Additional table 1. Clinical description of patients with VUS

| **Genetic variant** | **Treatment status** | **Major organ involvement** | **Signs and symptoms** | **Typical signs of FD** | **Comorbidity** | **Agea (years)** | **Time since *GLA* identification** |
| --- | --- | --- | --- | --- | --- | --- | --- |
| R118C c.352C>T | Untreated | PNS | Acroparesthesias | - | - | 29 | 4y 3m |
| R118C c.352C>T | Untreated | - | - | - | - | 38 | 3y 5m |
| R118C c.352C>T | Treated | Cardiac, renal, GI, CV, PNS | LVH, pain, vertigo, dizziness, diarrhoea, constipation, white matter lesion on MRI, TIA, albuminuria, anxiety | - | Carpal tunnel syndrome, obesity | 48 | 4y 1m |
| A143T c.427G>A | Untreated | - | Cataract  | Angiokeratoma | AHT | 72 | 1y 9m |
| A143T c.427G>A | Untreated | Cardiac | LVH, cardiac failure, atrial fibrillation, cataract, dyspnoea, RET | - | Hypothyroidism | 63 | 2m |
| S126G c.376A>G | Treated | CV, PNS | TIA, acroparesthesias, depression, anxiety | - | Heterozygosity for the factor V Leiden, AHT | 49 | 4y 8m |
| aAt study inclusionAbbreviations: AHT, arterial hypertension; CV, cerebrovascular; FD, Fabry disease; GI, gastrointestinal; m, month; MRI, magnetic resonance imaging; PNS, peripheral nervous system; RET, reduced exercise tolerance; TIA, transient ischemic attack; y, year. |