



Genetics for people

»» Genetic Testing Detoxification



My *Detox*

SUMMARY OF RESULTS

The following table shows the variants or haplotypes with non-standard metabolism detected in the patient:

| Gene | Metabolic process | State | Haplotype | Metabolizer |
|---------|-------------------|--------|---------------|--------------|
| APOE | Transport | E2E4 | Heterocigosys | INTERMEDIATE |
| CAT | Oxidative stress | *1*2 | Heterocigosys | INTERMEDIATE |
| CDA | Oxidative stress | *1*2 | Heterocigosys | SLOW |
| CYP1A2 | Cytochrome P450 | *1*1M | Heterocigosys | RAPID |
| CYP1B1 | Cytochrome P450 | *3*3 | Homocigosys | SLOW |
| CYP2B6 | Cytochrome P450 | *5*6 | Heterocigosys | INTERMEDIATE |
| CYP2C19 | Cytochrome P450 | *1*17 | Heterocigosys | RAPID |
| CYP2C8 | Cytochrome P450 | *1*3 | Heterocigosys | INTERMEDIATE |
| CYP2C9 | Cytochrome P450 | *1*2 | Heterocigosys | INTERMEDIATE |
| CYP2D6 | Cytochrome P450 | *4*10 | Heterocigosys | SLOW |
| CYP3A4 | Cytochrome P450 | *1*22 | Heterocigosys | INTERMEDIATE |
| CYP3A5 | Cytochrome P450 | *3*3 | Homocigosys | SLOW |
| CYP4F2 | Cytochrome P450 | *2*3 | Heterocigosys | INTERMEDIATE |
| GSTM1 | Glutathione | *1*D | Heterocigosys | SLOW |
| MTHFR | Methylation | *1*2 | Heterocigosys | INTERMEDIATE |
| NAT1 | Acetylation | *4*11A | Heterocigosys | RAPID |
| NAT2 | Acetylation | *6A*6A | Homocigosys | SLOW |
| OGG1 | Oxidative stress | *1*2 | Heterocigosys | INTERMEDIATE |
| SLC15A2 | Transport | *1*2 | Heterocigosys | INTERMEDIATE |
| SLCO1B1 | Transport | *1*14 | Heterocigosys | RAPID |
| MNSOD | Oxidative stress | *1*2 | Heterocigosys | INTERMEDIATE |
| UGT1A1 | Glucuronidation | *1*28 | Heterocigosys | INTERMEDIATE |
| UGT1A3 | Glucuronidation | *1*2 | Heterocigosys | INTERMEDIATE |
| UGT1A7 | Glucuronidation | *3*11 | Heterocigosys | INTERMEDIATE |
| UGT2B15 | Glucuronidation | *1*2 | Heterocigosys | INTERMEDIATE |
| UGT2B7 | Glucuronidation | *2*2 | Homocigosys | SLOW |

DETAILED RESULTS

The following table shows all genotypes included in this study, as well as their haplotype and corresponding enzyme activity:

| PHASE I | | | | | |
|--------------------|-------------------|-------------------|---------------------|-------------------|------------------|
| Gene | Metabolic Process | Variants Analyzed | Reference Haplotype | Patient Haplotype | Metabolizer Type |
| CYP1A1 | Cytochrome P450 | 6 | *1*1 | *1*1 | NORMAL |
| CYP1A2 | Cytochrome P450 | 25 | *1A*1A | *1*1M | RAPID |
| CYP1B1 | Cytochrome P450 | 12 | *1*1 | *3*3 | SLOW |
| CYP2A6 | Cytochrome P450 | 16 | *1*1 | *1*1 | NORMAL |
| CYP2B6 | Cytochrome P450 | 25 | *1*1 | *5*6 | INTERMEDIATE |
| CYP2C8 | Cytochrome P450 | 11 | *1*1 | *1*3 | INTERMEDIATE |
| CYP2C9 | Cytochrome P450 | 49 | *1*1 | *1*2 | INTERMEDIATE |
| CYP2C19 | Cytochrome P450 | 30 | *38*38 | *1*17 | RAPID |
| CYP2D6 | Cytochrome P450 | 86 | *1*1 | *4*10 | SLOW |
| CYP2E1 | Cytochrome P450 | 9 | *1*1 | *1*1 | NORMAL |
| CYP3A4 | Cytochrome P450 | 29 | *1*1 | *1*22 | INTERMEDIATE |
| CYP3A5 | Cytochrome P450 | 4 | *1*1 | *3*3 | SLOW |
| CYP3A7 | Cytochrome P450 | 8 | *1A*1A | *1A*1A | NORMAL |
| CYP4B1 | Cytochrome P450 | 4 | *1*1 | *1*1 | NORMAL |
| CYP4F2 | Cytochrome P450 | 3 | *1*1 | *2*3 | INTERMEDIATE |
| TBXAS1 (CYP5A1) | Cytochrome P450 | 4 | *1*1 | *1*1 | NORMAL |
| CYP19A1 | Cytochrome P450 | 4 | *1*1 | *1*1 | NORMAL |
| PTGIS (CYP8A1) | Cytochrome P450 | 1 | *1*1 | *1*1 | NORMAL |
| FMO1 | Cytochrome P450 | 2 | *1*1 | *1*1 | NORMAL |
| POR (CYPOR) | Cytochrome P450 | 17 | *1*1 | *1*1 | NORMAL |
| INTERMEDIATE PHASE | | | | | |
| Gene | Metabolic Process | Variants Analyzed | Reference Haplotype | Patient Haplotype | Metabolizer Type |
| CAT | Oxidative stress | 1 | *1*1 | *1*2 | INTERMEDIATE |
| CDA | Oxidative stress | 1 | *1*1 | *1*2 | SLOW |
| DPYD | Oxidative stress | 15 | *1*1 | *1*1 | NORMAL |
| G6PD | Oxidative stress | 90 | B | BB | NORMAL |
| OGG1 | Oxidative stress | 1 | *1*1 | *1*2 | INTERMEDIATE |
| SOD1 | Oxidative stress | 2 | *1*1 | *1*1 | NORMAL |
| MNSOD (SOD2) | Oxidative stress | 1 | *1*1 | *1*2 | INTERMEDIATE |

| PHASE II | | | | | |
|----------|-------------------|-------------------|---------------------|-------------------|------------------|
| Gene | Metabolic Process | Analyzed Variants | Reference Haplotype | Patient Haplotype | Metabolizer Type |
| NAT1 | Acetylation | 18 | *4*4 | *4*11A | RAPID |
| NAT2 | Acetylation | 28 | *4*4 | *6A*6A | SLOW |
| UGT1A1 | Glucuronidation | 3 | *1*1 | *1*28 | INTERMEDIATE |
| UGT1A3 | Glucuronidation | 6 | *1*1 | *1*2 | INTERMEDIATE |
| UGT1A4 | Glucuronidation | 7 | *1A*1A | *1A*1B | NORMAL |
| UGT1A7 | Glucuronidation | 2 | *1*1 | *3*11 | INTERMEDIATE |
| UGT1A8 | Glucuronidation | 1 | *1*1 | *1*1 | NORMAL |
| UGT1A9 | Glucuronidation | 3 | *1*1 | *1*1 | NORMAL |
| UGT1A10 | Glucuronidation | 4 | *1*1 | *1*1 | NORMAL |
| UGT2B7 | Glucuronidation | 2 | *1*1 | *2*2 | SLOW |
| UGT2B15 | Glucuronidation | 4 | *1*1 | *1*2 | INTERMEDIATE |
| GSTM1 | Glutathione | 1 | | *1*D | SLOW |
| GSTM3 | Glutathione | 1 | *1*1 | *1*1 | NORMAL |
| GSTP1 | Glutathione | 2 | *1A*1A | *1A*1A | NORMAL |
| GSTT1 | Glutathione | 1 | | *1*1 | NORMAL |
| COMT | Methylation | 1 | *1*1 | *1*1 | NORMAL |
| MTHFR | Methylation | 2 | *1*1 | *1*2 | INTERMEDIATE |
| TPMT | Methylation | 2 | *1*1 | *1*1 | NORMAL |
| NUDT15 | Methylation | 3 | *1*1 | *1*1 | NORMAL |
| SULT1A1 | Sulphation | 2 | *1*1 | *1*1 | NORMAL |
| SULT1A2 | Sulphation | 2 | *1*1 | *1*1 | NORMAL |
| OTHERS | | | | | |
| Gene | Metabolic Process | Analyzed Variants | Reference Haplotype | Patient Haplotype | Metabolizer Type |
| APOE | Transport | 4 | *E3*E3 | E2E4 | INTERMEDIATE |
| CFTR | Transport | 35 | *1*1 | *1*1 | NORMAL |
| SLC15A2 | Transport | 3 | *1*1 | *1*2 | INTERMEDIATE |
| SLCO1B1 | Transport | 14 | *1*1 | *1*14 | RAPID |

