



Calms inflammatory acne

- ACZONE® (dapson) Gel 5% is **dapson**—the well-known, powerful anti-inflammatory—in **topical** form for treatment of acne vulgaris
- ACZONE® is indicated for the **topical treatment of acne vulgaris**—a pea-sized dose should be applied twice daily
- **First in its class.** First new chemical entity for topical treatment of acne **in a decade**
- Rx acne treatment market is substantial—nearly 20 million prescriptions valued at more than \$2.7 billion in the 12 months from July 2007 to June 2008<sup>1</sup>
- Dermatologists look for new, effective treatments for inflammatory acne
- Unlike oral dapson, **NO blood test required.**<sup>2</sup> NO evidence of clinically relevant hemolysis in testing<sup>2</sup>
- 3010 subjects participated in clinical trials—substantial evidence of efficacy and safety<sup>3</sup>
- Favorable safety profile—equivalent to vehicle<sup>3</sup>
- Supplied in 30-g tubes

**48% reduction in inflammatory acne** in 12 weeks.<sup>3</sup>  
24% reduction in inflammatory acne in just 2 weeks!<sup>3</sup>

After 12 weeks, 41% of patients had minimal or no acne.<sup>3</sup>

**ACZONE® (dapson) Gel 5% is indicated for the topical treatment of acne vulgaris.**

### Important Safety Information WARNINGS AND PRECAUTIONS

**Hematological effects:** Oral dapson treatment has produced dose-related hemolysis and hemolytic anemia. Individuals with glucose-6-phosphate dehydrogenase (G6PD) deficiency are more prone to hemolysis with the use of certain drugs. There was no evidence of clinically relevant hemolysis or anemia in patients treated with ACZONE® Gel 5% including patients who were G6PD deficient. Some patients with G6PD deficiency using ACZONE® Gel 5% developed laboratory changes suggestive of mild hemolysis.

If signs and symptoms suggestive of hemolytic anemia occur, ACZONE® Gel 5% should be discontinued. ACZONE® Gel 5% should not be used in patients who are taking oral dapson or antimalarial medications because of the potential for hemolytic reactions. Combination of ACZONE® Gel 5% with trimethoprim/sulfamethoxazole (TMP/SMX) may increase the likelihood of hemolysis in patients with G6PD deficiency.

Supported nationally by more than 60 Allergan professional representatives—it is also Allergan’s intention to promote ACZONE® Gel to pediatricians and PCPs through a copromotional effort

Available in Q4 2008





Calms  
inflammatory  
acne

## Important Safety Information

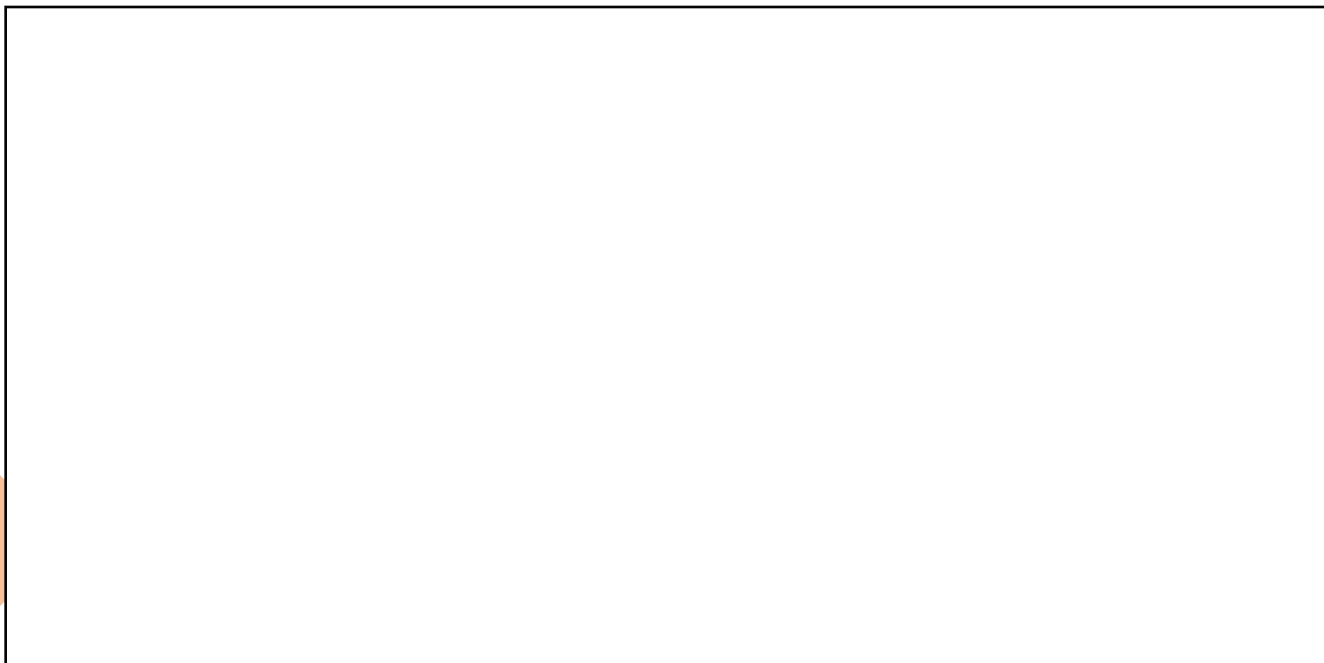
**Peripheral neuropathy:** Peripheral neuropathy (motor loss and muscle weakness) has been reported with oral dapsone treatment. No events of peripheral neuropathy were observed in clinical trials with topical ACZONE® Gel 5% treatment.

