Life-threatening errors with flecainide suspension in children

Flecainide is an oral class 1c antiarrhythmic drug that may be used to treat atrial fibrillation or supraventricular tachycardia, particularly when conventional treatment agents fail. Since it is available commercially only as 50 mg, 100 mg, and 150 mg tablets, it must be compounded into a suspension when needed for infants and small children. Unfortunately, errors during preparation and dosing of the suspension have occasionally led to serious overdoses that resulted in cardiac emergencies and required immediate therapeutic intervention. Overdoses can lead to seizures and cardiotoxicity, including ventricular tachycardia and fibrillation due to sodium channel blockade. Treatment includes sodium bicarbonate boluses or sodium chloride boluses and extracorporeal circulatory support.

We first learned of a flecainide suspension-related error from a report submitted to ISMP in 2007. A 4-month-old infant had been receiving 8 mg twice daily as an 8 mg/mL suspension (1 mL per dose). When the dose was later increased to 10 mg, a suspension purported to be 10 mg/mL was compounded. However, the baby’s mother complained that the suspension was too thick to withdraw from the bottle. The pharmacist asked the mother to return the suspension and elected to compound a 7 mg/mL suspension, instructing the parent to give 1.4 mL per dose. Due to a math error, this replacement suspension was actually compounded with 6 g (6,000 mg) of flecainide instead of 600 mg. Therefore, each dose represented 100 mg, not 10 mg. It is unclear whether the previous 10 mg/mL suspension may also have been prepared incorrectly.

Additional errors with compounded flecainide suspension have appeared in the literature. A 2-year-old child received a 5-fold overdose when unlabeled oral syringes of nadolol (CORGARD) and flecainide were used. Instead of withdrawing 5 mL of nadolol suspension (concentration not specified) and 1 mL of flecainide suspension (20 mg/mL), the nurse administered the opposite and the child received 5 mL of flecainide (100 mg). In another case, a pharmacy dispensed a 5 mg/mL suspension for a 4-week-old child who was supposed to receive 3 mg/0.6 mL TID. Pharmacy staff erroneously transcribed the dosing instructions to take “3 mL” instead of 3 mg (0.6 mL), resulting in an overdose that led to wide complex tachycardia. In a third case, an 18-day-old infant received 4 doses of flecainide 8 mg (0.8 mL of a 10 mg/mL suspension) instead of 4 mg (0.8 mL of a 5 mg/mL), also resulting in wide complex tachycardia and cardiac arrest from which the child recovered. A fourth case involved a 9-month-old infant whose parents were told to increase the dose of flecainide to 4 mL, assuming the concentration was 5 mg/mL as in the original prescription. But the parents refilled the prescription at another pharmacy, receiving the drug in a 20 mg/mL concentration. The patient received 80 mg/4 mL, a 4-fold overdose, resulting in wide complex tachycardia and QRS prolongation. Finally, an event published just this past December described an error in which the aunt of a 7-month-old child unreasonably transcribed the dosing instructions to take “3 mL” instead of 3 mg (0.6 mL), also resulting in wide complex tachycardia.3

To prevent these errors, prescribers should include both brand and generic names as well as the purpose of the medications on prescriptions and limit the use of telephone orders. If used, pharmacies should document telephone orders on a pharmacy prescription blank and read back, spelling the drug name(s) and stating the dose and concentration.4

ISMP has received 4 other reports involving this look- and sound-alike name pair dating back to 2002. The fact that both products are available in 50 mg dosage strengths only increases the risk of confusion. To help differentiate these drug names, the US Food and Drug Administration (FDA) applied tall man lettering to the pair as part of the FDA Name Differentiation Project (www.ismp.org/sc?id=520).

Confused drug names: clomiPHENE and clomiPRAMINE. Recently, a consumer reported that his wife received the wrong medication from a community pharmacy. The physician’s office telephoned a prescription to the pharmacy for the fertility drug clomiPHENE 50 mg, take 3 tablets by mouth daily. However, the pharmacy dispensed clomiPRAMINE 50 mg, a tricyclic antidepressant used to treat obsessive-compulsive disorder, with directions to take 3 capsules (150 mg) by mouth daily. After taking the first dose, the patient reported feeling sick and very sleepy. Later, she developed a headache, dizziness, and nausea. It is not known if any computerized alerts were generated for the high dose of clomiPRAMINE—the starting dose for clomiPRAMINE is 25 mg daily with a gradual titration over 2 weeks to 100 mg daily in divided doses.