CONCLUSIONS

APOE: E2E4 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CAT: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CDA: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate.

CYP1A2: *1*1M haplotype has been detected in patient's sample. This haplotype has increased activity and, therefore, upper metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme, as there is a greater likelihood of not obtaining the expected therapeutic results.

CYP1B1: *3*3 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP2B6: *5*6 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP2C19: *1*17 haplotype has been detected in patient's sample. This haplotype has increased activity and, therefore, upper metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme, as there is a greater likelihood of not obtaining the expected therapeutic results.

CYP2C8: *1*3 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP2C9: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP2D6: *4*10 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP3A4: *1*22 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP3A5: *3*3 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

CYP4F2: *2*3 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

GSTM1: *1*D haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate.

MTHFR: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

NAT1: *4*11A haplotype has been detected in patient's sample. This haplotype has increased activity and, therefore, upper metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme, as there is a greater likelihood of not obtaining the expected therapeutic results.

NAT2: *6A*6A haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects due to liver toxicity.

OGG1: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

SLC15A2: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate.

SLCO1B1: *1*14 haplotype has been detected in patient's sample. This haplotype has increased activity and, therefore, upper metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme, as there is a greater likelihood of not obtaining the expected therapeutic results.

MNSOD: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

UGT1A1: *1*28 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

UGT1A3: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

UGT1A7: *3*11 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

UGT2B15: *1*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

UGT2B7: *2*2 haplotype has been detected in patient's sample. This haplotype has reduced activity and, therefore, lower metabolic rate. It is therefore recommended to avoid the use of drugs or substances that are metabolised by this enzyme as there is a greater likelihood of adverse effects.

