



## RESULTS REPORT

### PATIENT IDENTIFICATION

NFGI00011 (TEST1)

Sample Code:

T21-I00011

Request date:

03/01/2021

Analysis #

0001

Entry date:

03/03/2021

REQUESTING DOCTOR:

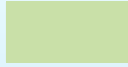
CTS Demo Doctor

Hospital/Clinic:

CTS NFG DEMO

## SUMMARY TABLE

An initial interpretation of the results obtained from the patients genetic profile is displayed in a table below. For each drug examined, the result is indicated according to the following code:



▶ No genetic variants relevant to the treatment have been found. Use as directed.



▶ Need for drug dose monitoring and/or less likelihood of positive response.



▶ Increased likelihood of positive response and/or lower risk of adverse drug reactions.



▶ Increased risk of adverse drug reactions.

Antidepressants		
Agomelatine		
Citalopram		
Desvenlafaxine		
Escitalopram		
Imipramine		
Nortriptyline		
Trazodone		
Vortioxetine		
Amitriptyline		
Clomipramine		
Doxepin		
Fluoxetine		
Mianserin		
Paroxetine		
Trimipramine		
Bupropion		
Desipramine		
Duloxetine		
Fluvoxamine		
Mirtazapine		
Sertraline		
Venlafaxine		

Antipsychotics		
Aripiprazole		
Haloperidol		
Olanzapine		
Pimozide		
Thioridazine		
Brexpiprazole		
Iloperidone		
Paliperidone		
Quetiapine		
Zuclopenthixol		
Clozapine		
Lurasidone		
Perphenazine		
Risperidone		

Stabilizers and anticonvulsants		
Carbamazepine		
Lamotrigine		
Oxcarbazepine		
Topiramate		
Zonisamide		
Clonazepam		
Levetiracetam		
Phenobarbital		
Valproic Acid		
Eslicarbazepine		
Lithium*		
Phenytoin		
Vigabatrin		

Anxiolytics / Hypnotics		
Alprazolam		
Eszopiclone		
Buspirone		
Lorazepam		
Clobazam		
Zolpidem		

Others		
Amphetamines		
Methadone		
Naltrexone		
Atomoxetine		
Methylphenidate		
Lisdexamfetamine		
Naloxone		

\* According to the ATC code, Lithium is considered an antipsychotic (N05AN01). By request of the physicians, the classification of lithium in the table has been modified and it is shown in the mood stabilizers section.

## RESULTS

This section contains the detailed list of drugs with the associated genetic results and interpretation. When different genetic results indicated in different colors occur at the same time for a given drug, the resulting color in the summary table will follow this safety priority rule: risk of adverse drug reactions (red) > dose monitoring (amber) > increased likelihood of positive response and/or lower risk of adverse drug reactions (green). The final evaluation of the analysis results is at the physician's discretion.

### Pharmacogenetics

#### DRUG

#### RESULTS AND INTERPRETATION

##### Agomelatine

**Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP1A2).

**Interpretation:**

The patient carries a variant that has been associated with an increased drug metabolism (CYP1A2). Therefore, he/she may experience a lower exposure to the drug.

##### Alprazolam

**Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

##### Amitriptyline

**Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

##### Amphetamines

**Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Aripiprazole****Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. If needed, increase the drug dosage.

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**Atomoxetine****Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. Be alert to reduced efficacy or select alternative drug.

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**Brexiprazole****Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore, use as directed and titrate dose in response to efficacy and adverse drug events.

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**Bupropion****Analysis result:**

■ Reduced metabolism of the drug (CYP2B6).

**Interpretation:**

The patient carries a variant that has been associated with reduced metabolism of the drug (CYP2B6), therefore a dose adjustment may be necessary.

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**Buspirone****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Carbamazepine****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Citalopram****Analysis result:**

- Higher likelihood of positive response to treatment (*BDNF*).