**Skin:** Skin reactions (toxic epidermal necrolysis, erythema multiforme, morbilliform and scarlatiniform reactions, bullous and exfoliative dermatitis, erythema nodosum, and urticaria) have been reported with oral dapsone treatment. These types of skin reactions were not observed in clinical trials with topical ACZONE® Gel 5% treatment.

## ADVERSE REACTIONS

The most common adverse reactions of ACZONE® Gel 5% (incidence  $\geq 10\%$ ) are oiliness/peeling, dryness, and erythema at the application site.

**Please see accompanying full prescribing information.**



1. Verispan LLC. *Vector One®: National (VONA)*. Yardley, PA: Verispan LLC; June 2008.

2. ACZONE® Prescribing Information.

3. Draelos ZD, Carter E, Maloney JM, et al; for United States/Canada Dapsone Gel Study Group. Two randomized studies demonstrate the efficacy and safety of dapsone gel, 5% for the treatment of acne vulgaris. *J Am Acad Dermatol*. 2007;56(3):439.e1-439.e10.

# ACZONE® (dapsone) Gel 5%



## HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use ACZONE® safely and effectively. See full prescribing information for ACZONE®.

ACZONE® (dapsone) Gel, 5%

For topical use only  
Initial U.S. Approval: 1955

### INDICATIONS AND USAGE

ACZONE® Gel is indicated for the topical treatment of acne vulgaris (1).

### DOSAGE AND ADMINISTRATION

- Apply twice daily (2).
- Apply approximately a pea-sized amount of ACZONE® Gel, 5%, in a thin layer to the acne affected area (2).
- If there is no improvement after 12 weeks, treatment with ACZONE® Gel, 5%, should be reassessed (2).

For topical use only. Not for oral, ophthalmic, or intravaginal use (2).

### DOSAGE FORMS AND STRENGTHS

ACZONE® (dapsone) Gel, 5%, is supplied in the following size tubes:

- Professional Sample: 3 gram laminate tube (3).
- Commercially: 30 gram laminate tube (3).

### CONTRAINDICATIONS

None.

## FULL PRESCRIBING INFORMATION: CONTENTS\*

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### WARNINGS AND PRECAUTIONS

Hematological Effects: Some subjects with G6PD deficiency using ACZONE® Gel developed laboratory changes suggestive of mild hemolysis. (5.1)(8.6)

The following are seen with oral dapsone treatment:

- Hematological Effects (5.1).
- Peripheral Neuropathy (5.2).
- Skin Reactions (5.3).

### ADVERSE REACTIONS

Most common adverse reactions (incidence ≥ 10%) are oiliness/peeling, dryness and erythema at the application site (6).

To report SUSPECTED ADVERSE REACTIONS, contact Allergan at 1-800-433-8871 or www.allergan.com or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

### DRUG INTERACTIONS

- Trimethoprim/sulfamethoxazole (TMP/SMX) increases the level of dapsone and its metabolites. (7.1)
- Topical benzoyl peroxide used at the same time as ACZONE® may result in temporary local yellow or orange skin discoloration. (7.2)

### USE IN SPECIFIC POPULATIONS

G6PD Deficiency (8.6).