ANNEX 1: GENES AND VARIANTS ANALYSED

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|------|-------------|----------------------|--------------------|--------|-------------|----------------------|--------------------|
| APOE | rs7412 | CC | CT | CYP2E1 | rs2070676 | GG | CC |
| APOE | rs429358 | TT | TC | CYP2E1 | rs3813867 | GG | GG |
| APOE | rs11083750 | CC | CC | CYP2E1 | rs6413419 | GG | GG |
| APOE | rs267606664 | GG | GG | CYP2E1 | rs6413420 | GG | GG |
| CAT | rs1001179 | CC | CT | CYP2E1 | rs55897648 | GG | GG |
| CDA | rs2072671 | AA | AC | CYP2E1 | rs72559710 | GG | GG |
| CFTR | rs11971167 | GG | GG | CYP3A4 | rs2242480 | CC | CC |
| CFTR | rs74503330 | GG | GG | CYP3A4 | rs2246709 | AA | GG |
| CFTR | rs74551128 | CC | CC | CYP3A4 | rs2740574 | CC | TT |
| CFTR | rs75039782 | CC | CC | CYP3A4 | rs3735451 | TT | TT |
| CFTR | rs75527207 | GG | GG | CYP3A4 | rs4646437 | GG | GG |
| CFTR | rs75541969 | GG | GG | CYP3A4 | rs4646440 | GG | GG |
| CFTR | rs76151804 | AA | AA | CYP3A4 | rs4986907 | CC | CC |
| CFTR | rs77834169 | CC | CC | CYP3A4 | rs4986908 | CC | CC |
| CFTR | rs77932196 | GG | GG | CYP3A4 | rs4986909 | GG | GG |
| CFTR | rs78655421 | GG | GG | CYP3A4 | rs4986910 | AA | AA |
| CFTR | rs78769542 | GG | GG | CYP3A4 | rs4986910 | AA | AA |
| CFTR | rs80224560 | GG | GG | CYP3A4 | rs4986913 | GG | GG |
| CFTR | rs80282562 | GG | GG | CYP3A4 | rs4987161 | AA | AA |
| CFTR | rs113993958 | GG | GG | CYP3A4 | rs12721627 | GG | GG |
| CFTR | rs115545701 | CC | CC | CYP3A4 | rs12721629 | GG | GG |
| CFTR | rs121908752 | TT | TT | CYP3A4 | rs12721634 | AA | AA |
| CFTR | rs121908753 | GG | GG | CYP3A4 | rs28371759 | AA | AA |
| CFTR | rs121908755 | GG | GG | CYP3A4 | rs35599367 | GG | GA |
| CFTR | rs121908757 | AA | AA | CYP3A4 | rs55785340 | AA | AA |
| CFTR | rs121909005 | TT | TT | CYP3A4 | rs55901263 | GG | GG |
| CFTR | rs121909013 | GG | GG | CYP3A4 | rs55951658 | TT | TT |
| CFTR | rs121909020 | GG | GG | CYP3A4 | rs56324128 | CC | CC |
| CFTR | rs121909041 | TT | TT | CYP3A4 | rs57409622 | GG | GG |
| CFTR | rs150212784 | TT | TT | CYP3A4 | rs67784355 | GG | GG |
| CFTR | rs186045772 | TT | TT | CYP3A4 | rs72552798 | CC | CC |
| CFTR | rs193922525 | GG | GG | CYP3A4 | rs72552799 | CC | CC |
| CFTR | rs202179988 | CC | CC | CYP3A4 | rs113667357 | TT | TT |
| CFTR | rs267606723 | GG | GG | CYP3A4 | rs138105638 | GG | GG |
| CFTR | rs368505753 | CC | CC | CYP3A4 | rs201821708 | TT | TT |
| CFTR | rs397508256 | GG | GG | CYP3A5 | rs776746 | TT | CC |
| CFTR | rs397508288 | AA | AA | CYP3A5 | rs10264272 | CC | CC |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|-------------|----------------------|--------------------|---------|------------|----------------------|--------------------|
| CFTR | rs397508387 | GG | GG | CYP3A5 | rs41303343 | DD | DD |
| CFTR | rs397508442 | CC | CC | CYP3A5 | rs55817950 | GG | GG |
| CFTR | rs397508759 | GG | GG | CYP3A7 | rs2257401 | GG | GG |
| CFTR | rs397508761 | AA | AA | CYP3A7 | rs11568824 | CC | CC |
| COMT | rs4680 | GG | GG | CYP3A7 | rs11568825 | AA | AA |
| CYP1A1 | rs1048943 | AA | TT | CYP3A7 | rs28451617 | CC | CC |
| CYP1A1 | rs1799814 | GG | GG | CYP3A7 | rs45446698 | TT | TT |
| CYP1A1 | rs28399430 | GG | GG | CYP3A7 | rs45465393 | GG | GG |
| CYP1A1 | rs41279188 | GG | GG | CYP3A7 | rs45494802 | AA | AA |
| CYP1A1 | rs56240201 | GG | GG | CYP3A7 | rs45575938 | AA | AA |
| CYP1A1 | rs56313657 | CC | CC | CYP3A7 | rs55798860 | CC | CC |
| CYP1A2 | rs2069514 | GG | GG | CYP4B1 | rs2297809 | CC | CC |
| CYP1A2 | rs2069526 | TT | TT | CYP4B1 | rs2297810 | GG | GG |
| CYP1A2 | rs2470890 | TT | CT | CYP4B1 | rs4646487 | CC | CC |
| CYP1A2 | rs2472304 | GG | GA | CYP4B1 | rs4646491 | CC | CC |
| CYP1A2 | rs4646425 | CC | CC | CYP4B1 | rs45467195 | AA | AA |
| CYP1A2 | rs4646427 | TT | TT | CYP4F2 | rs2108622 | CC | CT |
| CYP1A2 | rs12720461 | CC | CC | CYP4F2 | rs3093153 | CC | CC |
| CYP1A2 | rs16972381 | GG | GG | CYP4F2 | rs3093200 | GG | GG |
| CYP1A2 | rs17861157 | CC | CC | CYP8A1 | rs5622 | AA | AA |
| CYP1A2 | rs28399424 | CC | CC | CYP19A1 | rs700519 | GG | GG |
| CYP1A2 | rs35694136 | II | II | CYP19A1 | rs2236722 | AA | AA |
| CYP1A2 | rs35796837 | GG | GG | CYP19A1 | rs28757184 | GG | GG |
| CYP1A2 | rs45486893 | CC | CC | CYP19A1 | rs56658716 | AA | AA |
| CYP1A2 | rs55889066 | GG | GG | DPYD | rs1801158 | CC | CC |
| CYP1A2 | rs56107638 | GG | GG | DPYD | rs1801159 | TT | TT |
| CYP1A2 | rs56160784 | CC | CC | DPYD | rs1801160 | CC | CC |
| CYP1A2 | rs56276455 | GG | GG | DPYD | rs1801265 | AA | AA |
| CYP1A2 | rs59567621 | TT | TT | DPYD | rs1801266 | GG | GG |
| CYP1A2 | rs72547512 | GG | GG | DPYD | rs1801267 | CC | CC |
| CYP1A2 | rs72547513 | CC | CC | DPYD | rs1801268 | CC | CC |
| CYP1A2 | rs72547515 | GG | GG | DPYD | rs3918290 | CC | CC |
| CYP1A2 | rs72547516 | AA | AA | DPYD | rs55886062 | AA | AA |
| CYP1A2 | rs72547517 | GG | GG | DPYD | rs56038477 | CC | CC |
| CYP1A2 | rs138652540 | TT | CC | DPYD | rs72549303 | II | II |
| CYP1A2 | rs143193369 | CC | CC | DPYD | rs72549306 | CC | CC |
| CYP1A2 | rs144148965 | GG | GG | DPYD | rs72549309 | II | II |
| CYP1A2 | rs149928755 | CC | CC | DPYD | rs75017182 | GG | GG |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|--------------|----------------------|--------------------|------|--------------|----------------------|--------------------|
| CYP1B1 | rs10012 | GG | GG | DPYD | rs78060119 | CC | CC |
| CYP1B1 | rs1056827 | CC | CC | FMO1 | rs12720462 | CC | CC |
| CYP1B1 | rs1056836 | GG | CC | FMO1 | rs60639054 | CC | CC |
| CYP1B1 | rs1800440 | TT | TT | G6PD | rs1603411177 | GG | GG |
| CYP1B1 | rs4986888 | GG | GG | G6PD | rs137852348 | GG | GG |
| CYP1B1 | rs28936701 | GG | GG | G6PD | rs137852344 | GG | GG |
| CYP1B1 | rs55771538 | CC | CC | G6PD | rs72554664 | CC | CC |
| CYP1B1 | rs55989760 | CC | CC | G6PD | rs782608284 | CC | CC |
| CYP1B1 | rs56010818 | CC | CC | G6PD | rs72554665 | CC | CC |
| CYP1B1 | rs56175199 | GG | GG | G6PD | rs137852324 | CC | CC |
| CYP1B1 | rs56305281 | GG | GG | G6PD | rs398123546 | CC | GG |
| CYP1B1 | rs72549387 | CC | CC | G6PD | rs1557229572 | GG | CC |
| CYP2A6 | rs1801272 | AA | AA | G6PD | rs137852317 | AA | CC |
| CYP2A6 | rs5031016 | AA | AA | G6PD | rs1557229599 | CC | GG |
| CYP2A6 | rs5031017 | CC | CA | G6PD | rs137852337 | GG | CC |
| CYP2A6 | rs6413474 | TT | TT | G6PD | rs2230037 | CC | GG |
| CYP2A6 | rs28399433 | AA | AA | G6PD | rs782098548 | GG | CC |
| CYP2A6 | rs28399435 | CC | CC | G6PD | rs137852336 | GG | CC |
| CYP2A6 | rs28399440 | AA | AA | G6PD | rs137852323 | GG | CC |
| CYP2A6 | rs28399445 | II | CC | G6PD | rs137852325 | GG | CC |
| CYP2A6 | rs28399447 | AA | AA | G6PD | rs1557229683 | TT | GG |
| CYP2A6 | rs28399454 | CC | CC | G6PD | rs137852335 | CC | CC |
| CYP2A6 | rs28399468 | CC | CC | G6PD | rs137852316 | CC | CC |
| CYP2A6 | rs60563539 | GG | GG | G6PD | rs137852321 | GG | CC |
| CYP2A6 | rs376817657 | CC | CC | G6PD | rs137852334 | AA | GG |
| CYP2A6 | rs568811809 | II | II | G6PD | rs137852320 | AA | TT |
