Antibiotics and chemotherapeutics are used widely in swine feeds. They are effective in improving the rate and efficiency of growth and in reducing mortality and morbidity associated with respiratory and intestinal diseases in pigs. Certain feed additives require a withdrawal period prior to slaughter in order to insure that residues do not occur in the carcass. Approved feed additives that require withdrawal and their withdrawal times are given in Table 1. In addition, Table 1 includes those feed additives that do not require any withdrawal. A brief history of sulfa residues illustrates the common causes and preventive measures needed to provide residue-free pork.

Forms of Sulfonamides Used in Feeds

The only sulfonamides that can be used legally in feeds are sulfamethazine and sulfathiazole, and they are approved only in combination with certain other antibiotics and only at one level of inclusion (100 grams per ton). Some commercial products that include sulfonamides are Aureomix 500 (chlorotetracycline, penicillin and sulfamethazine), Tylan 40 Sulfa-G (tylosin and sulfamethazine), and CSP 500 Fermazole (chlorotetracycline, penicillin and sulfathiazole). These products are considered Type A drugs and they can only be handled and mixed by feed manufacturers that possess FD-1900 permits. However, producers can obtain and use blended products that have lower concentrations of these sulfa drugs. Sulfamethazine, sulfathiazole, and certain other sulfonamides sometimes are used as water medications for controlling pneumonia, scours and other bacterial infections.

Efficacy of Sulfonamides

Sulfonamides are used primarily for young pigs during the early growth stages. Most pig starter feeds and about 75% of grower feeds are medicated. It is estimated that about 10% to 20% of these medicated feeds contain sulfamethazine or sulfathiazole. One reason for the popularity of the sulfonamide-antibiotic combinations is that they are very effective growth promoters (Table 2). A summary of 453 experiments involving 13,632 pigs indicates that pigs fed sulfa-antibiotic combinations from 19 lb to 56 lb gained 20.5% faster and required 7.8% less feed per pound of gain than control pigs that received no antibiotics. For 10 other antibiotics, the average improvements in daily gain and efficiency of feed utilization were less than the sulfa-antibiotic combinations, 13.8% and 6.5%, respectively. Similar trends were found in a summary of 280 experiments involving slightly heavier pigs, fed from 37 lb to 109 lb (Table 2).

The sulfa-containing feed additives also have been shown to help maintain acceptable performance in herds having acute or chronic respiratory infections, such as atrophic rhinitis.

Sulfonamide Residues

The tolerance level for sulfamethazine and sulfathiazole in pork tissue (liver, kidney, or muscle) is set by the Food and Drug Administration at 0.1 ppm. Regu-
lations in the U.S. require that sulfamethazine be withdrawn from the feed for 15 days and sulfathiazole for 7 days prior to slaughter in order to insure that tissues do not exceed the tolerance level.

Shortly after a national monitoring program was initiated by the USDA in the 1970's, it was discovered that about 15% of hog carcasses had violative sulfonamide residues. In almost all cases, the sulfonamide found in the carcass tissues was sulfamethazine. A major effort was initiated in 1977 by the USDA, the Cooperative Extension Service and the National Pork Producers Council to solve this problem by means of research and educational programs. Additional testing of carcasses for residues at packing plants was implemented in 1987 with stiff penalties for producers marketing hogs with violative residues. Although the problem has not been completely solved, the violation rate for the past 5 years has been less than 1% (in 1994, it was .86%, 11 of 1,283 samples) according to the USDA surveillance data (Figure 1). The incidence of sulfa residues based on the Sulfa-on-Site (SOS) testing program in 1994 was .06% (104 of 166,091 samples), but this is based on muscle rather than liver samples. Sulfamethazine residues in liver are generally about four times higher than they are in muscle.