**Interpretation:**

The analysis indicates the presence of factors associated with a higher likelihood of positive response to treatment (*BDNF*), and therefore, if applicable, use of this drug is recommended in preference to other similar alternatives.

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**Clobazam****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Clomipramine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Clonazepam****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Clozapine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Desipramine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Desvenlafaxine****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Doxepin****Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Duloxetine****Analysis result:**

■ Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. However, there are no clinical data about the effect of this genotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Escitalopram****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Eslicarbazepine****Analysis result:**

■ The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under poly medication (ABCB1).

**Interpretation:**

Consider starting treatment with standard dose (ABCB1) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Eszopiclone****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Fluoxetine****Analysis result:**

- Higher likelihood of positive response to treatment (*BDNF*).
- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates the presence of factors associated with a higher likelihood of positive response to treatment (*BDNF*). Moreover, the analysis indicates the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Fluvoxamine****Analysis result:**

- Higher likelihood of positive response to treatment (*BDNF*).
- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates the presence of factors associated with a higher likelihood of positive response to treatment (*BDNF*). Moreover, the analysis indicates the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there are no clinical data about the effect of this genotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Haloperidol****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).
- Low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*).

**Interpretation:**

The analysis indicates that the patient has a low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*), therefore consider treatment with either a first or second generation antipsychotic as directed on the drug label. In addition, the analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Adjust maintenance dose in response to haloperidol plasma concentration or select an alternative drug.

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**Iloperidone****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Imipramine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Lamotrigine****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Levetiracetam****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Lisdexamfetamine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Lithium****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Lorazepam****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Lurasidone****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.



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**Methadone**

**Analysis result:**

- Reduced metabolism of the drug (*CYP2B6*).

**Interpretation:**

The patient carries a genotype associated with a reduction of methadone metabolism and clearance, increasing its plasma concentrations and therefore the risk of toxicity. Furthermore, carriers of this genotype in treatment with methadone for heroin addiction may require a decreased dose for effective treatment.

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**Methylphenidate**

**Analysis result:**

- Higher likelihood of positive response to treatment (*COMT*).

**Interpretation:**

The analysis indicates there is a higher likelihood of positive response to treatment (*COMT*), and therefore, if applicable, use of this drug is recommended in preference to other similar alternatives.

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**Mianserin**

**Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Mirtazapine**

**Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. This phenotype has been associated with increased clearance of the drug. Use as directed and titrate dose in response to efficacy and adverse drug events.

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**Naloxone**

**Analysis result:**

- Higher likelihood of positive response to treatment (*OPRM1*).

**Interpretation:**

The analysis indicates there is a higher likelihood of positive response to treatment (*OPRM1*), and therefore, if applicable, use of this drug is recommended in preference to other similar alternatives.

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**Naltrexone**

**Analysis result:**

- Higher likelihood of positive response to treatment (*OPRM1*).

**Interpretation:**

The analysis indicates there is a higher likelihood of positive response to treatment (*OPRM1*), and therefore, if applicable, use of this drug is recommended in preference to other similar alternatives.

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**Nortriptyline****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Olanzapine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP1A2*).

**Interpretation:**

The analysis suggests that the patient metabolizes the drug faster than average (*CYP1A2*), and therefore a higher dose than standard is recommended.

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**Oxcarbazepine****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under poly medication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Paliperidone****Analysis result:**

- Low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*).

**Interpretation:**

The analysis indicates that the patient has a low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*), therefore consider treatment with either a first or second generation antipsychotic as directed on the drug label.

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**Paroxetine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).
- Increased risk of drug-related adverse effects (*HTR2A*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Select an alternative drug not predominantly metabolized by this pathway. In addition, the analysis also indicates an increased risk of developing drug-related adverse effects (*HTR2A*). Therefore, select an alternative drug or use reduced doses.

