



3484391-X-88-01

RM. RES. INST. USA
or use by user-facilities,
distributors and manufacturers for
MANDATORY reporting

Approved by FDA on 09/25/95

Mfr report #	PRIUSA1999006582
UF/Dist report #	
FDA Use Only	

FDA MEDICAL PRODUCTS REPORTING PROGRAM

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A. Patient information

1. Patient identifier ? - ?	2. Age at time of event: 73 yr Date of birth: ??/??/??	3. Sex <input type="checkbox"/> female <input checked="" type="checkbox"/> male	4. Weight UNK lbs or UNK kgs
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B. Adverse event or product problem

1. Adverse event and/or Product problem (e.g., defects/malfunctions)

2. Outcomes attributed to adverse event (check all that apply)

<input checked="" type="checkbox"/> death ??/??/?? (mo/day/yr)	<input type="checkbox"/> disability
<input type="checkbox"/> life-threatening	<input type="checkbox"/> congenital anomaly
<input checked="" type="checkbox"/> hospitalization - initial or prolonged	<input checked="" type="checkbox"/> required intervention to prevent permanent impairment/damage
	<input type="checkbox"/> other:

3. Date of event (mo/day/yr) ??/??/??	4. Date of this report (mo/day/yr) 11/16/99
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5. Describe event or problem

Report published in 1993 Annual Report of the American Association of Poison Control Centers National Data Collection System (case 205). A 73-year-old patient (sex unspecified) died following intentional misuse of acetaminophen with oxycodone. Serum acetaminophen level 41 mcg/mL. Exposure to medication was chronic.

Additional information received 11-Nov-99: A 73-year-old man had a history of end stage liver disease and chronic pain. He had two pain medications available to him, both contained acetaminophen and oxycodone. His history revealed that pain medication had been taken every four hours for approximately one week prior to emergency department (ED) presentation. In the ED, he was central nervous system depressed. Admission laboratory values showed alkaline phosphatase 127 IU/L, AST 65 IU/L, albumin 2.5 g/dL, lactic dehydrogenase 971 IU/L, total bilirubin 3.8 mg/dL, prothrombin time 16 seconds (INR 1.8), and partial thromboplastin time 49 seconds. Toxicologic analysis revealed serum acetaminophen 41 mcg/mL. Oral N-acetylcysteine therapy was (Cont.)

6. Relevant tests/laboratory data, including dates

(Lab data cont.)

(Cont.)

7. Other relevant history, including preexisting medical conditions (e.g., allergies, race, pregnancy, smoking and alcohol use, hepatic/renal dysfunction, etc.)

Additional information received 11-Nov-99: end stage liver disease, chronic pain

C. Suspect medication(s)

1. Name (give labeled strength & mfr/labeler, if known)
#1 TYLOX (capsule) (OXYCODONE/ACETAMINOPHEN)
#2

2. Dose, frequency & route used #1 unk, 4 hour(s), oral #2	3. Therapy dates (if unknown, give duration) from/to (or best estimate) #1 1 week(s) #2
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4. Diagnosis for use (indication) #1 PAIN #2	5. Event abated after use stopped or dose reduced #1 <input type="checkbox"/> yes <input type="checkbox"/> no <input checked="" type="checkbox"/> doesn't apply #2 <input type="checkbox"/> yes <input type="checkbox"/> no <input type="checkbox"/> doesn't apply
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6. Lot # (if known) #1 #2	7. Exp. date (if known) #1 #2
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8. Event reappeared after reintroduction
#1 yes no doesn't apply
#2 yes no doesn't apply

9. NDC # - for product problems only (if known)

10. Concomitant medical products and therapy dates (exclude if not part of event)
No Concomitant Products Reported

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G. All manufacturers

1. Contact office - name/address (& mfrng site for devices) R.W. JOHNSON PHARM. RES. INST. USA DIV. OF ORTHO PHARMACEUTICAL CORP. 920 U.S. Route 202 P.O. Box 300 Raritan NJ 08869 USA (Informing Unit)	2. Phone number 908-704-4504	3. Report source (check all that apply) <input type="checkbox"/> foreign <input type="checkbox"/> study <input checked="" type="checkbox"/> literature <input type="checkbox"/> consumer <input checked="" type="checkbox"/> health professional <input type="checkbox"/> user facility <input type="checkbox"/> company representative <input type="checkbox"/> distributor <input type="checkbox"/> other:
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4. Date received by manufacturer (mo/day/yr) 11/11/99	5. (A)NDA # 88-790
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6. If IND, protocol #	IND # _____ PLA # _____
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7. Type of report (check all that apply) <input type="checkbox"/> 5-day <input checked="" type="checkbox"/> 15-day <input type="checkbox"/> 10-day <input type="checkbox"/> periodic <input type="checkbox"/> Initial <input checked="" type="checkbox"/> follow-up # 1	pre-1938 <input type="checkbox"/> yes OTC product <input type="checkbox"/> yes
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8. Adverse event term(s) 1) DEATH 2) COMA HEPATIC 3) COMA 4) STUPOR 5) DRUG ABUSE	9. Mfr. report number PRIUSA1999006582
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F. Initial reporter