Causes of Sulfonamide Residues

What was the reason for the high incidence of sulfamethazine residues, and why has it been so difficult to eliminate the problem? Initially, producers were blamed for not complying with the withdrawal period. However, it was soon realized that some of the violations were from farms where producers were making a sincere effort to follow proper withdrawal times. In some cases, violations were even being reported on farms where sulfonamides were not being used in feed or water.

Finally, results of research conducted at Iowa State University, the University of Illinois and the University of Kentucky shed new light on the problem. Their studies showed that only a very small amount of sulfamethazine in the feed would cause a residue in the tissue. An early study at the University of Kentucky indicated that as little as 1 gram of sulfamethazine per ton of feed could result in a high incidence of violative residues. Table 3 illus-

<table>
<thead>
<tr>
<th>Additives</th>
<th>Withdrawal time</th>
</tr>
</thead>
<tbody>
<tr>
<td>Carboxad</td>
<td>70 days</td>
</tr>
<tr>
<td>Apramycin</td>
<td>28 days</td>
</tr>
<tr>
<td>Neomycin</td>
<td>5-10 days</td>
</tr>
<tr>
<td>Sulfamethazine</td>
<td>15 days</td>
</tr>
<tr>
<td>Sulfathiazole</td>
<td>7 days</td>
</tr>
<tr>
<td>Lincomycin</td>
<td>6 days</td>
</tr>
<tr>
<td>Arsanilic acid, sodium arsanilate</td>
<td>5 days</td>
</tr>
<tr>
<td>Roxarsone (3-Nitro)</td>
<td>5 days</td>
</tr>
<tr>
<td>Dewormers</td>
<td></td>
</tr>
<tr>
<td>Thiabendazole</td>
<td>30 days</td>
</tr>
<tr>
<td>Hygromycin B</td>
<td>15 days</td>
</tr>
<tr>
<td>Ivomecin</td>
<td>5 days</td>
</tr>
<tr>
<td>Levamisole hydrochloride (Tramisol)</td>
<td>72 hours</td>
</tr>
<tr>
<td>Pyrantel tartrate (Banminth)</td>
<td>24 hours</td>
</tr>
</tbody>
</table>

Additives requiring withdrawal

Additives requiring no withdrawal

Antimicrobial Agents
- Bacitracin, zinc
- Bacitracin methylene disalicylate
- Banbermycins
- Chlorotetracycline
- Oxytetracycline
- Penicillin
- Tiamulin
- Tylosin
- Virginiamycin

Dewormers
- Dichlorvos (Atgard)
- Fenbendazole (Safe-Guard)
- Larvicide
- Rabon

Table 1. Feed additives that require withdrawal and those requiring no withdrawal from swine feed.

![Sulfanamide Residues in Pork Liver 1977 to 1994](image)

Figure 1. Incidence of sulfonamide residues in pork liver from 1977 to 1994.

Table 2. Comparison of antibiotics as growth promoters for young pigs.

<table>
<thead>
<tr>
<th>Item</th>
<th>Number of experiments</th>
<th>Avg. daily gain</th>
<th>Feed/gain</th>
</tr>
</thead>
</table>
| Starting pigs (19 to 56 lb.)
- Antibiotic-sulfa combinations
- Other antibiotics | 131 | 20.5 | 7.8 |
| 322 | 13.8 | 6.5 |
| Growing pigs (37 to 109 lb.)
- Antibiotic-sulfa combinations
- Other antibiotics | 32 | 15.4 | 5.7 |
| 48 | 10.7 | 4.6 |

1Feed Additive Compendium, 1996.
2Approved only in combination with certain other antimicrobial agents.
3Ten-day withdrawal when high level (140 grams per ton) is used.
4Five-day withdrawal when high level (500 grams per ton) used.
5Two-day withdrawal when high level (35 grams per ton) used.

1Data from 453 experiments, 13,632 pigs.
2Auro-SP-250, Chlorachel-250, Tylan-Sulfa and CSP-250.
3Data from 280 experiments, 5,763 pigs.
Table 3. Effects of form (sulfamethazine vs. sulfathiazole) and level of sulfa in finisher feed on sulfa residues in pork.