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**Perphenazine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Phenobarbital****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Phenytoin****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).
- Intermediate metabolizer of the drug (*CYP2C9*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2C9* intermediate metabolizer of this drug. Consider using a standard loading dose. Reduce maintenance dose by 25%. Evaluate response and serum concentration after 7-10 days. Be alert to adverse drug events such as ataxia, nystagmus, dysarthria or sedation. On the other hand, the patient may display pharmacoresistance (*ABCB1*), and thus it may be preferable to use another drug.

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**Pimozide****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Quetiapine****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Risperidone****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).
- Low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*).

**Interpretation:**

The analysis indicates that the patient has a low risk of developing extrapyramidal symptoms (*AKT1-DDIT4-FCHSD1-RPTOR*), therefore consider treatment with either a first or second generation antipsychotic as directed on the drug label. In addition, the analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Select an alternative drug or be extra alert to decreased efficacy and titrate dose in response to clinical effect.

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**Sertraline****Analysis result:**

- Higher likelihood of positive response to treatment (*BDNF*).

**Interpretation:**

The analysis indicates the presence of factors associated with a higher likelihood of positive response to treatment (*BDNF*), and therefore, if applicable, use of this drug is recommended in preference to other similar alternatives.

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**Thioridazine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this phenotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Topiramate****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Trazodone****Analysis result:**

- No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Trimipramine****Analysis result:**

- Ultrarapid metabolizer of the drug (*CYP2D6*).

**Interpretation:**

The analysis indicates that the patient is a *CYP2D6* ultrarapid metabolizer of this drug. Consider an alternative drug not metabolized by this pathway. If this drug is warranted, consider increasing the recommended starting dose.<sup>1</sup> Use therapeutic drug monitoring to guide dose adjustments<sup>3</sup>.

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**Valproic Acid****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (*ABCB1*).

**Interpretation:**

Consider starting treatment with standard dose (*ABCB1*) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Venlafaxine****Analysis result:**

- Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. Select an alternative drug or titrate dose to a maximum of 150% of the normal dose in response to efficacy and adverse drug events.

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**Vigabatrin****Analysis result:**

- The patient carries a variant that has been associated with resistance to antiepileptic drugs in adult patients under polymedication (ABCB1).

**Interpretation:**

Consider starting treatment with standard dose (ABCB1) and, in case of pharmacoresistance, evaluate the need for dose increase or change of drug always at the discretion of the physician.

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**Vortioxetine****Analysis result:**

- Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. However, there is no evidence suggesting a clinical effect of this genotype; therefore use as directed and titrate dose in response to efficacy and adverse drug events.

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**Zolpidem****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Zonisamide****Analysis result:**

No variations related to response and/or metabolism that are different from the population standard were found in the analyzed genes.

**Interpretation:**

Use as directed.

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**Zuclopenthixol****Analysis result:**

- Ultrarapid metabolizer of the drug (CYP2D6).

**Interpretation:**

The analysis indicates that the patient is a CYP2D6 ultrarapid metabolizer of this drug. Be alert to low zuclopenthixol plasma levels or select alternative drug.

The following clarifications apply only to tricyclic antidepressants, provided that they are referenced in the text of the recommendation:

(1) Patients may receive a low TCA starting dose, which will be increased over a number of days until the recommended steady-state dose has been reached. The starting dose in these guidelines refers to the recommended steady-state dose.

(3) Dosage recommendations apply to high starting doses, used in the treatment of conditions such as depression. For conditions in which this drug is used in lower doses, like neuropathic pain, there is also a risk of inefficacy for ultrarapid metabolizers; alternative agents should therefore also be considered.

## Folic Acid Conversion

### GENE

### RESULT AND INTERPRETATION

#### **MTHFR**

#### **Analysis result:**

Reduced MTHFR enzyme activity.

#### **Interpretation:**

The patient carries the T allele of the *MTHFR* C677T polymorphism in homozygosis. This genotype has been associated with reduced MTHFR enzyme activity, significantly reduced serum folate levels, and elevated serum homocysteine levels. L-methylfolate may be a preferred option of folate supplementation if clinically indicated.

