Pharmaceuticals and Medical Devices Safety Information

No. 209 January 2005

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This *Pharmaceuticals and Medical Devices Safety Information (PMDSI)* is issued based on safety information collected by the Ministry of Health, Labour and Welfare. It is intended to facilitate safer use of pharmaceuticals and medical devices by healthcare providers.
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This translation of the original Japanese text is for information purpose only (in the event of inconsistency, the Japanese text shall prevail).

1

Important Safety Information

This section presents contents of revisions, reference materials, and a case summary that served as the basis for these revisions to important adverse reactions included under the PRECAUTIONS section of package inserts of drugs that have been revised in accordance with the Notification after the previous bulletin (Pharmaceuticals and Medical Devices Safety Information No. 208).

1 Phtharal

Brand Name (name of company)	Disopa Solution 0.55% (Johnson & Johnson K.K.)
Therapeutic Category	Insecticides
Indications	Chemical disinfection and sanitization of medical devices

<<**PRECAUTIONS** (underlined parts are additions)>>

[PRECAUTIONS of Indications]

Shock and anaphylactoid symptoms have been reported in patients with a medical history of bladder cancer repeatedly examined with a cystoscope which was disinfected with this drug. This drug should not be used to disinfect medical devices used for transurethral examinations or treatment.

<Reference Information>

Company report

Case Summary

No.	Patient		Adverse reactions	Remarks
	Gender/ Age	Reason for use (complications)	Clinical course and therapeutic measures	
1	Male 70s	Sterilization/ Disinfection of flexible cystoscope (none)	**Anaphylactic shock** On day 1 of use: About 10 minutes after being examined with a flexible cystoscope, generalised itching and welts developed. Although itching was alleviated by the intravenous injection of a d-chlorpheniramine maleate and glycyrrhizin/glycine/cysteine, dysphoria developed about 20 minutes after the intravenous injection. As the patient recovered from the symptoms with drug therapy and oxygen inhalation, he was discharged from the hospital the following day. The physician in charge diagnosed the patient with anaphylactic shock. A flexible cystoscope which was disinfected using this drug was used.	Company report
	Drugs used at time of examination: chlorhexidine gluconate, glycerin, benzalkonium chloride			

No.	Patient		Adverse reactions	Remarks
	Gender/ Age	Reason for use (complications)	Clinical course and therapeutic measures	
2	Male 60s	Sterilization/ Disinfection of flexible cystoscope (hypertension, angina pectoris)	**Shock** On day 1 of use: 1 hour after being examined with the flexible cystoscope, oedema of the prepuce of the penis together with generalised itching occurred. 1 hour and 30 minutes after the examination by the flexible cystoscope, cold sweat and pallor facial developed. The patient went into a state of shock with a blood pressure of 60 mmHg level. The physician in charge diagnosed the patient with acute circulatory failure and allergic shock. The patient recovered after securing an intravenous line, administering oxygen, and implementing drug therapy. A flexible cystoscope which was disinfected using this drug was used.	Company report
	Drug used at time of examination: lidocaine hydrochloride Concomitant medications: amlodipine besilate, candesartan cilexetil, isosorbide mononitrate, isosorbide dinitrate			

2

Revision of PRECAUTIONS

(No. 162)

This section presents details of revisions to the PRECAUTIONS section of package inserts and brand names of drugs that have been revised according to the Notifications after the previous bulletin (Pharmaceuticals and Medical Devices Safety Information No. 208) (excluding those presented in "1. Important Safety Information" of this Bulletin), together with reference materials.

1 <Antihypertensives> Lisinopril

[Brand Name]	Longes Tablets 5 mg, 10 mg, and 20 mg (Shionogi & Co., Ltd.), and others
[Adverse Reactions (clinically significant adverse reactions)]	**Hepatic function disorder, jaundice:** Hepatic function disorder with significant elevations of AST (GOT), ALT (GPT), Al-P and γ-GTP levels etc. and jaundice may occur. Patients should be carefully monitored and if abnormalities are observed, administration should be immediately discontinued and appropriate measures should be taken. It was reported that patients resulted in hepatic failure rarely.
<Reference Information>	Company report

2 <Antitussives> Clofedanol Hydrochloride

[Brand Name]	Coldrin Granules, Coldrin Tablets (Nippon Shinyaku Co., Ltd.)
[Adverse Reactions (clinically significant adverse reactions)]	**Shock, anaphylactoid symptoms:** Shock and anaphylactoid symptoms may occur. Patients should be carefully monitored and if symptoms such as urticaria, cold sweat, dyspnoea, laryngeal oedema, and blood pressure decreased etc. are observed, administration should be immediately discontinued and appropriate measures should be taken.
<Reference Information>	Company report

3 <Antidotes> Calcium Leucovorin (calcium folinate) (5 mg tablets, injectable dosage form)

[Brand Name]	Leucovorin Tablets 5 mg, Leucovorin for Intramuscular Use (Wyeth K.K.)
[Contraindications]	Patients who have a history of serious hypersensitivity to ingredients of this drug
<Reference Information>	Company report

4 <Antimetabolites>
Tegafur/Gimeracil/Oteracil Potassium

[Brand Name] TS-1 Capsule 20 and 25 (Taiho Pharmaceutical Co., Ltd.)