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Revised: 09/2008

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\*Sections or subsections omitted from the full prescribing information are not listed.

Serious adverse reactions reported in patients treated with ACZONE® Gel, 5%, during clinical trials included but were not limited to the following:

- Nervous system/Psychiatric – Suicide attempt, tonic clonic movements.
- Gastrointestinal – Abdominal pain, severe vomiting, pancreatitis.
- Other – Severe pharyngitis

In the clinical trials, a total of 12 out of 4032 patients were reported to have depression (3 of 1660 treated with vehicle and 9 of 2372 treated with ACZONE® Gel, 5%). Psychosis was reported in 2 of 2372 patients treated with ACZONE® Gel, 5%, and in 0 of 1660 patients treated with vehicle.

Combined contact sensitization/irritation studies with ACZONE® Gel, 5%, in 253 healthy subjects resulted in at least 3 subjects with moderate erythema. ACZONE® Gel, 5%, did not induce phototoxicity or photoallergy in human dermal safety studies.

ACZONE® Gel, 5%, was evaluated for 12 weeks in four controlled studies for local cutaneous events in 1819 patients. The most common events reported from these studies include oiliness/peeling, dryness, and erythema. These data are shown by severity in Table 1 below.

Table 1 – Application Site Adverse Reactions by Maximum Severity

Application Site Event	ACZONE® (N=1819)			Vehicle (N=1660)		
	Mild	Moderate	Severe	Mild	Moderate	Severe
Erythema	9%	5%	<1%	9%	6%	<1%
Dryness	14%	3%	<1%	14%	4%	<1%
Oiliness/Peeling	13%	6%	<1%	15%	6%	<1%

The adverse reactions occurring in at least 1% of patients in either arm in the four vehicle controlled studies are presented in Table 2.

Table 2 – Adverse Reactions Occurring in at least 1% of Patients

	ACZONE® N=1819	Vehicle N=1660
Application Site Reaction NOS	18%	20%
Application Site Dryness	16%	17%
Application Site Erythema	13%	14%
Application Site Burning	1%	2%
Application Site Pruritus	1%	1%
Pyrexia	1%	1%
Nasopharyngitis	5%	6%
Upper Respiratory Tract Inf. NOS	3%	3%
Sinusitis NOS	2%	1%
Influenza	1%	1%
Pharyngitis	2%	2%
Cough	2%	2%
Joint Sprain	1%	1%
Headache NOS	4%	4%

NOS = Not otherwise specified

One patient treated with ACZONE® Gel in the clinical trials had facial swelling which led to discontinuation of medication.

In addition, 486 patients were evaluated in a 12 month safety study. The adverse event profile in this study was consistent with that observed in the vehicle-controlled studies.

## 6.2 Experience with Oral Use of Dapsone

Although not observed in the clinical trials with ACZONE® Gel (topical dapsone) serious adverse reactions have been reported with oral use of dapsone, including agranulocytosis, hemolytic anemia, peripheral neuropathy (motor loss and muscle weakness), and skin reactions (toxic epidermal necrolysis, erythema multiforme, morbilliform and scarlatiniform reactions, bullous and exfoliative dermatitis, erythema nodosum, and urticaria).

## 7 DRUG INTERACTIONS

### 7.1 Trimethoprim-Sulfamethoxazole

A drug-drug interaction study evaluated the effect of the use of ACZONE® Gel, 5%, in combination with double strength (160 mg/800 mg) trimethoprim-sulfamethoxazole (TMP/SMX). During co-administration, systemic levels of TMP and SMX were essentially unchanged. However, levels of dapsone and its metabolites increased in the presence of TMP/SMX. Systemic exposure (AUC<sub>0-24</sub>) of dapsone and N-acetyl-dapsone (NAD) were increased by about 40% and 20% respectively in presence of TMP/SMX. Notably, systemic exposure (AUC<sub>0-24</sub>) of dapsone hydroxylamine (DHA) was more than doubled in the presence of TMP/SMX. Exposure from the proposed topical dose is about 1% of that from the 100 mg oral dose, even when co-administered with TMP/SMX.

### 7.2 Topical Benzoyl Peroxide

Topical application of ACZONE® Gel followed by benzoyl peroxide in subjects with acne vulgaris resulted in a temporary local yellow or orange discoloration of the skin and facial hair (reported by 7 out of 95 subjects in a clinical study) with resolution in 4 to 57 days.