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Flecainide—continued from page 1

knowingly gave her nephew a 5-fold overdose of flecainide.\(^5\) She pulled the suspension into an oral syringe that had both a teaspoon and mL scale and measured 5 mL (one teaspoon) instead of 1 mL. The child was hospitalized and suffered a cardiac arrest but was successfully resuscitated. Fortunately, all of the children recovered without neurologic sequelae following life-threatening flecainide overdoses.

Unrecognized changes in drug concentration, math errors, labeling errors, and inaccurate dosing instructions led to the confusion in these cases. In one case, the authors indicated that all of their local pharmacies had agreed to compound flecainide in one standard concentration of 20 mg/mL.\(^1\) Along this line, a Michigan state-wide initiative agreed to standardize the flecainide concentration and other compounded liquid medication concentrations for pediatric patients. The initiative, which received a 2014 ISMP Cheers Award, provides the preparation directions, final concentration, stability data, storage information, and information for prescribers and families about the standards.\(^6\) Such efforts to standardize concentrations in both inpatient and outpatient pharmacies can help eliminate medication errors like those described above. The American Society of Health-System Pharmacists (ASHP), in cooperation with ISMP, the US Food and Drug Administration (FDA), and other stakeholders, is trying to broaden this initiative nationwide.

To reduce the risk of errors, prescribers should order flecainide in terms of the mg dose. This allows pharmacists to address the suspension concentration (mg/mL) and volume per dose, which should be expressed in mL (metric). For neonates and infants, a lower concentration may be required. If the drug is prescribed by volume (mL), the concentration MUST be specified, or the prescriber must be contacted for clarification.

Community pharmacies should provide liquid suspensions with a flow restrictor embedded in the neck of the bottle (Figure 1) along with a metric-only oral syringe to measure and administer doses. Be sure to remind the patient or parents to secure the child-resistant cap after each use. Label directions should include the dose in metric. This also gives pharmacists and parents an opportunity to catch an error.

Figure 1. Flow restrictors placed within neck of bottles of liquid medication.

Community pharmacies should provide liquid suspensions with a flow restrictor embedded in the neck of the bottle (Figure 1) along with a metric-only oral syringe to measure and administer doses. Be sure to remind the patient or parents to secure the child-resistant cap after each use. Label directions should include the dose in terms of mL (not teaspoonfuls), such as “Flecainide 0.25 mL by mouth every 8 hours.” The community pharmacy label should also include the concentration next to the drug name, below the instructions for use. To be sure parents give a proper dose, use “teach-back” methods to demonstrate how to measure and administer proper amounts. This also gives pharmacists and parents an opportunity to catch an error.

References

Health Alert! RB (formerly Reckitt Benckiser) has recalled certain lots of its MUCINEX FAST-MAX combination liquid cough and cold products. The affected lots, which correctly label the product on the front of the bottle and list all active ingredients, may not have the correct corresponding drug facts label on the back panel. This mislabeling could cause patients to be unaware of side effects and/or risks associated with the ingestion of certain product ingredients which include acetaminophen, dextromethorphan, guaifenesin, phenylephrine, and/or diphenhydramine. Patients may also be at risk of taking unintentional overdoses of these ingredients which can result in serious harm.

A press release (www.ismp.org/sc?id=521) from the company provides a list of the affected products and corresponding lot numbers. Check your shelves to make sure these affected products are removed. Post information about the recall at the pharmacy counter as well as on shelves where these products are stocked. Advise patients to safely dispose of any unused product by: 1) mixing liquid medicines with an unpalatable substance such as kitty litter or used coffee grounds; 2) placing the mixture in a container; and 3) disposing of the container in the household trash. For more information about the recall, go to: www.ismp.org/sc?id=519.

continued on page 3—SAFETY briefs>
FIRST brand oral vancomycin needs improved labeling

In our October 2014 issue, we wrote about errors that occurred after a long-term care (LTC) pharmacy elected to send vials of injectable vancomycin powder along with diluent for the LTC staff nurses to reconstitute and administer as an oral solution. This was due to the high cost of VANCOCIN brand capsules and occasional problems in obtaining specific vancomycin products. At two facilities, nurses were unfamiliar with the practice of using the injectable form of vancomycin for oral administration and instead administered a number of doses intramuscularly (IM).

Last year, CutisPharma launched unit-of-use compounding kits for vancomycin oral solution under the FIRST trademark (Figure 1). The kits contain a bottle of vancomycin powder along with a bottle of grape flavored diluent. Both 25 mg/mL and 50 mg/mL concentrations, after reconstitution, are available. Prior to use, the contents of each bottle must be properly reconstituted.

