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Safe handling of cytotoxics

Sewell, GJ

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Safe handling of cytotoxics

Despite the adoption of standard protective measures, healthcare workers are still exposed to cytotoxic drugs. The risks to the operator and the environment may be reduced by considering effective decontamination and intervention methods and BSCs, and ongoing use of ancillary equipment such as lighting and pump motors.10

Production of aerosols has also been demonstrated as a source of contamination when applying techniques used to reconstitute and dilute cytotoxic drugs.11,12 Oral ingestion or accidental dermal exposure from needle stick injuries, spillages or breakages could also occur.13 These events happen relatively less often, but can contribute to airborne and long-term workplace contamination.

Precontaminated sources taken into the environment before any manipulations are carried out will contaminate gloves and any surfaces subsequently touched. It has been demonstrated that the external surfaces of drug vials may be contaminated with the drug contained, which could also contaminate packaging. Levels found varied according to origin, indicating that precautionary measures taken are not standardised.14-17 Contamination on vials' external surfaces may occur due to splashing, foaming or dusting during the filling process. This may be reduced by vial washing and subsequent application of protective sleeves.15

In the UK and France, current practice in hospital pharmacies is to use negative-pressure isolators.18 Isolators offer containment, but there is no evidence for their superiority over BSCs. Isolators have their limitations and there is the potential for contamination to pass through hatches from the main chamber to the environment. Cytotoxic contamination has been reported on interior surfaces and on surfaces of the finished product in isolators and BSCs.19

If vapourisation were a common occurrence, the HEPA filters of both isolators and BSCs would not be effective in retaining molecules of cytotoxic vapour smaller than the filter's pore size. As a result, any cytotoxic aerosols generated would pass through the filter and be released into the environment. Therefore, whichever source is used, it is paramount to operator protection that the air be exhausted externally, away from the working environment.15

**Guidelines**

Recommendations on good practice have been produced by the Occupational Health and Safety

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**Sarah Roberts**
BSc
PhD Student
Department of Pharmacy and Pharmacology
University of Bath
Bath
UK

**Graham Sewell**
PhD
Professor
Department of Pharmacy
Kingston University
Kingston-upon-Thames
UK

E: s.roberts@kingston.ac.uk

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Figure 1. Contamination from a solution of quinine sulphate produced when withdrawing a syringe after addition to an infusion bag

Administration and the American Society of Health-System Pharmacists in the USA; by the International Society of Oncology Pharmacy Practitioners; by the Health and Safety Executive and the Royal College of Nursing, and the Management and Awareness of the Risks of Cytotoxic Handling group in the UK.20-24

Decontamination methods

EPA

Agents chosen for decontamination in areas where cytotoxic drugs are compounded are mainly intended for biological decontamination. Decontamination protocols should be carefully designed and validated to confirm a biological “kill” and likewise to confirm the removal or chemical degradation of cytotoxic drugs. Different biological agents have varying modes of action and efficacies against different microorganisms; the same applies to cytotoxic drugs, which represent a diverse range of chemical structures.

It is recommended that all surfaces be cleaned according to a protocol, which includes an appropriate deactivation agent to facilitate the removal and/or breakdown of biological and chemical contamination.25 Water-soluble cytotoxic drugs should be removed using wipes impregnated with an aqueous-based agent which binds the target drug, and the wipe should be disposed of afterwards.26 The drug would be more likely to be picked up by applying a detergent formulated at a pH at which the target drug is ionised. One should consider the possibility that surface contamination may take the form of multidrug chemical contamination, since cytotoxic drugs are frequently handled simultaneously; here, a combination of agents may be required. The chemistry of any breakdown products (which may retain cytotoxic activity) should also be considered alongside removal of the target drug. Wiping with industrial methylated spirit is common practice, but may also play a part in the removal of less water-soluble drugs and degradation products.27

Intervention with closed-system containment device

The US National Institute for Occupational Safety and Health recommends the use of a closed-system drug transfer device to prepare cytotoxic drugs, and this has been acknowledged by the relevant European directive and by international guidelines.13,23,26-28

Studies carried out at research centres worldwide, using the PhaSeal® closed-system drug transfer device have all demonstrated a significant reduction of contamination.29-30

Preventing contamination at source is more effective than trying to remove the contamination once it has occurred. Eliminating the primary contamination event will also prevent secondary contamination of areas outside the immediate drug environment.30

However, while closed-system devices may have a role in minimising staff exposure, one cannot recommend that they replace trained pharmacy staff using personal protective equipment and a pharmaceutical isolator.34 Containment by the device

Resources

US Occupational Health and Safety Administration  
W: www.osha.gov

American Society of Health-System Pharmacists  
W: www.ashp.org

International Society of Oncology Pharmacy Practitioners  
W: www.isopp.org

UK Health and Safety Executive  
W: www.hse.gov.uk

UK Royal College of Nursing  
W: www.rcn.org.uk

Management and Awareness of the Risks of Cytotoxic Handling (MARCH)  
W: www.marchguidelines.com
References (continued)


can only be achieved in the area in which the device is used – there would be no influence over pre-contaminated sources brought into the pharmacy.

Conclusions

Despite the adoption of standard protective measures, healthcare workers continue to be exposed to cytotoxic drugs. Targeting decontamination to the chemistry of the drugs used should be considered, as should the implementation of a closed-system device. Manufacturers should also be encouraged to improve their decontamination procedures and to guarantee the supply of vials free of cytotoxic contamination.

References


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