### 7.3 Drug Interactions with Oral Dapsone

Certain concomitant medications (such as rifampin, anticonvulsants, St. John's wort) may increase the formation of dapsone hydroxylamine, a metabolite of dapsone associated with hemolysis. With oral dapsone treatment, folic acid antagonists such as pyrimethamine have been noted to possibly increase the likelihood of hematologic reactions.

## 8 USE IN SPECIFIC POPULATIONS

### 8.1 Pregnancy

#### Teratogenic Effects: Pregnancy Category C

There are no adequate and well controlled studies in pregnant women. Dapsone has been shown to have an embryocidal effect in rats and rabbits when administered orally in doses of 75 mg/kg/day and 150 mg/kg/day (approximately 800 and 500 times the systemic exposure observed in human females as a result of use of the maximum recommended topical dose, based on AUC comparisons), respectively. These effects were probably secondary to maternal toxicity. ACZONE® Gel, 5%, should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

### 8.3 Nursing Mothers

Although systemic absorption of dapsone following topical application of ACZONE® Gel, 5%, is minimal relative to oral dapsone administration, it is known that dapsone is excreted in human milk. Because of the potential for oral dapsone to cause adverse reactions in nursing infants, a decision should be made whether to discontinue nursing or to discontinue ACZONE® Gel, 5%, taking into account the importance of the drug to the mother.

### 8.4 Pediatric Use

Safety and efficacy was evaluated in 1169 children aged 12-17 years old treated with ACZONE® Gel, 5%, in the clinical studies. The adverse event rate for ACZONE® Gel, 5%, was similar to the vehicle control group. Safety and efficacy was not studied in pediatric patients less than 12 years of age, therefore ACZONE® Gel, 5%, is not recommended for use in this age group.

### 8.5 Geriatric Use

Clinical studies of ACZONE® Gel, 5%, did not include sufficient number of patients aged 65 and over to determine whether they respond differently from younger patients.

### 8.6 G6PD Deficiency

ACZONE® Gel, 5% and vehicle were evaluated in a randomized, double-blind, cross-over design clinical study of 64 patients with G6PD deficiency and acne vulgaris. Subjects were Black (88%), Asian (6%), Hispanic (2%) or of other racial origin (5%). Blood samples were taken at Baseline, Week 2, and Week 12 during both vehicle and ACZONE® Gel, 5% treatment periods. There were 56 out of 64 subjects who had a Week 2 blood draw and applied at least 50% of treatment applications. Table 3 contains results from testing of relevant hematology parameters for these two treatment periods. ACZONE® Gel was associated with a 0.32 g/dL drop in hemoglobin after two weeks of treatment, but hemoglobin levels generally returned to baseline levels at Week 12.

## FULL PRESCRIBING INFORMATION

### 1 INDICATIONS AND USAGE

ACZONE® Gel, 5%, is indicated for the topical treatment of acne vulgaris.

### 2 DOSAGE AND ADMINISTRATION

For topical use only. Not for oral, ophthalmic, or intravaginal use.

After the skin is gently washed and patted dry, apply approximately a pea-sized amount of ACZONE® Gel, 5%, in a thin layer to the acne affected areas twice daily. Rub in ACZONE® Gel, 5%, gently and completely. ACZONE® Gel, 5%, is gritty with visible drug substance particles. Wash hands after application of ACZONE® Gel, 5%.

If there is no improvement after 12 weeks, treatment with ACZONE® Gel, 5%, should be reassessed.

### 3 DOSAGE FORMS AND STRENGTHS

ACZONE® (dapsone) Gel, 5%, is supplied in the following size tubes:

- Professional Sample: 3 gram laminate tube
- Commercially: 30 gram laminate tube

### 4 CONTRAINDICATIONS

None.

### 5 WARNINGS AND PRECAUTIONS

#### 5.1 Hematological Effects

Oral dapsone treatment has produced dose-related hemolysis and hemolytic anemia. Individuals with glucose-6-phosphate dehydrogenase (G6PD) deficiency are more prone to hemolysis with the use of certain drugs. G6PD deficiency is most prevalent in populations of African, South Asian, Middle Eastern and Mediterranean ancestry.

There was no evidence of clinically relevant hemolysis or anemia in patients treated with ACZONE® Gel, 5%, including patients who were G6PD deficient. Some subjects with G6PD deficiency using ACZONE® Gel developed laboratory changes suggestive of mild hemolysis.