<table>
<thead>
<tr>
<th>Form and level of sulfa</th>
<th>Sulfa residue Liver ppm</th>
<th>Muscle ppm</th>
<th>Violations Liver %</th>
<th>Muscle %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sulfamethazine&lt;sup&gt;3&lt;/sup&gt; in feed, g/ton</td>
<td>0</td>
<td>&lt;.01</td>
<td>&lt;.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>1</td>
<td>.04</td>
<td>.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>2</td>
<td>.09</td>
<td>.09</td>
<td>38</td>
</tr>
<tr>
<td></td>
<td>4</td>
<td>.20</td>
<td>.05</td>
<td>100</td>
</tr>
<tr>
<td></td>
<td>8</td>
<td>.43</td>
<td>.09</td>
<td>100</td>
</tr>
<tr>
<td></td>
<td>16</td>
<td>.88</td>
<td>.19</td>
<td>100</td>
</tr>
<tr>
<td></td>
<td>100</td>
<td>4.55</td>
<td>1.52</td>
<td>100</td>
</tr>
<tr>
<td>Sulfathiazole&lt;sup&gt;4&lt;/sup&gt; in feed, g/ton</td>
<td>0</td>
<td>.1</td>
<td>&lt;.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>1</td>
<td>&lt;.01</td>
<td>&lt;.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>2</td>
<td>&lt;.01</td>
<td>&lt;.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>4</td>
<td>&lt;.01</td>
<td>&lt;.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>8</td>
<td>.03</td>
<td>.01</td>
<td>0</td>
</tr>
<tr>
<td></td>
<td>16</td>
<td>.03</td>
<td>.02</td>
<td>20</td>
</tr>
<tr>
<td></td>
<td>100</td>
<td>.03</td>
<td>.05</td>
<td>78</td>
</tr>
</tbody>
</table>

<sup>1</sup>University of Kentucky and University of Nebraska. 1981, 16 pigs/treatment
<sup>2</sup>Percent of samples having 0.1 ppm or more of sulfa, based on two assay methods: colorimetric (corrected for background) and GLC.
<sup>3</sup>Sixteen pigs per treatment were fed 100 grams of sulfamethazine per ton for 2 weeks, then these levels were fed for 15 days prior to slaughter.
<sup>4</sup>Sixteen pigs per treatment were fed 100 grams of sulfathiazole per ton for 2 weeks, then these levels were fed for 7 days prior to slaughter.

Table 3 indicates that feed containing up to 16 grams of sulfathiazole per ton, on a continuous basis, before a violative residue occurs. A major cause of the high incidence of sulfonamide residues was, and still is, due to the cross-mixing of clean feed with sulfonamide-containing feed. Drug carry-over can occur in commercial feed mills and on the farm. It can also result from the inadvertent purchase of sulfonamide-containing premixes and supplements. As little as 40 lb of a sulfamethazine-medicated feed (containing 100 grams per ton), if mixed into a ton of “clean” feed, will result in a feed containing 2 grams of sulfamethazine per ton—a carry-over level that can leave a violative residue of sulfamethazine in liver tissue.

**Preventing Drug Carry-Over in Feeds**

Drug carry-over in feeds can occur in a number of ways. Feed manufacturing equipment such as mixers, pellet mills, augers, elevator legs, dust control devices and storage bins can harbor dust and residual feed, which can carry-over into clean feed (Figures 2, 3, 4 and 5). A vertical screw mixer may contain 40 lb to 50 lb of residual feed in the boot after the feed is discharged. Failure to remove this residual feed will cause the next batch to be contaminated. In some farm mixers, such as portable grinder-mixers, 100 lb of residual feed per batch can remain. A thorough clean-out or flush of all mixing equipment, conveyors, augers, elevator legs, and similar equipment is imperative to reduce the chance of drug carry-over. Some producers use a second set of equipment for mixing finishing feeds containing feed additives with a required withdrawal to solve the drug residue problem.
Figure 4. Elevator legs can hold sizable amounts of residual feeds or ingredients. Some of this material can be incorporated in the next batch of feed.

