



**Figure S1. Correlation between median age of onset and mutated alleles detected in *SLC7A9*.** (A) Five truncating (red) and three missense alleles (black) sorted by median age of onset. Note that there is no correlation between the presence of a truncating variant and median age of onset in the affected individuals. (B) In contrast, the presence of two mutated variants in *SLC7A9* results in a significantly earlier age of onset compared to 1 mutated variant (\* $p=0.029$ , unpaired Student's t-test). NL, nephrolithiasis; NC, nephrocalcinosis; yr, years.

**Supplemental Table 1. Cohort characteristics of 272 individuals from 268 different families with NL/NC**

| General / Age of onset           |         |                                  |                                                  |                                              |                | Gender      |             | Ethnicity / Origin        |                          |           |          |             |                 |                          |
|----------------------------------|---------|----------------------------------|--------------------------------------------------|----------------------------------------------|----------------|-------------|-------------|---------------------------|--------------------------|-----------|----------|-------------|-----------------|--------------------------|
| Inclusion criterion <sup>a</sup> | Total # | Median age of onset (range - yr) | # of pediatric individuals (age of onset <18 yr) | # of adult individuals (age of onset ≥18 yr) | Con-sanguinity | Female      | Male        | Western / Northern Europe | Eastern Europe / Balkans | EA        | Asia     | Middle East | Southern Europe | South America / Hispanic |
| <b>NL</b>                        | 256     | 30 (1-81)                        | 91                                               | 165                                          | 4              | 95          | 161         | 170                       | 62                       | 17        | 4        | 1           | 1               | 1                        |
| <b>NC</b>                        | 16      | 8 (1-46)                         | 15                                               | 1                                            | 2              | 6           | 10          | 4                         | 7                        | 1         | 2        | 2           | 0               | 0                        |
| <b>Total #</b>                   | 272     | 28 (1-81)                        | 106 (39.0%)                                      | 166 (61.0%)                                  | 6 (2.2%)       | 101 (37.1%) | 171 (62.9%) | 174 (64.0%)               | 69 (25.4%)               | 18 (6.6%) | 6 (2.2%) | 3 (1.1%)    | 1 (0.4%)        | 1 (0.4%)                 |

<sup>a</sup>Note that, the inclusion criteria nephrocalcinosis and hypercalcuria refer to isolated conditions without a reported history of stone disease. Another 11 individuals, who primarily presented with kidney stone disease also showed nephrocalcinosis on renal ultrasound or CT scan. Abbreviations: EA, European American; NC, isolated nephrocalcinosis; NL, nephrolithiasis; yr, years