If signs and symptoms suggestive of hemolytic anemia occur, ACZONE® Gel, 5% should be discontinued. ACZONE® Gel, 5% should not be used in patients who are taking oral dapsone or antimalarial medications because of the potential for hemolytic reactions. Combination of ACZONE® Gel, 5%, with trimethoprim/sulfamethoxazole (TMP/SMX) may increase the likelihood of hemolysis in patients with G6PD deficiency.

#### 5.2 Peripheral Neuropathy

Peripheral neuropathy (motor loss and muscle weakness) has been reported with oral dapsone treatment. No events of peripheral neuropathy were observed in clinical trials with topical ACZONE® Gel, 5% treatment.

#### 5.3 Skin

Skin reactions (toxic epidermal necrolysis, erythema multiforme, morbilliform and scarlatiniform reactions, bullous and exfoliative dermatitis, erythema nodosum, and urticaria) have been reported with oral dapsone treatment. These types of skin reactions were not observed in clinical trials with topical ACZONE® Gel, 5% treatment.

### 6 ADVERSE REACTIONS

#### 6.1 Clinical Studies Experience

Because clinical trials are conducted under prescribed conditions, adverse reaction rates observed in the clinical trials of a drug cannot be directly compared to rates in the clinical trials of another drug and may not reflect the rates observed in practice.

Table 3 – Mean Hemoglobin, Bilirubin, and Reticulocyte Levels in Acne Subjects with G6PD Deficiency in ACZONE®/Vehicle Cross-Over Study

		ACZONE®		Vehicle	
		N	Mean	N	Mean
Hemoglobin (g/dL)	Pre-treatment	53	13.44	56	13.36
	2 weeks	53	13.12	55	13.34
	12 weeks	50	13.42	50	13.37
Bilirubin (mg/dL)	Pre-treatment	54	0.58	56	0.55
	2 weeks	53	0.65	55	0.56
	12 weeks	50	0.61	50	0.62
Reticulocytes (%)	Pre-treatment	53	1.30	55	1.34
	2 weeks	53	1.51	55	1.34
	12 weeks	50	1.48	50	1.41

There were no changes from baseline in haptoglobin or lactate dehydrogenase during ACZONE® or vehicle treatment at either the 2-week or 12-week time point.

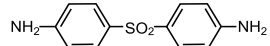
The proportion of subjects who experienced decreases in hemoglobin  $\geq 1$  g/dL was similar between ACZONE® Gel, 5% and vehicle treatment (8 of 58 subjects had such decreases during ACZONE® treatment compared to 7 of 56 subjects during vehicle treatment among subjects with at least one on-treatment hemoglobin assessment). Subgroups based on gender, race, or G6PD enzyme activity did not display any differences in laboratory results from the overall study group. There was no evidence of clinically significant hemolytic anemia in this study. Some of these subjects developed laboratory changes suggestive of mild hemolysis.

## 10 OVERDOSAGE

ACZONE® Gel, 5%, is not for oral use. If oral ingestion occurs, medical advice should be sought.

## 11 DESCRIPTION

ACZONE® Gel, 5%, contains dapsonone, a sulfone, in an aqueous gel base for topical dermatologic use. ACZONE® Gel, 5% is a gritty translucent material with visible drug substance particles. Chemically, dapsonone has an empirical formula of C<sub>12</sub>H<sub>12</sub>N<sub>2</sub>O<sub>2</sub>S. It is a white, odorless crystalline powder that has a molecular weight of 248. Dapsonone's chemical name is 4,4'-diaminodiphenylsulfone and its structural formula is:



Each gram of ACZONE® Gel, 5%, contains 50 mg of dapsonone, USP, in a gel of carbomer 980; diethylene glycol monoethyl ether, NF; methylparaben, NF; sodium hydroxide, NF; and purified water, USP.

## 12 CLINICAL PHARMACOLOGY

### 12.1 Mechanism of Action

The mechanism of action of dapsonone gel in treating acne vulgaris is not known.