| CYP2A6 | rs763469584 | AA | AA | G6PD | rs137852322 | AA | AA |
| CYP2A6 | rs1967144166 | GG | GG | G6PD | rs371489738 | AA | CC |
| CYP2B6 | rs2279343 | AA | AG | G6PD | rs1057518975 | AA | TT |
| CYP2B6 | rs3211371 | CC | CT | G6PD | rs137852329 | TT | GG |
| CYP2B6 | rs3745274 | GG | GT | G6PD | rs137852345 | CC | GG |
| CYP2B6 | rs8192709 | CC | CC | G6PD | rs2148328996 | AA | CC |
| CYP2B6 | rs12721655 | AA | AA | G6PD | rs137852333 | AA | GG |
| CYP2B6 | rs28399499 | TT | TT | G6PD | rs2070355916 | AA | CC |
| CYP2B6 | rs33926104 | TT | CC | G6PD | rs34193178 | AA | CC |
| CYP2B6 | rs34223104 | TT | TT | G6PD | rs398123544 | AA | TT |
| CYP2B6 | rs34698757 | CC | CC | G6PD | rs137852342 | AA | GG |
| CYP2B6 | rs34826503 | CC | CC | G6PD | rs1557229854 | GG | GG |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|-------------|----------------------|--------------------|------|--------------|----------------------|--------------------|
| CYP2B6 | rs35010098 | CC | CC | G6PD | rs5030869 | GG | CC |
| CYP2B6 | rs35303484 | AA | AA | G6PD | rs1379306569 | TT | GG |
| CYP2B6 | rs35773040 | GG | GG | G6PD | rs76723693 | GG | AA |
| CYP2B6 | rs35979566 | TT | TT | G6PD | rs137852347 | GG | AA |
| CYP2B6 | rs36056539 | CC | CC | G6PD | rs137852339 | CC | CC |
| CYP2B6 | rs36060847 | GG | GG | G6PD | rs2070360793 | CC | CC |
| CYP2B6 | rs45482602 | CC | CC | G6PD | rs137852327 | TT | CC |
| CYP2B6 | rs145884402 | GG | GG | G6PD | rs74575103 | GG | CC |
| CYP2B6 | rs186335453 | GG | GG | G6PD | rs192737996 | GG | GG |
| CYP2B6 | rs193922917 | CC | CC | G6PD | rs137852318 | CC | CC |
| CYP2B6 | rs193922918 | GG | GG | G6PD | rs137852346 | TT | CC |
| CYP2B6 | rs281864907 | TT | TT | G6PD | rs2070375134 | GG | GG |
| CYP2B6 | rs373489637 | TT | TT | G6PD | rs782757170 | TT | GG |
| CYP2B6 | rs564083989 | GG | GG | G6PD | rs137852328 | TT | CC |
| CYP2C8 | rs1058930 | GG | GG | G6PD | rs137852319 | TT | AA |
| CYP2C8 | rs3832694 | DD | II | G6PD | rs137852326 | TT | CC |
| CYP2C8 | rs10509681 | TT | TC | G6PD | rs782754619 | GG | TT |
| CYP2C8 | rs11572103 | TT | TT | G6PD | rs781865768 | GG | TT |
| CYP2C8 | rs41286886 | CC | CC | G6PD | rs137852330 | CC | GG |
| CYP2C8 | rs45438799 | GG | GG | G6PD | rs782170731 | GG | GG |
| CYP2C8 | rs72558195 | GG | GG | G6PD | rs5030868 | GG | GG |
| CYP2C8 | rs72558196 | II | II | G6PD | rs267606836 | TT | GG |
| CYP2C8 | rs78637571 | CC | CC | G6PD | rs5030872 | TT | TT |
| CYP2C8 | rs188934928 | CC | CC | G6PD | rs137852343 | TT | AA |
| CYP2C8 | rs769460274 | TT | TT | G6PD | rs137852331 | GG | TT |
| CYP2C9 | rs1057910 | AA | AA | G6PD | rs137852314 | CC | CC |
| CYP2C9 | rs1799853 | CC | CT | G6PD | rs370918918 | GG | CC |
| CYP2C9 | rs1934969 | AA | TT | G6PD | rs782487723 | AA | CC |
| CYP2C9 | rs2185570 | TT | TC | G6PD | rs137852313 | GG | CC |
| CYP2C9 | rs2256871 | AA | AA | G6PD | rs979416826 | AA | GG |
| CYP2C9 | rs4917636 | AA | AG | G6PD | rs782322505 | CC | TT |
| CYP2C9 | rs7900194 | GG | GG | G6PD | rs137852341 | TT | CC |
| CYP2C9 | rs9332119 | GG | GC | G6PD | rs78365220 | GG | AA |
| CYP2C9 | rs9332130 | AA | AA | G6PD | rs1050829 | TT | TT |
| CYP2C9 | rs9332131 | DD | II | G6PD | rs782130334 | GG | CC |
| CYP2C9 | rs9332239 | CC | CC | G6PD | rs5030870 | TT | CC |
| CYP2C9 | rs12414460 | GG | GG | G6PD | rs1557230626 | CC | AA |
| CYP2C9 | rs12772884 | AA | TT | G6PD | rs267606835 | GG | GG |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|---------|--------------|----------------------|--------------------|-------|--------------|----------------------|--------------------|
| CYP2C9 | rs17847037 | CC | CC | G6PD | rs181277621 | GG | CC |
| CYP2C9 | rs28371685 | CC | CC | G6PD | rs782308266 | GG | CC |
| CYP2C9 | rs28371686 | CC | CC | G6PD | rs782090947 | GG | TT |
| CYP2C9 | rs56165452 | TT | TT | G6PD | rs137852349 | CC | AA |
| CYP2C9 | rs57505750 | TT | TT | G6PD | rs1050828 | GG | CC |
| CYP2C9 | rs71486745 | DD | II | G6PD | rs2070404146 | CC | AA |
| CYP2C9 | rs72558187 | TT | TT | G6PD | rs2070404412 | GG | II |
| CYP2C9 | rs72558189 | GG | GG | G6PD | rs137852315 | AA | CC |
| CYP2C9 | rs72558190 | CC | CC | G6PD | rs1000937138 | GG | CC |
| CYP2C9 | rs72558192 | AA | AA | G6PD | rs2070404778 | CC | GG |
| CYP2C9 | rs72558193 | AA | AA | G6PD | rs76645461 | GG | AA |
| CYP2C9 | rs114071557 | AA | AA | G6PD | rs78478128 | GG | GG |
| CYP2C9 | rs142240658 | CC | CC | G6PD | rs1163458456 | TT | TT |
| CYP2C9 | rs182132442 | CC | CC | G6PD | rs137852340 | GG | TT |
| CYP2C9 | rs199523631 | CC | CC | G6PD | rs2070705276 | TT | CC |
| CYP2C9 | rs200183364 | GG | GG | G6PD | rs797043472 | GG | CC |
| CYP2C9 | rs200965026 | CC | CC | G6PD | rs1273138455 | GG | GG |
| CYP2C9 | rs202201137 | AA | AA | GSTM1 | CNV | II | ID |
| CYP2C9 | rs367826293 | GG | GG | GSTM3 | rs7483 | CC | CC |
| CYP2C9 | rs371055887 | GG | GG | GSTP1 | rs1695 | AA | AA |
| CYP2C9 | rs564813580 | AA | AA | GSTP1 | rs1138272 | CC | CC |
| CYP2C9 | rs749060448 | GG | GG | GSTT1 | CNV | II | II |
| CYP2C9 | rs754487195 | GG | GG | MTHFR | rs1801133 | GG | GA |
| CYP2C9 | rs762239445 | GG | GG | MTHFR | rs373398993 | AA | AA |
| CYP2C9 | rs764211126 | AA | AA | NAT1 | rs15561 | AA | AC |
| CYP2C9 | rs767284820 | TT | TT | NAT1 | rs4986782 | GG | GG |
| CYP2C9 | rs767576260 | CC | CC | NAT1 | rs4986783 | TT | TG |
| CYP2C9 | rs769942899 | GG | GG | NAT1 | rs4986988 | CC | CT |
| CYP2C9 | rs774550549 | CC | CC | NAT1 | rs4986989 | AA | AT |
| CYP2C9 | rs781583846 | GG | GG | NAT1 | rs4986990 | GG | GA |
| CYP2C9 | rs868182778 | GG | GG | NAT1 | rs4986991 | TT | TT |
| CYP2C9 | rs988617574 | CC | CC | NAT1 | rs4986992 | TT | TT |
| CYP2C9 | rs1237225311 | CC | CC | NAT1 | rs4987076 | GG | GA |
| CYP2C9 | rs1250577724 | CC | CC | NAT1 | rs5030839 | CC | CC |
| CYP2C9 | rs1274535931 | CC | CC | NAT1 | rs55793712 | AA | AA |
| CYP2C9 | rs2031308986 | AA | AA | NAT1 | rs56172717 | AA | AA |
| CYP2C19 | rs3758581 | AA | GG | NAT1 | rs56318881 | CC | CC |
| CYP2C19 | rs4244285 | GG | GG | NAT1 | rs56379106 | CC | CC |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|---------|--------------|----------------------|--------------------|--------|-------------|----------------------|--------------------|
| CYP2C19 | rs4986893 | GG | GG | NAT1 | rs72554609 | AA | AA |
| CYP2C19 | rs6413438 | CC | CC | NAT1 | rs72554610 | GG | GG |
| CYP2C19 | rs11188072 | CC | CT | NAT1 | rs72554611 | AA | AA |
| CYP2C19 | rs12248560 | CC | CT | NAT1 | rs146727732 | TT | TT |
| CYP2C19 | rs12769205 | AA | AA | NAT2 | rs1208 | GG | AA |
| CYP2C19 | rs17879685 | CC | CC | NAT2 | rs1041983 | CC | TT |
| CYP2C19 | rs17884712 | GG | GG | NAT2 | rs1799929 | CC | CC |
| CYP2C19 | rs17885179 | AA | AA | NAT2 | rs1799930 | GG | AA |
| CYP2C19 | rs28399504 | AA | AA | NAT2 | rs1799931 | GG | GG |
| CYP2C19 | rs41291556 | TT | TT | NAT2 | rs1801279 | GG | GG |
| CYP2C19 | rs55640102 | AA | AA | NAT2 | rs1805158 | CC | CC |
| CYP2C19 | rs55752064 | TT | TT | NAT2 | rs4271002 | GG | GG |
