#### SUPPLEMENTARY MATERIAL

Supplementary Table 1:

Data from the clinical history that could lead to refer or not the patient to perform an allergy study

## Hypersensitivity reaction analysis: when should we refer to an allergist?

- 1. When there is a history of a previous HSR:
  - For most patients with an HSR to  $\beta$ -lactams, NSAIDs, or anaesthetics (including local anaesthetics).
  - For other drugs, depending on the type of drug, the severity of the reaction and the availability of alternative drugs with a similar efficacy.
- 2. When there is a history of a previous severe HSR and the study is being performed for other structurally related drugs.

# Hypersensitivity reaction analysis: when is an evaluation not required?

- 1. Cases without a drug allergy causality:
  - Incompatible symptomatology.
  - Inconsistent timeline.
  - Subsequently tolerated drug.
  - Similar reaction without taking the drug.
  - Alternative diagnosis (e.g., herpesvirus rash, chronic urticaria, etc.).
- 2. For provocation testing, whenever the original reaction was too severe (e.g., an uncontrollable reaction or severe life-threatening reactions).

### Supplementary Table II

# Relevant points in clinical histories in relation to drug allergies.

- 1. Reaction date.
- 2. Reason for the drug prescription.
- 3. Drugs received.
- 4. Administration route.
- 5. Number of doses administered until the onset of symptoms.
- 6. Time elapsed between the drug administration and onset of the symptoms.
- 7. Symptoms and signs presented during the allergic reaction.
- 8. Treatment given during the allergic reaction.
- 9. Same symptoms without the drug administration.
- 10. Previous drug exposures.
- 11. Subsequently tolerated drugs.
- 12. Other data.

### Supplementary Table III

List of common side effects that can lead to a false diagnosis of drug allergy.

- Dizziness, nausea, or vomiting with tramadol and other opiates.
- Mucosal itching (genital, oral, or anal) due to broad-spectrum antibiotic candidiasis.
- Extrapyramidalism with sulpiride or metoclopramide.
- Diarrhoea after antibiotic intake.
- Tremors after administration of an iodinated contrast.
- Malar erythema after administration of high-potency prolonged-release corticosteroids.
- Hydrocortisone administered in the Emergency Room causing paresthesia.
- Autonomic symptoms after administration of local anaesthetics.

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