Unfortunately, while this product is convenient to use, we have received reports about the product not being clearly labeled. One issue is that the diluent container, which has no vancomycin, and the powder container have “vancomycin 25” or “vancomycin 50” listed at the top of the label. Someone unfamiliar with the product might see the diluent as a bottle of active drug, especially when the containers have been removed from the carton.

Administering the diluent alone has happened in the past with other oral antibiotics that require reconstitution with a special diluent. In 1999, ISMP alerted practitioners to a similar situation with CIPRO Oral Suspension (ciprofloxacin). The active antibiotic powder container and the diluent container were both labeled “Cipro Oral Suspension.” This led to the diluent being identified as a ready-to-use product, and it was dispensed without reconstituting the actual antibiotic. At the time, ISMP called upon the US Food and Drug Administration (FDA) and the manufacturer to improve label clarity, and the diluent container labels were then revised to state, “DILUENT for Cipro Oral Suspension.” Similarly, the new vancomycin product diluent bottle should be prominently labeled, “DILUENT for vancomycin 25” or “DILUENT for vancomycin 50.”

We also received reports in which the powder alone was dispensed without dilution for oral antibiotic products. This can happen with this product because the powder container is marked with an icon that states, “Oral Solution.” Another issue with this product is that the bottle of vancomycin powder, which acts as the dispensing container after reconstitution, is not prominently labeled with the expected final concentration or the total volume after reconstitution. This information may not be available unless the bottle happens to be relabeled with that information once reconstituted.

The powder container should state, “Vancomycin 25 Powder—Must be Diluted.” Until this change is made by the company, pharmacies using this product are advised to reinforce the need to reconstitute with the diluent.
> Vancomycin—continued from page 3
to label containers appropriately so the diluent and/or powder are not dispensed and administered as the drug itself without dilution. We also recommend labeling the reconstituted powder containers with the final concentration per mL and total volume, along with the notation, “Reconstitution Completed.” We have contacted both FDA and CutisPharma to request improvements to the way the product carton and containers are labeled for this kit, as well as their other oral suspension compounded kits (e.g., lansoprazole, omeprazole).

Special Announcements

Updated list of confused drug names
An updated list of confused drug names was recently added to the ISMP website at: www.ismp.org/sc?id=515. You may also purchase a full color wallchart from our online store at: http://onlinestore.ismp.org/shop/item.aspx?itemid=133. As always, our hope is that you will review this list and consider which require special safeguards at your practice site.

ISMP webinars
Join us on May 27 for Expanding Barcode Medication Administration: Making a Difference in the Emergency Department (ED) and Other Outpatient Settings. Despite widespread adoption of barcode medication administration in hospitals, the safety benefits of this technology have not been as quickly adopted in many associated outpatient clinical locations, such as the ED, oncology clinics, and dialysis locations, where a large number of high-alert medications are administered. Our distinguished faculty will provide an inside look at the implementation of bedside barcode scanning in clinical areas once thought to be “off limits” for this type of technology.

Join us on June 24 for our 2015 Update on The Joint Commission Medication-Related Standards. Learn the most troublesome Medication Management Standards and National Patient Safety Goals along with successful approaches taken by healthcare organizations to accomplish the intent of these Standards. A second presenter will provide personal insight into the Standards, based on a recent Joint Commission survey at a large, nonprofit, teaching hospital. The speaker will share opportunities at both the pharmacy and health-system level.

For details on this and other webinars, please visit: www.ismp.org/sc?id=349.

New ISMP safety CE opportunities
ISMP is now offering two free CE opportunities: Improving Medication Safety in the Perioperative Setting and Addressing a Trifecta of Overlooked IV Medication Risks. Each program is approved for 1.5 contact hours. For details, go to: www.ismp.org/sc?id=509.

SAFETY briefs cont’d from page 3

SAFETY briefs

Look-alike opioid bottles. A pharmacist recently alerted us to a potential hazard. The bottles of oxyCODONE 10 mg and oxymorphone 10 mg manufactured by KBK-Tech (see image) look very similar to one another. Both products have similar looking names and overlapping dosage strengths. The containers use similar colors and are likely to be stored near one another on pharmacy shelves. When possible, explore ordering one of the products from a different manufacturer to prevent a mix-up from occurring. We have notified the US Food and Drug Administration (FDA) and the manufacturer of this issue.