A proper feed mixing sequence will reduce the degree of drug carry-over. For example, a finishing feed should never immediately follow a feed containing medication requiring a withdrawal time. Instead, it should be followed with a feed that is less likely to cause residue problems, such as a grower feed.

The powdered form of the sulfonamides tends to be electrostatic and will cling to metal surfaces. Grounding of equipment will reduce this characteristic, but will not completely eliminate it. Fortunately, granulated forms of sulfanamide (the forms that are present in all commercial antibiotic-sulfanamide mixes) have helped to reduce this problem. In a study at the University of Kentucky, the sulfa level in feed dust taken from the inside surface of a mixer was 276 ppm when powdered sulfamethazine was used as compared with only 59 ppm when granulated sulfamethazine was used.

Feed should never be medicated with powdered sulfonamides. This is an illegal practice and it is likely to cause residue problems. Excessive dust and waste feed should not be allowed to accumulate around feed mixing and handling equipment, as they can be a source of drug carry-over. Accumulated dust should be removed at regular intervals and discarded; it should not be included in mixed finishing feed.

Bulk delivery trucks also can be responsible for drug carry-over in feeds if medicated and nonmedicated feeds are hauled at the same time or if the conveying system on these trucks is not cleaned out well between delivery of medicated feed and delivery of nonmedicated feed (Figure 6). Bulk storage bins on the farm should never be used for both medicated feed and nonmedicated feed unless they are thoroughly cleaned between batches. Feed tends to cling to the sides and corners of the bins (Figures 7 and 8) and in the discharge augers. Drug carry-over can occur in these structures if they are not completely emptied and properly cleaned between batches of feed. Pig feeders should be emptied and cleaned after sulfanamide-medicated feeds are used, if the same feeders are used to finish out pigs. If feeders are not cleaned out completely, medicated feed can build up in certain parts of feeders (Figure 9) and can contaminate several batches of nonmedicated feed. If a thorough clean-out and flushing of the feed delivery system in a
building is not possible, then separate delivery systems are recommended for sulfonamide-medicated and nonmedicated feeds. Another alternative is to completely avoid the use of any sulfonamide-medicated feed in every building that houses finishing pigs.

The same contamination principles hold true for water medicators. Care should be taken to prevent contamination of clean water with medicated water. Also, one should not medicate the feed and the water with the same drug at the same time. This practice could cause high intakes of the compound and could result in a residue, even with published withdrawal times.

Proper Mixing of Feeds

Producers who mix their own feed on the farm must follow good feed mixing practices to insure uniform dispersal of drugs and other microingredients in feed. Adequate mixing time is a must. Both undermixing and overmixing should be avoided. Recommended mixing time for vertical mixers is 15 to 20 minutes and for horizontal mixers is 5 to 6 minutes per batch. Accurate scales must be used. Volumetric mills must be calibrated often (at least once a week) to insure proper inclusion rates of ingredients. Producers must be certain that only approved levels of drugs and approved combinations of drugs are used in feeds. Levels and combinations of drugs are regulated by the FDA and are published in the Feed Additive Compendium (Miller Publishing Co., Minnetonka, MN 55343).

Producers should use a record system to keep track of their medicated feeds. An example of one is shown in Figure 10. A good record system also will help to avoid mixing errors.

Preventing Access to Feed Additive-Containing Manure

Studies at the University of Illinois and Iowa State University indicate that sulfonamide residues in pork can be caused by pigs having access to sulfamethazine-containing manure. Pigs housed on solid floors that allow accumulation of manure and urine are more likely to pick up sulfonamides from the floors than those housed on slotted floors. Lagoons that receive waste from buildings where sulfonamides are being used can be a source of contamination when lagoon water is used in finishing house flush systems.