**Supplemental Table 2. Thirty genes, known to cause monogenic forms of NL/NC that were included in the study.**

|    | Gene Symbol <sup>a</sup> | Gene Name                                                              | Accession #    | Disease entity                                                                           | MIM-Phenotype # | Mode | Coding Exons | Ref. |
|----|--------------------------|------------------------------------------------------------------------|----------------|------------------------------------------------------------------------------------------|-----------------|------|--------------|------|
| 1  | <b><u>ADCY10/SAC</u></b> | adenylate cyclase 10 (soluble)                                         | NM_018417.4    | Idiopathic (absorptive) hypercalciuria, susceptibility                                   | 143870          | AD   | 32           | 1    |
| 2  | <b><u>AGXT</u></b>       | alanine-glyoxylate aminotransferase                                    | NM_000030.2    | Primary hyperoxaluria, type 1                                                            | 259900          | AR   | 11           | 2    |
| 3  | <b><u>APRT</u></b>       | adenine phosphoribosyltransferase                                      | NM_000485.2    | Adenine phosphoribosyltransferase deficiency, APRT                                       | 614723          | AR   | 5            | 3    |
| 4  | <b><u>ATP6V0A4</u></b>   | ATPase, H <sup>+</sup> transporting, lysosomal V0 subunit a4           | NM_020632.2    | dRTA                                                                                     | 602722          | AR   | 20           | 4    |
| 5  | <b><u>ATP6V1B1</u></b>   | ATPase, H <sup>+</sup> transporting, lysosomal 56/58kDa, V1 subunit B1 | NM_001692.3    | distal renal tubular acidosis (dRTA) with deafness                                       | 267300          | AR   | 14           | 5    |
| 6  | <b><u>CA2</u></b>        | carbonic anhydrase II                                                  | NM_000067.2    | Osteopetrosis + d/pRTA                                                                   | 259730          | AR   | 7            | 6    |
| 7  | <b><u>CASR</u></b>       | calcium-sensing receptor                                               | NM_001178065.1 | Hypocalcemia with Bartter syndrome / hypocalcemia, autosomal dominant                    | 601198          | AD   | 6            | 7    |
| 8  | <b><u>CLCN5</u></b>      | chloride channel, voltage-sensitive 5                                  | NM_001127898.3 | Dent disease / Nephrolithiasis, type 1                                                   | 300009 / 310468 | XR   | 14           | 8    |
| 9  | <b><u>CLCNKB</u></b>     | chloride channel, voltage-sensitive Kb                                 | NM_000085.4    | Bartter syndrome, type 3                                                                 | 607364          | AR   | 19           | 9    |
| 10 | <b><u>CLDN16</u></b>     | claudin 16                                                             | NM_006580.3    | Familial hypomagnesemia with hypercalciuria & nephrocalcinosis, FHHNC                    | 248250          | AR   | 5            | 10   |
| 11 | <b><u>CLDN19</u></b>     | claudin 19                                                             | NM_001123395.1 | Familial hypomagnesemia with hypercalciuria & nephrocalcinosis with ocular abnormalities | 248190          | AR   | 4            | 11   |
| 12 | <b><u>CYP24A1</u></b>    | cytochrome P450, family 24, subfamily A, polypeptide 1                 | NM_000782.4    | 1,25-(OH) D-24 hydroxylase deficiency , infantile Hypercalcemia                          | 143880          | AR   | 11           | 12   |
| 13 | <b><u>FAM20A</u></b>     | family with sequence similarity 20, member A                           | NM_017565.3    | Enamel-Renal syndrome, amelogenesis imperfect and nephrocalcinosis                       | 204690          | AR   | 12           | 13   |
| 14 | <b><u>GRHPR</u></b>      | glyoxylate reductase/hydroxypyruvate reductase                         | NM_012203.1    | Primary hyperoxaluria, type 2                                                            | 260000          | AR   | 9            | 14   |
| 15 | <b><u>HNF4A</u></b>      | hepatocyte nuclear factor 4, alpha                                     | NM_000457.4    | MODY + Fanconi syndrome + Nephrocalcinosis (p.R76W)                                      | 125850          | AD   | 1            | 15   |
| 16 | <b><u>HOGA1</u></b>      | 4-hydroxy-2-oxoglutarate aldolase 1                                    | NM_138413.3    | Primary hyperoxaluria, type 3                                                            | 613616          | AR   | 7            | 16   |
| 17 | <b><u>HPRT1</u></b>      | hypoxanthine phosphoribosyltransferase 1                               | NM_000194.2    | Kelley-Seegmiller syndrome, partial HPRT deficiency, HPRT-related gout                   | 300323          | XR   | 9            | 17   |
| 18 | <b><u>KCNJ1</u></b>      | potassium inwardly-rectifying channel, subfamily J, member 1           | NM_000220.4    | Bartter syndrome, type 2                                                                 | 241200          | AR   | 2            | 18   |
| 19 | <b><u>OCRL</u></b>       | oculocerebrorenal syndrome of Lowe                                     | NM_000276.3    | Lowe syndrome / Dent disease 2                                                           | 309000 / 300555 | XR   | 24           | 19   |
| 20 | <b><u>SLC12A1</u></b>    | solute carrier family 12, member 1                                     | NM_000338.2    | Bartter syndrome, type 1                                                                 | 601678          | AR   | 27           | 20   |

