 17 (60.7%) | 307 (49.4%) |
| **Diabetes duration** | |  |
| 0-1 years | 1 (3.6%) | 67 (10.8%) |
| 2-5 years | 5 (17.9%) | 110 (17.7%) |
| 6-9 years | 3 (10.7%) | 88 (14.1%) |
| 10+ year | 19 (67.9%) | 357 (57.4%) |
| **Charlson comorbidity index** |  |  |
| 0 | 14 (50.0%) | 397 (63.8%) |
| 1 | 2 (7.1%) | 96 (15.4%) |
| ≥2 | 12 (42.9%) | 129 (20.7%) |
| **Laboratory values** |  |  |
| **eGFR**d,e, mean (SD) | 60·8 (26.1) | 70·0 (22.6) |
| **HbA1c**g,f, mean (SD) | 7·0 (1.2) | 7·0 (1.2) |

a. Non-use of metformin is “never use of metformin or occurrence of a metformin prescription dated more than 365 days before admission with lactic acidosis”.

b. Recent use of metformin is “occurrence of a metformin prescription in the past dated 91 to 365 days before admission with lactic acidosis”.

c. Current use is “occurrence of a metformin prescription dated within the past 90 days before admission with lactic acidosis”.

d. Two cases (7.1%) and 111 controls (17.8%) had a missing value for eGFR.

e. eGFR calculated by the MDRD formula.

f. Two cases (7.1%) and 108 controls (17.4%) had a missing value for HbA1c.

g. HbA1c (glycated hemoglobin), % of total hemoglobin.