| CYP2C19 | rs56337013 | CC | CC | NAT2 | rs4986996 | GG | GG |
| CYP2C19 | rs58973490 | GG | GG | NAT2 | rs12720065 | CC | CC |
| CYP2C19 | rs72552267 | GG | GG | NAT2 | rs45477599 | TT | TT |
| CYP2C19 | rs72558186 | TT | TT | NAT2 | rs45518335 | CC | CC |
| CYP2C19 | rs113934938 | GG | AA | NAT2 | rs45618543 | GG | GG |
| CYP2C19 | rs118203756 | GG | GG | NAT2 | rs55700793 | AA | AA |
| CYP2C19 | rs118203759 | CC | CC | NAT2 | rs56011192 | CC | CC |
| CYP2C19 | rs138142612 | GG | GG | NAT2 | rs56054745 | AA | AA |
| CYP2C19 | rs140278421 | GG | GG | NAT2 | rs56387565 | TT | TT |
| CYP2C19 | rs145328984 | CC | CC | NAT2 | rs56393504 | GG | GG |
| CYP2C19 | rs192154563 | CC | CC | NAT2 | rs72466456 | TT | TT |
| CYP2C19 | rs375781227 | GG | GG | NAT2 | rs72466459 | CC | CC |
| CYP2C19 | rs1288601658 | AA | AA | NAT2 | rs72466460 | CC | CC |
| CYP2C19 | rs1564656981 | AA | AA | NAT2 | rs72466461 | AA | AA |
| CYP2C19 | rs1564657013 | AA | AA | NAT2 | rs72554615 | TT | TT |
| CYP2C19 | rs1564660997 | CC | CC | NAT2 | rs72554616 | AA | AA |
| CYP2D6 | CNV | II | II | NAT2 | rs72554617 | GG | GG |
| CYP2D6 | rs1058172 | CC | - | NAT2 | rs79050330 | CC | CC |
| CYP2D6 | rs1081003 | GG | GG | NAT2 | rs138707146 | CC | CC |
| CYP2D6 | rs1135822 | AA | - | NAT2 | rs139351995 | AA | AA |
| CYP2D6 | rs1135823 | CC | AA | NUDT15 | rs186364861 | GG | GG |
| CYP2D6 | rs1135824 | TT | TT | NUDT15 | rs116855232 | CC | CC |
| CYP2D6 | rs1135833 | GG | GG | NUDT15 | rs147390019 | GG | GG |
| CYP2D6 | rs1135835 | TT | TT | OGG1 | rs1052133 | CC | CG |
| CYP2D6 | rs1135837 | CC | CC | POR | rs1057868 | CC | CC |
| CYP2D6 | rs1135838 | AA | AA | POR | rs17853284 | CC | CC |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|-------------|----------------------|--------------------|---------|--------------|----------------------|--------------------|
| CYP2D6 | rs4078249 | CC | CC | POR | rs28931607 | GG | GG |
| CYP2D6 | rs5030655 | II | II | POR | rs28931608 | GG | GG |
| CYP2D6 | rs5030862 | CC | CC | POR | rs56256515 | TT | TT |
| CYP2D6 | rs5030865 | CC | CC | POR | rs56355228 | GG | GG |
| CYP2D6 | rs5030867 | TT | TT | POR | rs72552772 | GG | GG |
| CYP2D6 | rs28371703 | GG | TT | POR | rs121912974 | GG | GG |
| CYP2D6 | rs28371704 | TT | CC | POR | rs121912975 | AA | AA |
| CYP2D6 | rs28371705 | GG | GG | POR | rs145782750 | GG | GG |
| CYP2D6 | rs28371706 | GG | GG | POR | rs199634961 | CC | CC |
| CYP2D6 | rs28371710 | CC | CC | POR | rs201513102 | GG | GG |
| CYP2D6 | rs28371717 | CC | CC | POR | rs567904247 | GG | GG |
| CYP2D6 | rs28371725 | CC | CC | POR | rs781946801 | CC | CC |
| CYP2D6 | rs28371730 | CC | CC | POR | rs782128221 | GG | GG |
| CYP2D6 | rs28371733 | CC | CC | POR | rs1304915832 | AA | AA |
| CYP2D6 | rs28371735 | GG | GG | POR | rs1312625886 | TT | TT |
| CYP2D6 | rs35028622 | AA | CC | SLC15A2 | rs1143672 | GG | GA |
| CYP2D6 | rs35742686 | II | II | SLC15A2 | rs2257212 | CC | CT |
| CYP2D6 | rs59421388 | CC | CC | SLC15A2 | rs1143671 | CC | CT |
| CYP2D6 | rs72549346 | DD | DD | SLCO1B1 | rs2306283 | AA | AG |
| CYP2D6 | rs72549347 | GG | GG | SLCO1B1 | rs4149056 | TT | TT |
| CYP2D6 | rs72549348 | TT | TT | SLCO1B1 | rs11045819 | CC | CA |
| CYP2D6 | rs72549349 | CC | CC | SLCO1B1 | rs34671512 | AA | AA |
| CYP2D6 | rs72549353 | II | II | SLCO1B1 | rs55737008 | AA | AA |
| CYP2D6 | rs72549354 | DD | DD | SLCO1B1 | rs55901008 | TT | TT |
| CYP2D6 | rs72549358 | CC | CC | SLCO1B1 | rs56061388 | TT | TT |
| CYP2D6 | rs74478221 | CC | CC | SLCO1B1 | rs56101265 | TT | TT |
| CYP2D6 | rs74802369 | TT | TT | SLCO1B1 | rs56199088 | AA | AA |
| CYP2D6 | rs75386357 | CC | CC | SLCO1B1 | rs56387224 | AA | AA |
| CYP2D6 | rs75467367 | GG | GG | SLCO1B1 | rs59502379 | GG | GG |
| CYP2D6 | rs77312092 | CC | CC | SLCO1B1 | rs72559745 | AA | AA |
| CYP2D6 | rs78482768 | GG | GG | SLCO1B1 | rs72559748 | AA | AA |
| CYP2D6 | rs79292917 | CC | CC | SLCO1B1 | rs373327528 | GG | GG |
| CYP2D6 | rs118203758 | CC | CC | SOD1 | rs121912442 | CC | CC |
| CYP2D6 | rs138100349 | GG | GG | SOD1 | rs121912443 | AA | AA |
| CYP2D6 | rs148769737 | GG | GG | MNSOD | rs4880 | AA | AG |
| CYP2D6 | rs199535154 | AA | - | SULT1A1 | rs750155 | CC | TT |
| CYP2D6 | rs201377835 | CC | CC | SULT1A1 | rs1042028 | CC | CC |
| CYP2D6 | rs267608276 | CC | CC | SULT1A2 | rs1136703 | AA | AA |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|--------------|----------------------|--------------------|---------|-------------|----------------------|--------------------|
| CYP2D6 | rs267608279 | II | II | SULT1A2 | rs10797300 | GG | GG |
| CYP2D6 | rs267608295 | GG | GG | TBXAS1 | rs4528 | CC | CC |
| CYP2D6 | rs267608297 | GG | GG | TBXAS1 | rs4529 | CC | CC |
| CYP2D6 | rs267608308 | CC | CC | TBXAS1 | rs5763 | CC | CC |
| CYP2D6 | rs267608310 | GG | GG | TBXAS1 | rs8192868 | GG | GG |
| CYP2D6 | rs267608311 | GG | GG | TPMT | rs1142345 | TT | TT |
| CYP2D6 | rs267608313 | GG | GG | TPMT | rs1800460 | CC | CC |
| CYP2D6 | rs267608319 | CC | CC | TPMT | rs1800462 | CC | CC |
| CYP2D6 | rs367543000 | GG | GG | UGT1A1 | rs887829 | CC | CT |
| CYP2D6 | rs368858603 | DD | II | UGT1A1 | rs4148323 | GG | GG |
| CYP2D6 | rs374616348 | CC | CC | UGT1A1 | rs35350960 | CC | CC |
| CYP2D6 | rs375135093 | AA | AA | UGT1A3 | rs3821242 | TT | TC |
| CYP2D6 | rs532668079 | CC | CC | UGT1A3 | rs28898619 | GG | GG |
| CYP2D6 | rs535642512 | CC | CC | UGT1A3 | rs45449995 | AA | AA |
| CYP2D6 | rs536109057 | GG | GG | UGT1A3 | rs45625338 | CC | CC |
| CYP2D6 | rs567606867 | CC | CC | UGT1A3 | rs61764030 | CC | CC |
| CYP2D6 | rs568495591 | GG | GG | UGT1A3 | rs758737792 | TT | TT |
| CYP2D6 | rs569229126 | TT | TT | UGT1A4 | rs2011219 | CC | CC |
| CYP2D6 | rs569439709 | CC | CC | UGT1A4 | rs2011404 | TT | TC |
| CYP2D6 | rs569926140 | AA | AA | UGT1A4 | rs3732218 | GG | GG |
| CYP2D6 | rs730882170 | II | II | UGT1A4 | rs3732219 | CC | CC |
| CYP2D6 | rs730882251 | GG | GG | UGT1A4 | rs3732220 | GG | GG |
| CYP2D6 | rs748712690 | TT | TT | UGT1A4 | rs6755571 | CC | CC |
| CYP2D6 | rs751092905 | CC | CC | UGT1A4 | rs12468274 | TT | TT |
| CYP2D6 | rs753126547 | II | GG | UGT1A7 | rs61261057 | GG | GG |
| CYP2D6 | rs763964554 | GG | GG | UGT1A7 | rs61261057 | GG | GG |
| CYP2D6 | rs765776661 | DD | DD | UGT1A8 | rs17863762 | GG | GG |
| CYP2D6 | rs766507177 | TT | TT | UGT1A9 | rs58597806 | GG | GG |
| CYP2D6 | rs770790629 | CC | CC | UGT1A9 | rs72551330 | TT | TT |
| CYP2D6 | rs773790593 | GG | GG | UGT1A9 | rs66915469 | TT | TT |
| CYP2D6 | rs774943042 | CC | CC | UGT1A10 | rs10187694 | GG | GG |
| CYP2D6 | rs949717872 | TT | TT | UGT1A10 | rs28969685 | CC | CC |
| CYP2D6 | rs1450378700 | GG | GG | UGT1A10 | rs56935833 | GG | GG |
| CYP2D6 | rs1555888899 | AA | AA | UGT1A10 | rs58704432 | CC | CC |
| CYP2D6 | rs1555888910 | AA | AA | UGT2B7 | rs7439366 | TT | TT |
| CYP2D6 | rs1930945471 | AA | AA | UGT2B7 | rs12233719 | GG | GG |
| CYP2D6 | rs1931013246 | TT | TT | UGT2B15 | rs1531022 | GG | GA |
| CYP2D6 | rs2146934648 | TT | TT | UGT2B15 | rs1902023 | AA | AC |