|    |                        |                                                                                                                                                          |                |                                                                                           |                 |       |    |    |
|----|------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------|----------------|-------------------------------------------------------------------------------------------|-----------------|-------|----|----|
| 21 | <u><b>SLC22A12</b></u> | solute carrier family 22 (organic anion/urate transporter), member 12                                                                                    | NM_144585.3    | Renal hypouricemia, RHUC1                                                                 | 220150          | AD/AR | 10 | 21 |
| 22 | <u><b>SLC2A9</b></u>   | solute carrier family 2 (facilitated glucose transporter), member 9                                                                                      | NM_001001290.1 | Renal hypouricemia, RHUC2                                                                 | 612076          | AD/AR | 13 | 22 |
| 23 | <u><b>SLC34A1</b></u>  | solute carrier family 34 (sodium phosphate), member 1                                                                                                    | NM_003052.4    | Hypophosphatemic nephrolithiasis/osteoporosis-1, NPHLOP1 / Fanconi renotubular syndrome 2 | 612286 / 613388 | AD/AR | 13 | 23 |
| 24 | <u><b>SLC34A3</b></u>  | solute carrier family 34 (sodium phosphate), member 3                                                                                                    | NM_001177316.1 | Hypophosphatemic rickets with hypercalciuria                                              | 241530          | AR    | 12 | 24 |
| 25 | <u><b>SLC3A1</b></u>   | solute carrier family 3 (cystine, dibasic and neutral amino acid transporters, activator of cystine, dibasic and neutral amino acid transport), member 1 | NM_000341.3    | Cystinuria, type A                                                                        | 220100          | AR    | 10 | 25 |
| 26 | <u><b>SLC4A1</b></u>   | solute carrier family 4, anion exchanger, member 1 (erythrocyte membrane protein band 3, Diego blood group)                                              | NM_000342.3    | Primary distal renal tubular acidosis, dominant / recessive                               | 179800 / 611590 | AD/AR | 19 | 26 |
| 27 | <u><b>SLC7A9</b></u>   | solute carrier family 7 (glycoprotein-associated amino acid transporter light chain, bo,+ system), member 9                                              | NM_014270.4    | Cystinuria, type B                                                                        | 220100          | AD/AR | 12 | 27 |
| 28 | <u><b>SLC9A3R1</b></u> | solute carrier family 9, subfamily A (NHE3, cation proton antiporter 3), member 3 regulator 1                                                            | NM_004252.4    | Hypophosphatemic nephrolithiasis/osteoporosis-2, NPHLOP2                                  | 612287          | AD    | 6  | 28 |
| 29 | <b>VDR</b>             | vitamin D (1,25- dihydroxyvitamin D3) receptor                                                                                                           | NM_000376.2    | Idiopathic hypercalciuria                                                                 | 277440          | AD    | 11 | 29 |
| 30 | <b>XDH</b>             | xanthine dehydrogenase                                                                                                                                   | NM_000379.3    | Xanthinuria, type 1                                                                       | 278300          | AR    | 36 | 30 |

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<sup>a</sup>Gene symbols are underlined whenever putatively causative mutations were detected in the present study. For HNF4A the MIM-phenotype number denotes MODY type 1, as occurrence of Fanconi syndrome and NC has only been shown in the presence of a specific allele (p.R76W).

Abbreviations: AD, autosomal dominant; AR, autosomal recessive; NC, isolated nephrocalcinosis; NL, nephrolithiasis; Ref., reference; XR, x-chromosomal recessive.

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**Supplemental Table 3. Coverage and variant statistics for 336 DNA samples and 428 amplicons representing the coding exons of 30 genes known to be mutated in NL/NC/HC.**

### Coverage statistics <sup>a</sup>

|                                       | Total #     | Percentage |
|---------------------------------------|-------------|------------|
| <b>Median coverage per individual</b> | <b>261x</b> | -          |
| - individuals with coverage 0         | 0           | 0.0%       |
| - individuals with coverage <10x      | 1           | 0.3%       |
| - individuals with coverage <20x      | 1           | 0.3%       |
| - individuals with coverage <100x     | 4           | 1.2%       |
| <b>Median coverage per amplicon</b>   | <b>264x</b> | -          |
| - amplicons with coverage 0           | 9           | 2.1%       |
| - amplicons with coverage <10x        | 14          | 3.3%       |
| - amplicons with coverage <20x        | 21          | 4.9%       |
| - amplicons with coverage <50x        | 35          | 8.2%       |

<sup>a</sup> Median coverage per individual/sample was 261x. One individual/sample failed to undergo multiplex amplification (0.3%), highlighted in grey. Median coverage per amplicon reached 264x, with 14 amplicons (3.3%) showing less than 10x coverage in total (highlighted in grey). These amplicons were *FAM20A*-exon1\_1, *FAM20A*-exon1\_2, *SLC9A3R1*-exon1\_1, *SLC9A3R1*-exon1\_2, *CA2*-exon1, *CA2*-exon3, *APRT*-exon1, *SLC34A3*-exon2, *SLC34A3*-exon9, *OCRL*-exon1, *CLCNKB*-exon20, *HPRT1*-exon1, *HPRT1*-exon4, and *VDR*-exon2. Note that among failed exons, the first coding exon of a gene was highly overrepresented, presumably due to generally greater GC-content.