### 12.3 Pharmacokinetics

An open-label study compared the pharmacokinetics of dapsonone after ACZONE® Gel, 5%, (110  $\pm$  60 mg/day) was applied twice daily (~BSA 22.5%) for 14 days (n=18) with a single 100 mg dose of oral dapsonone administered to a subgroup of patients (n=10) in a crossover design. On Day 14 the mean dapsonone AUC<sub>0-24h</sub> was 415  $\pm$  224 ng•h/mL for ACZONE® Gel, 5%, whereas following a single 100 mg dose of oral dapsonone the AUC<sub>0-24h</sub> was 52,641  $\pm$  36,223 ng•h/mL. Exposure after the oral dose of 100 mg dapsonone was approximately 100 times greater than after the topical ACZONE® Gel, 5% dose, twice a day.

In a long-term safety study of ACZONE® Gel, 5% treatment, periodic blood samples were collected up to 12 months to determine systemic exposure of dapsonone and its metabolites in approximately 500 patients. Based on the measurable dapsonone concentrations from 408 patients (M=192, F=216), obtained at month 3, neither gender, nor race appeared to affect the pharmacokinetics of dapsonone. Similarly, dapsonone exposures were approximately the same between the age groups of 12-15 years (N=155) and those greater than or equal to 16 years (N=253). There was no evidence of increasing systemic exposure to dapsonone over the study year in these patients.

### 12.4 Microbiology

*In Vivo* Activity: No microbiology or immunology studies were conducted during dapsonone gel clinical trials.

**Drug Resistance:** No dapsonone resistance studies were conducted during dapsonone gel clinical trials. Because no microbiology studies were done, there are no data available as to whether dapsonone treatment may have resulted in decreased susceptibility of *Propionibacterium acnes*, an organism associated with acne, to other antimicrobials that may be used to treat acne. Therapeutic resistance to dapsonone has been reported for *Mycobacterium leprae*, when patients have been treated with oral dapsonone.

## 13 NONCLINICAL TOXICOLOGY

### 13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility

Dapsonone was not mutagenic in a bacterial reverse mutation assay (Ames test) using *S. typhimurium* and *E. coli*, with and without metabolic activation and was negative in a micronucleus assay conducted in mice. Dapsonone increased both numerical and structural aberrations in a chromosome aberration assay conducted with Chinese hamster ovary (CHO) cells.

Dapsonone was not carcinogenic to rats when orally administered to females for 92 weeks or males for 100 weeks at dose levels up to 15 mg/kg/day (approximately 160 times the systemic exposure observed in human males and 300 times the systemic exposure observed in human females as a result of use of the maximum recommended topical dose, based on AUC comparisons).

No evidence of potential to induce carcinogenicity was obtained in a dermal study in which dapsonone gel was topically applied to Tg.AC transgenic mice for approximately 26 weeks. Dapsonone concentrations of 3%, 5%, and 10% were evaluated; 3% material was judged to be the maximum tolerated dosage.

ACZONE® Gel, 5%, did not increase the rate of formation of ultra violet light-induced skin tumors when topically applied to hairless mice in a 12-month photocarcinogenicity study.

The effects of dapsonone on fertility and general reproduction performance were assessed in male and female rats following oral (gavage) dosing. Dapsonone reduced sperm motility at dosages of 3 mg/kg/day or greater (approximately 17 times the systemic exposure observed in human males as a result of use of the maximum recommended topical dose, based on AUC comparisons). The mean numbers of embryo implantations and viable embryos were significantly reduced in untreated females mated with males that had been dosed at 12 mg/kg/day or greater (approximately 70 times the systemic exposure observed in human males as a result of use of the maximum recommended topical dose, based on AUC comparisons), presumably due to reduced numbers or effectiveness of sperm, indicating impairment of fertility. Dapsonone had no effect on male fertility at dosages of 2 mg/kg/day or less (approximately 13 times the systemic exposure observed in human males as a result of use of the maximum recommended topical dose, based on AUC comparisons). When administered to female rats at a dosage of 75 mg/kg/day (approximately 800 times the systemic exposure observed in human females as a result of use of the maximum recommended topical dose, based on AUC comparisons) for 15 days prior to mating and for 17 days thereafter, dapsonone reduced the mean number of implantations, increased the mean early resorption rate, and reduced the mean litter size. These effects were probably secondary to maternal toxicity.

Dapsonone was assessed for effects on perinatal/postnatal pup development and postnatal maternal behavior and function in a study in which dapsonone was orally administered to female rats daily beginning on the seventh day of gestation and continuing until the twenty-seventh day postpartum. Maternal toxicity (decreased body weight and food consumption) and developmental effects (increase in stillborn pups and decreased pup weight) were seen at a dapsonone dose of 30 mg/kg/day (approximately 500 times the systemic exposure observed in human females as a result of use of the maximum recommended topical dose, based on AUC comparisons). No effects were observed on the viability, physical development, behavior, learning ability, or reproductive function of surviving pups.