| Gene | Marker | Reference's genotype | Patient's genotype | Gene | Marker | Reference's genotype | Patient's genotype |
|--------|-----------|----------------------|--------------------|---------|-------------|----------------------|--------------------|
| CYP2E1 | rs2031920 | CC | CC | UGT2B15 | rs148583958 | GG | GG |
| CYP2E1 | rs2070672 | AA | AA | UGT2B15 | rs368012995 | AA | AA |
| CYP2E1 | rs2070673 | AA | TT | | | | |

ANNEX 2: DRUGS WITH FDA AND EMA RECOMMENDATIONS FOR GENETIC ANALYSIS

| Antidepressant | |
|-------------------|---|
| Gene | Drug |
| CYP2D6 | Amitriptyline, Citalopram, Clomipramine, Desipramine, Fluoxetine, Fluvoxamine, Imipramine, Nefazodone, Nortryptiline, Paroxetine, Protryptiline, Timipramine, Venlafaxine, Vortioxetine |
| CYP2C19 | Citalopram |
| SLCO1B1 | Viloxacin |
| Antiarrhythmic | |
| Gene | Drug |
| CYP2D6 | Propafenone, Quinine |
| CYP3A4 | Droneradone |
| Antianginal | |
| Gene | Drug |
| NAT2 | Isosorbide |
| Antibiotic | |
| Gene | Drug |
| CYP3A4 | Telitromicina |
| NAT2 | Isoniazid, Pyrazinamide, Rifampicin |
| Antiviral | |
| Gene | Drug |
| CYP2B6 | Efavirenz, Tenofovir, Emtricitabine |
| CYP3A4 | Darunavir, Fosamprenavir, Nelfinavir, Indinavir, Ritonavir, Dolutegravir |
| SLCO1B1 | Letermovil |
| Antineoplastic | |
| Gene | Drug |
| UGT1A1 | Irinotecan, Nilotinib, Pazopanib |
| CYP1A2 | Rucaparib |
| CYP2A6 | Letrozole |
| CYP2A6 | Cabazitaxel, Sunitinib, Sirolimus, Ruxolitinib |
| DPYD | Fluorouracil, Capecitabine, Tegafur, Gimeracil, Oteracil |
| MTHFR | Ethinyl Estradiol |
| TPMT | Cisplatin, Azathioprine, Thioguanine, Mercaptopurine |
| Attention Deficit | |
| Gene | Drug |
| CYP2D6 | Atomoxetine |

| Analgesic | |
|-----------------------|---|
| Gene | Drug |
| CYP2D6 | Codeine, Tramadol |
| CYP2C9 | Celecoxib |
| DPYD | Salicylic acid |
| Proton Pump Inhibitor | |
| Gene | Drug |
| CYP2C19 | Dexlansoprazole, Esomeprazole, Lansoprazole, Omeprazole, Rabeprazole , Pantoprazole |
| Antifungal | |
| Gene | Drug |
| CYP2C19 | Voriconazole |
| CYP3A4 | Posaconazole |
| Cholesterol Modifier | |
| Gene | Drug |
| SLCO1B1 | Rousvastatine, Simvastatine, Atorvastatine, Pitavastatine, Fenofibrate |
| Anticoagulant | |
| Gene | Drug |
| CYP2C9 | Prasugrel, Warfarin |
| CYP2C19 | Prasugrel, Ticagrelor |
| CYP3A4 | Prasugrel |
| CYP3A5 | Prasugrel |
| NAT2 | Clopidogrel |
| Anxiolytic | |
| Gene | Drug |
| CYP2C19 | Diazepam, Doxepine |
| CYP2D6 | Doxepina |
| Antimalarial | |
| Gene | Drug |
| CYP2D6 | Quinine sulphate |
| Contraceptive | |
| Gene | Drug |
| CYP2C19 | Drosperinone |
| CYP2B6 | Ospemifen |
| CYP3A4 | Tamsulosin |
| MTHFR | Norelgestromin |