### Variant statistics <sup>b</sup>

|                                                      | Total #       | Percentage  |
|------------------------------------------------------|---------------|-------------|
| <b>All variants</b>                                  | <b>13,835</b> | <b>100%</b> |
| <b>SNV</b>                                           | <b>12,589</b> | <b>91%</b>  |
| <b>DIV</b>                                           | <b>1,246</b>  | <b>9%</b>   |
| <b>Retained deleterious variants after filtering</b> | <b>162</b>    | <b>1.2%</b> |
| <b>Sanger confirmed variants</b>                     | <b>139</b>    | <b>1.0%</b> |
| <b>Disease causing variants</b>                      | <b>50</b>     | <b>0.4%</b> |

<sup>b</sup> Total number of variant calls (grey) in 336 DNA samples and percentage of SNV/DIV calls after mapping to the concatenated reference sequence of 30 NL/NC/HC genes. The variant filtering process is described in the 'Methods' section. Abbreviations: SNV, single nucleotide variant; DIV, deletion/insertion variant.

**Supplemental Table S4. Clinical characteristics of 41 individuals (40 families) with newly established genetic/molecular diagnoses and their allele frequency compared to the general population.**

| Gene, Individual <sup>a</sup> | CKD-stage (ESRD at age) | Urine abnormalities | Serum abnormalities | Imaging (US/CT/KUB)                         | Other                                                   | Ethnicity     | Nucleotide change | AA change        | EVS (All alleles)      |
|-------------------------------|-------------------------|---------------------|---------------------|---------------------------------------------|---------------------------------------------------------|---------------|-------------------|------------------|------------------------|
| <b><u>SLC7A9</u></b>          |                         |                     |                     |                                             |                                                         |               |                   |                  |                        |
| B208-21                       | 1                       | cystine↑            | -                   | -                                           | -                                                       | Albanian      | c.313G>A          | p.Gly105Arg      | AA=0/AG=4/GG=6499      |
| JAS-C8                        | 1                       | -                   | uric acid↑          | 6 mm calculus RUJ                           | CaOx stone, normal urinary Ox + Ca2+, CU not quantified | White British | c.313G>A          | p.Gly105Arg      | AA=0/AG=4/GG=6499      |
| JAS-E5                        | 1                       | cystine↑            | -                   | NC                                          | urinary tract infections                                | White British | c.313G>A          | p.Gly105Arg      | AA=0/AG=4/GG=6499      |
| F1029-21                      | 1                       | -                   | -                   | solitary renal cyst, increased echogenicity | -                                                       | German        | c.313G>A          | p.Gly105Arg      | AA=0/AG=4/GG=6499      |
|                               |                         |                     |                     |                                             |                                                         |               | c.544G>A          | p.Ala182Thr      | AA=1/AG=42/GG=6460     |
| JAS-D30                       | 5 (45)                  | cystine↑            | -                   | -                                           | cystine stones including staghorn calculi               | White British | c.313G>A          | p.Gly105Arg      | AA=0/AG=4/GG=6499      |
| JAS-D31                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.614del          | p.Lys205Argfs*59 | A1A1=0/A1R=2/RR=6257   |
| JAS-D47                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.411_412del      | p.Pro139Leufs*69 | A1A1=69/A1R=78/RR=6112 |
| JAS-D34                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.411_412del      | p.Pro139Leufs*69 | A1A1=69/A1R=78/RR=6112 |
| JAS-D28                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.544G>A          | p.Ala182Thr      | AA=1/AG=42/GG=6460     |
| JAS-F41                       | 1                       | Ca2+↑               | -                   | renal calculus seen on KUB, radioopaque     | -                                                       | White British | c.544G>A          | p.Ala182Thr      | AA=1/AG=42/GG=6460     |
| JAS-F50                       | 1                       | -                   | uric acid↑          | radiolucent stone                           | cystine not quantified                                  | White British | c.544G>A          | p.Ala182Thr      | AA=1/AG=42/GG=6460     |
| JAS-D57                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.544G>A          | p.Ala182Thr      | AA=1/AG=42/GG=6460     |
|                               |                         |                     |                     |                                             |                                                         |               | c.614dup          | p.Asn206Glufs*3  | A1A1=0/A1R=2/RR=6257   |
| JAS-F87                       | 1                       | Ca2+↑               | -                   | radioopaque stones, large > 15 mm           | CaPO- stones                                            | White British | c.614dup          | p.Asn206Glufs*3  | A1A1=0/A1R=2/RR=6257   |
| JAS-G10                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          |               | c.614dup          | p.Asn206Glufs*3  | A1A1=0/A1R=2/RR=6257   |
| JAS-D29                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.671C>T          | p.Ala224Val      | -                      |
| JAS-D55                       | 1                       | cystine↑            | -                   | -                                           | cystine stones                                          | White British | c.671C>T          | p.Ala224Val      | -                      |
| JAS-F19                       | 1                       | Ca2+↑ (mild)        | PO-↓                | tiny bilateral calculi on KUB               | -                                                       | White British | c.671C>T          | p.Ala224Val      | -                      |
|                               |                         |                     |                     |                                             |                                                         |               | c.1369T>C         | p.Tyr457His      | CC=0/CT=2/TT=6499      |