## 14 CLINICAL STUDIES

Two randomized, double blind, vehicle controlled, clinical studies were conducted to evaluate ACZONE® Gel, 5%, for the treatment of patients with acne vulgaris (N=1475 and 1525). The studies were designed to enroll patients 12 years of age and older with 20 to 50 inflammatory and 20 to 100 non-inflammatory lesions at baseline. In these studies patients applied either ACZONE® Gel, 5%, or vehicle control twice daily for up to 12 weeks. Efficacy was evaluated in terms of success on the Global Acne Assessment Score (no or minimal acne) and in the percent reduction in inflammatory, non-inflammatory, and total lesions.

The Global Acne Assessment Score was a 5-point scale as follows:

- 0 None: no evidence of facial acne vulgaris
- 1 Minimal: few non-inflammatory lesions (comedones) are present; a few inflammatory lesions (papules/pustules) may be present
- 2 Mild: several to many non-inflammatory lesions (comedones) are present; a few inflammatory lesions (papules/pustules) are present
- 3 Moderate: many non-inflammatory (comedones) and inflammatory lesions (papules/pustules) are present; no nodulo-cystic lesions are allowed
- 4 Severe: significant degree of inflammatory disease; papules/pustules are a predominant feature; a few nodulo-cystic lesions may be present; comedones may be present.

The success rates on the Global Acne Assessment Score (no or minimal acne) at Week 12 are presented in Table 4.

Table 4 - Success (No or Minimal Acne) on the Global Acne Assessment Score at Week 12

	Study 1*		Study 2*	
	ACZONE® N=699	Vehicle N=687	ACZONE® N=729	Vehicle N=738
Subjects with No or Minimal Acne	291 (42%)	223 (32%)	253 (35%)	206 (28%)

\*Analysis excludes subjects classified with minimal acne at baseline

Table 5 presents the mean percent reduction in inflammatory, non-inflammatory, and total lesions from baseline to Week 12.

Table 5 - Percent Reduction in Lesions from Baseline to Week 12

	Study 1		Study 2	
	ACZONE® N=745	Vehicle N=740	ACZONE® N=761	Vehicle N=764
Inflammatory	46%	42%	48%	40%
Non-inflammatory	31%	24%	30%	21%
Total	38%	32%	37%	29%

The clinical studies enrolled about equal proportions of male and female subjects. Female patients tended to have greater percent reductions in lesions and greater success on the Global Acne Assessment Score than males. The breakdown by race in the clinical studies was about 73% Caucasian, 14% Black, 9% Hispanic, and 2% Asian. Efficacy results were similar across the racial subgroups.

## 16 HOW SUPPLIED/STORAGE AND HANDLING

ACZONE® (dapsonone) Gel, 5%, is supplied in the following size tubes:

Professional Sample  
5% NDC 0023-3670-03  
3 gram laminate tube

Commercially Available as:  
5% NDC 0023-3670-30  
30 gram laminate tube

**KEEP OUT OF THE REACH OF CHILDREN LESS THAN 12 YEARS OLD.**

Storage conditions:

Store at controlled room temperature, 20 - 25° C (68 - 76° F), excursions permitted to 15 - 30° C (59 - 86° F). Protect from freezing.

## 17 PATIENT COUNSELING INFORMATION

See FDA Approved-Patient Labeling (17.2)

### 17.1 Information for Patients

1. Patients should use ACZONE® Gel, 5%, as directed by the physician. ACZONE® Gel, 5%, is for external topical use only. ACZONE® Gel, 5%, is not for oral, ophthalmic or intravaginal use.
2. Patients should not use this medication for any disorder other than that for which it was prescribed.
3. Patients should report any signs of adverse reactions to their physician.
4. Protect ACZONE® Gel, 5%, from freezing.
5. See Patient Labeling for additional information on safety, efficacy, general use, and storage of ACZONE® Gel, 5%.

### 17.2 FDA-Approved Patient Labeling

ACZONE® (dapsonone) Gel 5%

Read this important information before you start using ACZONE® (AK-zon) Gel and each time you refill your prescription. There may be new information that you need to know. This summary is not meant to take the place of your doctor's advice. If you have any questions or want more information about ACZONE® Gel, ask your doctor or pharmacist.