1. Name, address & phone #
Dr. Toby Litovitz
National Capital Poison Center
Georgetown University Hospital
3800 Reservoir Road NW
Washington, DC 20007
USA
NOV 19 1999

2. Health professional? <input checked="" type="checkbox"/> yes <input type="checkbox"/> no	3. Occupation Physician	4. Initial reporter who sent report to FDA <input type="checkbox"/> yes <input type="checkbox"/> no <input checked="" type="checkbox"/> unk
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Submission of a report does not constitute an admission that medical personnel, user facility, distributor, manufacturer or product caused or contributed to the event.



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B. Adverse event or product problem

B.5 Describe event or problem (Cont...)

started. Fifteen hours after presentation, laboratory and toxicologic analysis showed alkaline phosphatase 116 IU/L, AST 73 IU/L, ALT 40 IU/L, lactic dehydrogenase 1,119 IU/L, prothrombin time 16.9 seconds (INR 2.0), partial thromboplastin time 45 seconds, and acetaminophen 12 mcg/mL. The patient's condition continued to deteriorate and by the fifth hospital day he was unresponsive and in hepatic encephalopathy. He was intubated and mechanically ventilated. Laboratory values on the sixth hospital day showed lactic dehydrogenase 3,224 IU/L, prothrombin time 17.5 seconds (INR 2.2), blood urea nitrogen 40 mg/dL, and serum creatinine 0.8 mg/dL. Life support measures were discontinued and he died on the tenth hospital day. Post-mortem examination was not done.

B.6 Relevant tests/laboratory data, including dates (Cont...)

Lab Result:

Sl.No.	Test date	Test name	Test result	Normal value
1	??/??/??	ALANINE AMINOTRANSFERASE	40 IU/L (international unit/liter)	
		at 15 hours		
		ALBUMIN	2.5 g/dL (grams/deciliter)	
		on admission		
		ALKALINE PHOSPHATASE	127 IU/L (international unit/liter)	
		on admission		
		ALKALINE PHOSPHATASE	116 IU/L (international unit/liter)	
		at 15 hours		
		ASPARTATE AMINOTRANSFERASE	65 IU/L (international unit/liter)	
		on admission		
		ASPARTATE AMINOTRANSFERASE	73 IU/L (international unit/liter)	
		at 15 hours		
		BILIRUBIN, TOTAL	3.8 mg/dL (milligram/deciliter)	
		on admission		
		BLOOD UREA NITROGEN	40 mg/dL (milligram/deciliter)	
		day 6		
		CREATININE	0.8 mg/dL (milligram/deciliter)	
		day 6		
		DRUG LEVEL	41 mcg/mL (microgram/milliliter)	
		acetaminophen, on admission		
		DRUG LEVEL	12 mcg/mL (microgram/milliliter)	
		acetaminophen: at 15 hours		
		LACTIC DEHYDROGENASE	971 IU/L (international unit/liter)	
		on admission		
		LACTIC DEHYDROGENASE	1,119 IU/L (international unit/liter)	
		at 15 hours		
		LACTIC DEHYDROGENASE	3,224 IU/L (international unit/liter)	
		day 6		
		PARTIAL THROMBOPLASTIN TIME	49 sec (second)	
		on admission		
		PARTIAL THROMBOPLASTIN TIME	45 sec (second)	
		at 15 hours		
		PROTHROM TIME	16 sec (second)	
		INR 1.8, on admission		
		PROTHROM TIME	16.9 sec (second)	
		INR 2.0, at 15 hours		
		PROTHROM TIME	17.5 sec (second)	
		INR 2.2, day 6		

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PHARMACY REPORTING SYSTEM

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Source of report (Literature):



3404391-X-00-03

Continuation Sheet for FDA-3500A Form

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Date of this report: 11/16/99

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Author	:	Toby Litovitz
Journal title	:	1993 Annual Report of the American Association of Poison Control Centers National Data Collection System
Year	:	94
Edition	:	12(5)
Page number	:	From 546 To 515
Article title	:	American Journal of Emergency Medicine

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DIVERSE EVENT REPORTING SYSTEM

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