|                        |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
|------------------------|---|-------------------------------|------------|----|----------------------------------------------------|---------------------------------------------|-------------------|-------------|-------------------|----------------------|
| B114-21                | 1 | cystine↑                      | -          |    | bladder stone on US                                | cystine stones                              | European American | c.814del    | p.Val272Cysfs*6   | -                    |
|                        |   |                               |            |    |                                                    |                                             |                   | c.997C>T    | p.Arg333Trp       | TT=0/TC=2/CC=6501    |
| JAS-E9                 | 1 | cystine↑                      | -          | -  |                                                    | cystine stones                              | White British     | c.1353C>A   | p.Tyr451*         | -                    |
|                        |   |                               |            |    |                                                    |                                             |                   | c.1400-2A>G | 3' splice         | -                    |
| <b><u>ADCY10</u></b>   |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| JAS-F8                 | 1 | Ca2+↑                         | -          | -  | -                                                  |                                             | Polish            | c.1263C>A   | p.Tyr421*         | -                    |
| JAS-F68                | 1 | Ca2+↑                         | -          |    | radioopaque renal calculus 6 x 4 mm – suspected NC | CaOx stone                                  | White British     | c.1282G>A   | p.Asp428Asn       | AA=0/AG=1/GG=6502    |
| JAS-F29                | 1 | -                             | PO-↓       |    | bilateral calculi on CT, 2 small kidney cysts      | normal urinary Ca                           | White British     | c.4477del   | p.Leu1493Serfs*24 | A1A1=0/A1R=3/RR=6257 |
| <b><u>SLC2A9</u></b>   |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| B179-21                | 1 | -                             | -          | -  | -                                                  |                                             | Albanian          | c.1343C>T   | p.Pro448Leu       | TT=0/TC=1/CC=6502    |
| B230-21                | 1 | fractionary uric acid↑, Ca2+↑ | uric acid↑ | -  | -                                                  |                                             | Macedonian        | c.1419+1G>A | 5' splice         | -                    |
| <b><u>SLC9A3R1</u></b> |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| B224-21                | 1 | Ca2+↑                         | -          |    | stone on US                                        | normal serum PO-                            | Macedonian        | c.673G>A    | p.Glu225Lys       | AA=0/AG=31/GG=6472   |
| B109-21                | 1 | -                             | -          |    | stone on CT                                        | Ulcerative colitis                          | European American | c.888+2T>C  | 5' splice site    | CC=0/CT=6/TT=6497    |
| <b><u>SLC22A12</u></b> |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| JAS-F98                | 1 | -                             | uric acid↑ | -  |                                                    | no HC, serum CK↑, muscle pains, CaOx stones | White British     | c.431T>C    | p.Leu144Pro       | CC=0/CT=5/TT=6493    |
| B155-12                | 1 | -                             | -          | -  |                                                    | reported Ca-stones                          | European American | c.1300C>T   | p.Arg434Cys       | TT=0/TC=1/CC=6484    |
| <b><u>SLC4A1</u></b>   |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| JAS-E8                 | 1 | Ca2+↑                         | -          | NC | -                                                  |                                             | White British     | c.2716G>C   | p.Glu906Gln       | -                    |
| <b><u>SLC3A1</u></b>   |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| JAS-B21 <sup>a</sup>   | 1 | cystine↑                      | -          | -  |                                                    | cystine stones                              | White British     | c.1354C>T   | p.Arg452Trp       | no                   |
| JAS-B22 <sup>a</sup>   | 1 | cystine↑                      | -          | -  |                                                    | cystine stones                              | White British     | c.1354C>T   | p.Arg452Trp       | no                   |
| JAS-G7                 | 1 | cystine↑                      | -          | -  |                                                    | cystine stones                              | White British     | c.1400T>C   | p.Met467Thr       | CC=0/CT=27/TT=6476   |
| <b><u>ATP6V1B1</u></b> |   |                               |            |    |                                                    |                                             |                   |             |                   |                      |
| B214-21                | 1 | Ca2+↑                         | HCO3-↓     | -  |                                                    | no known hearing problems                   | Gypsy             | c.481G>A    | p.Glu161Lys       | AA=4/AG=294/GG=6205  |



