

CYTOTOXIC AGENTS : SKIN TESTS FOR THE DIAGNOSIS OF

DRUG HYPERSENSITIVITY

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BACKGROUND

In our hospital, a dermatologist required us to produce skin tests (and more precisely patch tests) with cytotoxic and anti-HER2 antibody agents (docetaxel, pertuzumab, trastuzumab) in order to evaluate the hypersensitivity of a patient who developed a photosensitive dermatosis after a second cycle of chemotherapy.

Key points of drug hypersensitivity reactions :

- ✓ unpredictible
- ✓ 15% of adverse drug reactions (ADR)
- ✓ drug allergy or non-immunological drug hypersensitivity reaction
- ✓ immediate reactions (manifestations within 1-6 hours following drug intake) or delayed reactions (manifestations several hours to days later)

Because of lack of data about skin tests with cytotoxic agents, the aim of this project was to realize a feasibility study for the production of skin tests preparations with these three molecules.

PURPOSE

METHODS



tests

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Physical



RESULTS

- review Literature tests
- pH drug must be superior to 6 and inferior to 9
 - Excipient used for patch preparation petroleum jelly in most of publication (or water)
 - Drug concentration : 10% of pure active drug or 30% if commercialized form is used
 - Positive control : histamine or codeine phosphate
 - Negative control : 0,9% saline

- Preparation realized in other clinical services in most of the time
- the ear e prick tests
- ition Ω - 1 hospital : prick, IDR, and 60 **Q** patch tests but mainly for esti platinum salt
- Docetaxel is too acid (pH drug pur = 3 ; diluted with 0,9 percent sodium chloride drug pH = 4)
- Docetaxel immiscible with water
- pH of trastuzumab and pertuzumab = 7
- pH of the diluant = 7
- Trastuzumab and pertuzumab miscible with

- Skin tests performed :
- \checkmark Docetaxel \rightarrow prick test
 - ¼ dilution
 - Sterile preparation in syringe
- ✓ Trastuzumab and pertuzumab : patch tests
 - 30% petroleum jelly

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- Vertical laminar airflow hood
- ✓ Conservation : 1 day









CONCLUSION

The literature deals mainly with platinum agents, and more often with skin prick tests and intradermal tests. We were confronted with the difficulty to have poor data to face the request of the dermatologist. Moreover, according to the literature, cytotoxic patch tests commonly reveal false negative results. This activity in routin needs the development of a local thesaurus and an economic study because of its expansive cost. http://www.eahp.eu/2

Guidelines for performing skin tests with drugs in the investigation of cutaneous adverse drug reactions, Contact Dermatitis, 2001 Skin test concentrations for systemically administered drugs – an ENDA/EAACI Drug Allergy Interest Group position paper, Brockow et al. Allergy 2013 Diagnostic and predictive value of skin testing in platinum salt hypersensitivity, Leguy-Seguin et al. J Allergy Clin Immunol 2007