#### What is ACZONE® Gel?

ACZONE® Gel is a prescription medicine used on your skin (topical) to treat acne in people 12 years and older.

ACZONE® Gel has not been studied in children under 12 years of age.

#### Who should not use ACZONE® Gel?

Do not use ACZONE® Gel if you are allergic to any of the ingredients in ACZONE® Gel or if you are younger than 12 years of age.

**Active ingredient:** dapsonone.

**Inactive ingredients:** Carbomer 980, diethylene glycol monoethyl ether (DGME), methylparaben, sodium hydroxide, and purified water.

#### What should I tell my doctor before using ACZONE® Gel?

**Tell your doctor about all of your medical conditions, including if you:**

- are pregnant or planning to become pregnant. It is not known if ACZONE® Gel may harm your unborn baby. You and your doctor will need to decide if ACZONE® is right for you.
- are breastfeeding. ACZONE® Gel passes into your milk and may harm your baby. You should choose either to use ACZONE® Gel, or breastfeed, but not both. Talk to your doctor about the best way to feed your baby while using ACZONE® Gel.
- have glucose-6-phosphate dehydrogenase deficiency.

**Tell your doctor about all the medicines you are taking including prescription and nonprescription medicines, vitamins and herbal supplements.** Especially, tell your doctor if you are using any other medicines applied to the skin, such as acne medicines with benzoyl peroxide.

#### How do I use ACZONE® Gel?

- Use ACZONE® Gel exactly as prescribed by your doctor. ACZONE® Gel is usually used on your affected skin twice a day, once in the morning and once in the evening.
- Wash the areas of your skin where you will apply ACZONE® Gel. Gently pat your skin dry with a clean towel.
- Apply a thin layer of ACZONE® Gel to the areas of your skin that have acne. A pea-sized amount of ACZONE® Gel will usually be enough.
- Rub the medicine in gently and completely
- Make sure to put the cap back on the ACZONE® Gel tube. Close it tightly.
- Wash your hands after applying ACZONE® Gel.
- Keep ACZONE® Gel away from your mouth and eyes. Do not swallow ACZONE® Gel. If you swallow ACZONE® Gel, call your doctor or poison control center right away.
- If your acne does not get better after using ACZONE® Gel for 12 weeks, talk to your doctor about other treatments for acne.

#### What are the possible side effects of ACZONE® Gel?

Like all medicines, ACZONE® Gel can cause some side effects. The most common side effects of ACZONE® Gel are dryness, redness, oiliness and peeling of the skin being treated.

When the active ingredient of ACZONE® Gel (called dapsonone) is taken orally as a pill, it has been related to the abnormal breakdown of red blood cells (hemolytic anemia). If you have glucose-6-phosphate dehydrogenase deficiency, you may have a greater risk for lowering your hemoglobin level. However, using ACZONE® Gel on the skin is not expected to put enough dapsonone in the blood to cause clinical symptoms of hemolytic anemia. You are advised to be alert for signs and symptoms suggestive of this type of anemia (sudden onset of: back pain, breathlessness, tiredness/weakness with daily activities, dark-brown urine, high fever and yellow or pale skin). If you experience these signs and symptoms, stop use and call your doctor immediately.

Use of benzoyl peroxide together with ACZONE® Gel at the same time may cause your skin to temporarily turn yellow or orange at the site of application.

This is not a complete list of all the possible side effects. Call your doctor if you have any side effects that do not go away or bother you. If you have any questions, ask your doctor or pharmacist.

#### How should I store ACZONE® Gel?

Store ACZONE® Gel at room temperature 68 to 76°F. Do not freeze ACZONE® Gel.

Keep ACZONE® Gel out of the reach of children less than 12 years of age.

#### Where can I find more information about ACZONE® Gel?

If you have any questions or want more information about ACZONE® Gel, ask your doctor or pharmacist. Your doctor or pharmacist can also give you a copy of the ACZONE® Gel Package Insert written for health professionals. Ask them to explain anything you do not understand.

You may call 1-800-433-8871 or visit [www.allergan.com](http://www.allergan.com) to obtain more information about ACZONE® Gel.

## ACZONE® (dapsonone) Gel, 5%

Manufactured for: Allergan, Inc., Irvine, CA 92612, U.S.A.

Manufactured by: TOLMAR Inc., Fort Collins, CO 80526, U.S.A.

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