Following drug withdrawal, pigs should be moved to a clean pen or the pen should be thoroughly cleaned at the time of withdrawal. These pens should be cleaned 3 to 7 days following medication withdrawal. Pigs should not be allowed to have access to manure in trucks, holding pens, etc., where other hogs that may have had withdrawal requiring products in their feed were kept. Holding pens that allow pooling of urine should be avoided before and during marketing.

Adherence to Withdrawal

Producers must be certain that they comply with the proper withdrawal periods. Water medications may require longer withdrawal periods. To be on the safe side, it is best to include sulfonamides only in the starter feed. If sulfonamides are used in grower feed, they should not be used beyond 125 lb body weight and should be left out of the finisher feed. Some producers finish their pigs in a separate building and avoid the use of sulfonamides and other feed additives in the finishing building. This practice also solves the recycling problem caused by contaminated manure. Sows and gilts that are sent to slaughter also can contribute to residue problems if withdrawal times are not adhered to.

Some have suggested that finishing pigs be fed only corn for several days prior to slaughter. While this practice might help insure that sulfonamides will not be present in the preslaughter feed, it may be a costly
Date mixed | Tank number | Description of feed | Tons | Medication | g/ton
--- | --- | --- | --- | --- | ---
6-14-90 | 2 | Gestation, 14% | 3 | — | —
6-16-90 | 1 | Starter, 18% | 1 | Tylan-Sulfapyridine | 100-100
6-16-90 | 3 | Lactation, 14% | 3 | Neo-tetraacycline | 50-50
6-21-90 | 4 | Finisher, 13% | 5 | Aureomycin | 50

Figure 10. A feed mixing record sheet will help to eliminate mixing errors and help producers keep track of medicated feeds.

practice. Corn is extremely low in lysine and other amino acids, so growth rate and feed conversion will be markedly reduced by feeding only shelled corn for any extended period of time, even to finishing pigs. However, this practice might be feasible as a last resort for producers having a serious residue problem, provided manure or urine are not allowed to accumulate in the pen.

Testing Live Hogs for Sulfonamide Residues

Kits are available for on-the-farm testing of live pigs for potential residues. In these tests, urine is collected and tested. Producers who anticipate a potential problem with sulfonamide residues or who are interested in their sulfonamide residue status can test a few pigs before they go to slaughter to insure that they are free of residues. When help is needed, swine practitioners can perform these tests or have access to persons who can perform the tests. Sulfamethazine test kits are available from the following company:

Editek, Inc.
1238 Anthony Road
Burlington, NC 77215

Pork Quality Assurance

The National Pork Producers Council has developed a three-stage Pork Quality Assurance program which is designed to assist pork producers in eliminating drug residues. The basis for the program is education and management changes followed by voluntary testing for residues. Information on the program is available from the National Pork Producers Council, Box 10383, Des Moines, IA 50306.

Summary

Drug residues in pork carcasses are a deterrent to consumer acceptability of pork and to international sales of pork. Drug residues can be greatly reduced and even be eliminated by adherence to the following practices.

1. Use only approved levels and combinations of drugs.
2. Follow good feed mixing practices (especially adequate mixing time) to insure that feed is mixed properly.
3. Maintain a record system to keep track of drug premixes and medicated feed usage.
4. Mix batch feeds in proper sequence to reduce the chance of carry-over of drugs into finishing feed.
5. Clean out or flush feed mixing, conveying and feeding equipment to reduce drug carry-over into finishing feeds.
6. Adhere to proper withdrawal periods for drugs.
7. Prevent recycling of drugs via manure and urine.
8. Use an on-farm testing program to insure freedom from drug residues.
10. Inform other pork producers how to check for residues and the problems associated with residues.

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