| Antispasmodic | |
|-----------------------|---|
| Gene | Drug |
| CYP3A4 | Tolteridone |
| Narcolepsy Treatment | |
| Gene | Drug |
| CYP2D6 | Modafinil |
| Gaucher disease | |
| Gene | Drug |
| CYP2D6 | Eliglustat |
| Cardiovascular System | |
| Gene | Drug |
| CYP3A4 | Ivabradine, Losartan |
| SLCO1B1 | Ezetimiba, Amlodipine, Perindopril Arginine |
| Anti-epileptic | |
| Gene | Drug |
| CYP2C19 | Clobazam |
| CYP3A4 | Zonamide |
| Antihypertensive | |
| Gene | Drug |
| CYP2D6 | Cavedilol, Metoprolol, Propranolol |
| Anti-inflammatory | |
| Gene | Drug |
| CYP2C9 | Flurbiprofen |
| Antipsychotic | |
| Gene | Drug |
| CYP2D6 | Aripiprazole, Iloperidone, Clozapine, Perphenazine, Pimozide, Risperidone, Thioridazine |
| Muscle Relaxant | |
| Gene | Drug |
| CYP2C19 | Carisoprodol |
| CYP2D6 | Tolterodine |
| NAT2 | Hydralazine |
| Antitussive | |
| Gene | Drug |
| CYP2D6 | Dextromethorphan |
| Bronchodilator | |

| Gene | Drug |
|------------------------|--|
| CYP2D6 | Arformoterol |
| UGT1A1 | Arformoterol, Indacaterol |
| ALS Treatment | |
| Gene | Drug |
| SOD1 | Tofersen |
| Treatment Korea | |
| Gene | Drug |
| CYP2D6 | Tetrabenazine |
| Dry Mouth Treatment | |
| Gene | Drug |
| CYP2D6 | Cevimeline |
| Amino Acid Derivatives | |
| Gene | Drug |
| CYP2D6 | Cevimeline |
| Respiratory System | |
| Gene | Drug |
| CFTR | Ivacaftor, Tezacaftor, Lumacaftor, Elexacaftor |
| Respiratory System | |
| Gene | Drug |
| SLCO1B1 | Elagolix |

ANNEX 3: FOODS AND FOOD-DERIVED COMPONENTS THAT MODULATE METABOLIC DETOXIFICATION

Cytochrome enzyme inducers

| CYP1A1 | | |
|---|----------------|---|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Green tea | <i>In vivo</i> | 45 mL/d green tea |
| Black tea | <i>In vivo</i> | 54 mL/d black tea |
| Curcumin | <i>In vivo</i> | 1 g/kg/d curcumin, found in turmeric and curry powderder |
| Soy | <i>In vivo</i> | 100 mg/kg soybean extract |
| Garlic | <i>In vivo</i> | 30-200 mg/kg garlic oil |
| Fish oil | <i>In vivo</i> | 20.5 g/kg fish oil |
| Rosemary | <i>In vivo</i> | Diet of 0.5 % rosemary extract |
| Astaxanthin | <i>In vivo</i> | Diet of between 0.001 % and 0.03 % astaxanthin for 15 days |
| CYP1A2 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Cruciferous vegetables | Clinic | 7-14 g/kg cruciferous vegetables, including frozen broccoli and cauliflower, radish sprouts, fresh daikon and shredded raw cabbage, 250 g/d Brussels sprouts and broccoli |
| Green tea | <i>In vivo</i> | Green tea (2,5 % p/v) |
| Black tea | <i>In vivo</i> | Do not exceed 3 cups per day |
| Chicory root | <i>In vivo</i> | 10 % dried chicory root diet |
| Astaxanthin | <i>In vivo</i> | Diet of between 0.001 % and 0.03 % astaxanthin for 15 days |
| CYP1B1 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Curcumin | <i>In vivo</i> | 1 g/kg/d curcumin, found in turmeric and curry powderder |
| Cruciferous vegetables | <i>In vivo</i> | 25-250 mg/kg indole-3-carbinol, occurring in cruciferous vegetables |
| CYP2A | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Chicory root | <i>In vivo</i> | 10 % dried chicory root diet |
| CYP2A6 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Quercetin | Clinic | 500 mg/d quercetin, found in apples, apricots, blueberries, alfalfa sprouts, broccoli, black tea, green beans, kale, and chili powder |
| Broccoli | Clinic | 10 % dried chicory root diet |

| CYP2E1 | | |
|---|----------------|--|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Fish oil | <i>In vivo</i> | 20.5 g/kg fish oil |
| Chicory root | <i>In vivo</i> | 10% dried chicory root diet |
| CYP3A4 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Curcumin | <i>In vivo</i> | 1 g/kg/d curcumin, found in turmeric and curry powderder |
| CYP4B1 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Caffeic acid | <i>In vivo</i> | 179 mg/kg caffeic acid, present in coffee |

Enzyme inducers involved in phase II of liver detoxification

| NRF2 | | |
|---|----------------|--|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Curcumin | <i>In vivo</i> | 200 mg/kg/d de cucumina, presenta en la cúrcuma o en el curry |
| Cruciferous vegetables | <i>In vivo</i> | 0.5 mg/kg/d sulforaphane (compound found in cruciferous vegetables) |
| Garlic | <i>In vivo</i> | 250 mg/kg/d raw garlic |
| Catechins | <i>In vivo</i> | 15 mg/kg epicatechins, found in blackberries, cherries, red wine or dark chocolate |
| Resveratrol | <i>In vivo</i> | 10 mg/kg/d, present in grapes, wine, nuts, soya or tea itadori |
| Ginger | <i>In vivo</i> | Between 10 and 100 mg/kg ginger extract, kudze root |
| Purple sweet potato | <i>In vivo</i> | 100-200 mg/kg anthocyanin, present in purple sweet potato |
| Isoflavones | <i>In vivo</i> | 80 mg/kg/d of isoflavones, found in soya and red clover |
| Coffee | <i>In vivo</i> | 1% of your weight in mL of coffee per day |
| Rosemary | <i>In vivo</i> | 50-100 mg/kg carnosic acid, present in rosemary |
| Blueberries | <i>In vivo</i> | Between 0.6 and 10 g per day |
| Pomegranate | <i>In vivo</i> | 1 to 10 mg/kg of pomegranate extract |
| Naringenin | <i>In vivo</i> | 50 mg/kg/d of naringenin, found in citrus fruits |
| Ellagic acid | <i>In vivo</i> | Dieta de 0.4 % de ácido elágico, presente en bayas, granada, uvas, nueces y grosellas negras |
| Astaxanthin | <i>In vivo</i> | 15 mg/kg astaxanthin, found in algae, yeast, salmon, trout, krill, shrimps and crayfish |
| γ -tocoferol | <i>In vivo</i> | 80 mg/kg of γ -tocoferol |