|                       |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
|-----------------------|-----------|------------------------------------|---------------------------------|----------------------|------------------------------------------|-------------------|--------------|------------------|------------------------|--|
| <u><b>CLCN5</b></u>   |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| B111-21               | 1         | -                                  | -                               | NC                   | -                                        | Filipino          | c.344G>A     | p.Trp115*        | no                     |  |
| B167-21               | 5<br>(25) | oxalate↑                           | Mg2+↓                           | NC                   | -                                        | Macedonian        | c.1009G>A    | p.Glu337Lys      | no                     |  |
| <u><b>CLDN16</b></u>  |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| JAS-C1                | 5<br>(38) | Ca2+↑                              | Mg2+↓                           | NC                   | recurrent stones                         | Arabian           | c.445C>T     | p.Arg149*        | no                     |  |
| B178-21               | 2         | Ca2+↑                              | Mg2+↓                           | NC                   | -                                        | Macedonian        | c.453G>T     | p.Leu151Phe      | no                     |  |
| <u><b>CYP24A1</b></u> |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| B223-21               | 1         | Ca2+↑                              | Mg2+↓                           | NC                   | no FTT, no infantile onset (age 12 yr)   | Albanian          | c.428_430del | p.Glu143del      | no                     |  |
| <u><b>AGXT</b></u>    |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| B106-21               | 1         | oxalate↑                           | -                               | -                    | CaOx stones, not responsive to pyridoxin | European American | c.416_418del | p.Val139del      | no                     |  |
|                       |           |                                    |                                 |                      |                                          |                   | c.846+1G>T   | 5' splice site   | no                     |  |
| <u><b>SLC34A1</b></u> |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| B168-21               | 1         | oxalate↑<br>(pyridoxine sensitive) | PO-↓, severe metabolic acidosis | NC                   | FTT, cow-milk allergy                    | Macedonian        | c.271_291del | p.Val91Ala97del7 | A1A1=1/A1R=229/RR=6029 |  |
|                       |           |                                    |                                 |                      |                                          |                   | c.1534C>T    | p.Arg512Cys      | no                     |  |
| <u><b>SLC34A3</b></u> |           |                                    |                                 |                      |                                          |                   |              |                  |                        |  |
| JAS-F43               | 1         | -                                  | PO-↓                            | bilat. calculi on US | -                                        | White British     | c.1454G>A    | p.Arg485His      | AA=0/AG=18/GG=6469     |  |
|                       |           |                                    |                                 |                      |                                          |                   | c.1585A>T    | p.Ile529Phe      | TT=0/TA=15/AA=6341     |  |

<sup>a</sup>JAS-B21 and JAS-B22 are siblings (same family). All other listed individuals are unrelated.

Abbreviations: AA, amino acid; AKI, acute kidney injury; bilat., bilateral; Ca, calcium; CaOx, calcium oxalate; CaPO, calcium phosphate; CKD, chronic kidney disease; CT, computed tomography; CU, cystinuria; ESRD, end-stage renal disease; EVS, Exome variant sever (<http://evs.gs.washington.edu/EVS/>); FTT, failure to thrive; HC, hypercalciuria; HO, hyperoxaluria; HCO3-, bicarbonate; HU, hypouricemia; KUB, x-ray of kidney, ureter, and bladder; Mg2+, magnesium; NC, nephrocalcinosis; NL, nephrolithiasis; PO-, phosphate; PH, primary hyperoxaluria; RUJ, right pelvicalyceal-ureteric junction; RTX, renal transplant; US, ultrasound; yr, years.