[Adverse Reactions (clinically significant adverse reactions)]

Psychoneurotic disorder including leukoencephalopathy etc.:
Leukoencephalopathy (main symptoms of consciousness disturbed, cerebellar ataxia, dementia-like symptoms etc.), consciousness disturbed, disorientation, somnolence, memory decline, extrapyramidal symptoms, language disorder, quadriplegia, gait disturbance, urinary incontinence, and perceptual disturbance etc. may occur. Patients should be carefully monitored and if such symptoms are observed, administration should be discontinued.

<Reference Information> Company report

5 <Antineoplastics-Miscellaneous>
Mitoxantrone Hydrochloride

[Brand Name] Novantron Injection 10 mg and 20 mg (Wyeth K.K.)

[Important Precautions]

Serious adverse reactions such as myelosuppression and myocardial disorder etc. may occur in the administration of this drug. Caution should be exercised for the following points:

1) This drug should be administered at a medical institution sufficiently capable of responding to emergency situations by a doctor sufficiently experienced with cancer chemotherapies, and only for cases judged to be appropriate targets for administration.

2) Patient should be carefully monitored through frequent clinical examinations (blood tests, hepatic/renal function tests, and cardiac function tests etc.) etc. If abnormalities are observed, take appropriate measures such as reduction or withdrawal of administration etc. Moreover, as strong adverse reactions may occur and become prolonged if the drug is administered over the long term, this drug should be carefully administrated.

The patient should not be vaccinated with live vaccines during the administration of this drug as there is risk that vaccinating a patient with suppressed immune function with a live vaccine may increase or sustain infection originating from the vaccine.

[Adverse Reactions (clinically significant adverse reactions)]

Congestive cardiac failure, myocardial disorder, myocardial infarction:
When the total dosage of this drug is 160 mg/m^2 in cases not administered an anthracycline agent beforehand, or when the total dosage of this drug exceeds 100 mg/m^2 in cases administered an anthracycline agent beforehand, serious cardiac disorder such as congestive cardiac failure etc. may occur. In addition, myocardial disorder may occur in cases which were given an anthracycline agent beforehand, irrespective of the dosage of this drug, cardiac function tests should be frequently conducted. If abnormalities are observed, administration should be discontinued.

Shock, anaphylactoid symptoms: Shock and anaphylactoid symptoms may occur. Patients should be carefully monitored and if symptoms such as rash, dyspnoea, and blood pressure decreased etc. are observed, administration should be immediately discontinued and appropriate measures should be taken.

<Reference Information> Company report

6 <Kampo medicines>

Kamishoyosan

[Brand Name] TSUMURA Kamishoyosan Extract Granules for Ethical Use (Tsumura & Co.), and others

[Adverse Reactions (clinically significant adverse reactions)] **Hepatic function disorder and jaundice:** Hepatic function disorder with significant elevations of AST (GOT), ALT (GPT), Al-P, and γ-GTP levels etc. and jaundice may occur. Patients should be carefully monitored and if abnormalities are observed, administration should be discontinued and appropriate measures should be taken.

<Reference Information> Company report

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7 <Acting mainly on gram-positive and gram-negative bacteria>

Sulbactam Sodium/Cefoperazone Sodium

[Brand Name] Sulperazon for Intravenous Use 0.5 g and 1 g (Pfizer Japan Inc.), and others

[Adverse Reactions (clinically significant adverse reactions)] **Oculomucocutaneous syndrome (Stevens-Johnson syndrome), toxic epidermal necrolysis (Lyell syndrome):** Oculomucocutaneous syndrome (Stevens-Johnson syndrome) or toxic epidermal necrolysis (Lyell syndrome) may occur. Patients should be carefully monitored and if abnormalities are observed, administration should be discontinued and appropriate measures should be taken.

<Reference Information> Company report

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8 <Acting mainly on gram-positive and gram-negative bacteria>

Biapenem

[Brand Name] Omegacin 0.3 g for Intravenous Drip Infusion (Wyeth K.K.), and others

[Adverse Reactions (clinically significant adverse reactions)] **Acute renal failure:** Serious renal disorder such as acute renal failure may occur. Patients should be carefully monitored through periodic tests, etc. and if abnormalities are observed, administration should be discontinued and appropriate measures should be taken.

<Reference Information> Company report

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9 Over the counter drugs

Kamishoyosan

[Brand Name] TSUMURA Kampo Kamishoyosan Extract Granules (Tsumura & Co.), and others

[Consultation] In case of the following, immediately discontinue administration and bring this document to your doctor or pharmacist for consultation.

If the following symptoms are observed after taking this drug

In rare instances, the following serious symptoms may occur. Visit a physician immediately in such a case.

Hepatic function disorder: General fatigue, jaundice (skin and white of the eyes become yellow) etc. may occur.

<Reference Information> Company report