Cytochrome enzyme inhibitors (phase I of hepatic detoxification)

| CYP1A1 | | |
|---|---------------------|---|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Black raspberry | <i>In vivo</i> | 2.5% black raspberry diet |
| Blueberries | <i>In vivo</i> | 2.5% blueberry diet |
| Ellagic acid | <i>In vivo</i> | 30 mg/kg/d ellagic acid, found in berries, pomegranates, grapes, walnuts and blackcurrants |
| Soy | <i>In vivo</i> | 1 mg/kg/d black soybean seed coat extract |
| Black tea | <i>In vivo</i> | 20 mg/kg thiaflavins, found in black tea |
| Turmeric | <i>In vivo</i> | Turmeric diet 1%. |
| CYP1A2 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Apiaceous Vegetables | Clinic | 4 g/kg of celery vegetables, including frozen carrots and fresh celery, dill, parsley, parsnips and parsnips |
| Quercetin | Clinic | 500 mg/d quercetin, found in apples, apricot, blueberries, yellow onion, kale, alfalfa sprouts, green beans, broccoli, black tea and chilli powder. |
| Daidzein Soy | Clinic | 200 mg twice daily dose of daidzein |
| Grapefruit | Clinic | 300 ml grapefruit juice |
| Kale | <i>In vivo</i> | 2 g/kg/d kale or as freeze-dried kale drink |
| Garlic | <i>In vivo</i> | 30 to 200 mg/kg garlic oil, 3 times a week |
| Camomile | <i>In vivo</i> | Camomile tea at 2% |
| Peppermint | <i>In vivo</i> | Peppermint tea at 2 Dandelion |
| <i>In vivo</i> | Dandelion tea at 2% | |
| Curcumin | <i>In vivo</i> | Curcumin diet at 1% |
| CYP2C6 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Ácido elágico | <i>In vivo</i> | Dieta de 1% de ácido elágico presente en bayas, granadas, uvas, nueces y grosellas negras |
| CYP2C9 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Resveratrol | Clinic | 1 g/d of resveratrol (high dose used). Found in wine, grapes, peanuts, soybeans and itadori tea |
| Myricetin | <i>In vivo</i> | 2-8 mg/kg myricetin, found in onions, berries, grapes and red wine |

| CYP2C19 | | |
|---|----------------|---|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Kale | <i>In vivo</i> | 2 g/kg/d kale or as freeze-dried kale drink |
| CYP2D6 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Resveratrol | Clinic | 1 g/d of resveratrol (high dose used). Found in wine, grapes, peanuts, soybeans and itadori tea |
| Garden cress | Clinic | 7.5 g/d in the form of seed powder, divided into two doses |
| Kale | <i>In vivo</i> | 2 g/kg/d kale or as freeze-dried kale drink |
| CYP2E1 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Garlic | <i>In vivo</i> | Diet between 2-5 % garlic powder |
| N-acetyl cysteine | <i>In vivo</i> | 25 mg/kg N-acetyl cysteine, found in vegetables of the genus <i>Allium</i> , onions, garlic, leeks and shallots |
| Ellagic acid | <i>In vivo</i> | 10-30 mg/kg ellagic acid, found in berries, pomegranate, grapes, walnuts and blackcurrants |
| Green tea | <i>In vivo</i> | 45 mL/d green tea |
| Black tea | <i>In vivo</i> | 54 mL/d black tea |
| Dandelion | <i>In vivo</i> | 0.5 to 2 g/kg aqueous dandelion leaf extract |
| Crisina | <i>In vivo</i> | 20-40 mg/kg/d chrysin, present in honey |
| Medium Chain Triglycerides (MCTs) | <i>In vivo</i> | 32 % of calories from MCTs, found in coconut and coconut oil. |
| CYP3A4 | | |
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Grapefruit | Clinic | 300 ml grapefruit juice |
| Resveratrol | Clinic | 1 g/d of resveratrol (note high dose used). Found in wine, grapes, peanuts, soya and itadori tea. |
| Garden cress | Clinic | 7.5 g/d in the form of seed powder, divided into two doses |
| Soy bean | <i>In vivo</i> | Soybean extract 100 mg/kg |
| Kale | <i>In vivo</i> | 2 g/kg/d kale or as freeze-dried kale drink |
| Myricetin | <i>In vivo</i> | 2-8 mg/kg myricetin, found in onions, berries, grapes and red wine |

Inhibitors of enzymes involved in phase II of liver detoxification

| NRF2 | | |
|---|----------------|--|
| Bioactive foods, beverages or compounds | Type of study | Dosages |
| Luteolin | <i>In vivo</i> | 40 mg/kg luteolin three times a week, found in oranges, artichokes, carrots, broccoli and celery |
| Quercetin | <i>In vivo</i> | 50 mg/kg/d quercetin, found in onions, berries, apples, red wine and green tea |

Foods, beverages and bioactive compounds with demonstrated or potential clinical impact on detoxification systems

| Food or beverage | Compounds with bioactive nutrients |
|------------------------|------------------------------------|
| Verdura allium | Astaxanthin |
| Apiculent vegetables | Caffeic acid |
| Black raspberry | Catechins (including EGCG) |
| Black tea | Crisine |
| Blueberries | Curcumin |
| Camomille | Daidzein |
| Chicory root | Ellagic acid |
| Citrus fruit | Feluric acid |
| Coffee | Fish oil |
| Cruciferous vegetables | Genistein |
| Dandelion tea | Luteolin |
| Garlic | Lycopene |
| Ghee | MCTs |
| Ginger | Myricetin |
| Grapefruit | N-acil cisteine |
| Green tea | Naringenine |
| Honeybush tea | Quercetin |
| Peppermint | Resveratrol |
| Pomegranate | vitamin A |
| Sweet potatoes | |
| Tea Rooibos | |
| Rosemary | |
| Soy bean | |

TECHNOLOGY

DNA Microarray technology consists of a solid surface with microscopic reactions (microreactions) or DNA chip, on which molecular probes are attached to detect the presence of target DNA molecules. Probe-target hybridization is usually detected and quantified by measuring the intensity of a given fluorescence provided by the molecular probe in samples. This type of technology allows the detection of thousands of specific DNA fragments present in a DNA sample. On the other hand, the specificity in terms of DNA sequence recognition is very high since single nucleotide exchange (single-base resolution) can be detected using short oligonucleotide probes (20-25 nucleotides). As a result, DNA Microarray technology has also evolved to be applied as a DNA sequencing technique to genotype several hundred thousand single nucleotide variants (SNVs) in target genes located throughout the genome (Whole Genome DNA Microarray).

Bead Chip Infinium Global Screening Array Orion (GSA Orion) is a line of DNA chips developed by Illumina for its DNA Microarray iScan platform, widely used in population genetic studies and precision medicine, providing optimized content with 100 % reliable and reproducible high-quality genotyping results. The construction of the GSA Chip was carried out in collaboration with a consortium of experts, and for the selection of SNVs, information from prestigious scientific databases such as gnomAD, NHGRI-EBI-GWAS Catalog, ClinVar, MHC-HLA-KIR and PharmGKB has been used. The GSA allows the analysis of approximately 700,000 SNVs that cover variants of interest (hot spots) throughout the entire genome, impacting a wide range of genetic traits with physiological and pathophysiological implications. In addition, it allows the customization by users to incorporate Ad Hoc 50,000-100,000 variants of interest.

QUALITY

The analysis laboratory has standard and effective procedures to protect against technical and operational problems. However, results can be altered due to problems with sample collection (contamination) and labelling (identification), delay in receiving the sample in the laboratory (integrity), among other problems. This could lead to invalidation of the test results. In such cases, the patient would be asked to repeat the entire procedure to perform the test.

As with all clinical analytical tests, there is a small chance that the laboratory may report inaccurate information. If there is a suspicion of an error in the genotype detected, further verification testing may be requested.

LIMITATIONS

The results of the detox test should be used as one more tool in a wide range of factors to be taken into consideration when making therapeutic decisions.

Metabolic response is affected by other factors such as concomitant treatments with other drugs, diseases, toxic habits, age, gender, etc. Treatment decisions should be made according to the judgement of the responsible physician.

REFERENCES

[1] Hodges RE, Minich DM. *Modulation of Metabolic Detoxification Pathways Using Foods and Food-Derived Components: A Scientific Review with Clinical Application*. Journal of Nutrition and Metabolism Volume 2015, Article ID 760689 <http://dx.doi.org/10.1155/2015/760689>

