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(54) Titre : UNE COMPOSITION MICROBICIDE DE N-METHYL-1,2-BENZISOTHIAZOLIN-3-ONE

(54) Title: AN N-METHYL-1,2-BENZISOTHIAZOLIN-3-ONE MICROBICIDAL COMPOSITION

(57) Abrégé/Abstract:

Synergistic microbicidal compositions containing N-methyl-1,2-benzisothiazolin- 3-one.



ABSTRACT OF THE DISCLOSURE

Synergistic microbicidal compositions containing N-methyl-1,2-benzisothiazolin-3-one.

AN N-METHYL-1,2-BENZISOTHIAZOLIN-3-ONE MICROBICIDAL COMPOSITION

This invention relates to a synergistic combination of selected microbicides having greater activity than would be observed for the individual microbicides.

5 In some cases, commercial microbicides cannot provide effective control of microorganisms, even at high use concentrations, due to weak activity against certain types of microorganisms, e.g., those resistant to some microbicides, or due to aggressive environmental conditions. Combinations of different microbicides are sometimes used to provide overall control of microorganisms in
10 a particular end use environment. For example, U.S. Pat. App. Pub. No. 2007/0078118 discloses synergistic combinations of N-methyl-1,2-benzisothiazolin-3-one (MBIT) with other biocides. However, there is a need for additional combinations of microbicides having enhanced activity against various strains of microorganisms to provide effective control of the microorganisms.
15 Moreover, there is a need for combinations containing lower levels of individual microbicides for environmental and economic benefit. The problem addressed by this invention is to provide such additional combinations of microbicides.

STATEMENT OF THE INVENTION

20 The present invention is directed to a microbicidal composition comprising: (a) N-methyl-1,2-benzisothiazolin-3-one; and (b) at least one microbicide selected from the group consisting of ethyl lauroyl arginate hydrochloride, cocamidopropyl-N-2-hydroxyethylcarbamoyl methyl dimethyl ammonium chloride, Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium
25 chloride, didecyl dimethyl ammonium carbonate and bicarbonate, Cu(II) 2-aminoethanolate, glycerol monolaurate, propylene glycol mono-laurate, and propylene glycol caprylate.

DETAILED DESCRIPTION OF THE INVENTION

30 As used herein, the following terms have the designated definitions, unless the context clearly indicates otherwise. "MBIT" is N-methyl-1,2-benzisothiazolin-3-one. The term "microbicide" refers to a compound capable of

killing, inhibiting the growth of or controlling the growth of microorganisms at a locus; microbicides include bactericides, fungicides and algacides. The term "microorganism" includes, for example, fungi (such as yeast and mold), bacteria and algae. The term "locus" refers to an industrial system or product subject to contamination by microorganisms. The following abbreviations are used throughout the specification: ppm = parts per million by weight (weight/weight), mL = milliliter, ATCC = American Type Culture Collection, MBC = minimum biocidal concentration, and MIC = minimum inhibitory concentration. Unless otherwise specified, temperatures are in degrees centigrade (°C), and references to percentages (%) are by weight. Amounts of organic microbicides are given on an active ingredient basis in ppm (w/w).

The compositions of the present invention unexpectedly have been found to provide enhanced microbicidal efficacy at a combined active ingredient level lower than that of the individual microbicides. Additional microbicides beyond those listed in the claims may be present in the composition.

In one embodiment of the invention, the antimicrobial composition comprises N-methyl-1,2-benzisothiazolin-3-one and ethyl lauroyl arginate hydrochloride. Preferably, a weight ratio of ethyl lauroyl arginate hydrochloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:94 to 1:0.0137.

In one embodiment of the invention, the antimicrobial composition comprises N-methyl-1,2-benzisothiazolin-3-one and cocamidopropyl-N-2-hydroxyethylcarbamoyl methyl dimethyl ammonium chloride. Preferably, a weight ratio of cocamidopropyl-N-2-hydroxyethylcarbamoyl methyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:188 to 1:0.0022.

In one embodiment of the invention, the antimicrobial composition comprises N-methyl-1,2-benzisothiazolin-3-one and Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride. Preferably, a weight ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:176 to 1:0.08.

In one embodiment of the invention, the antimicrobial composition comprises N-methyl-1,2-benzisothiazolin-3-one and didecyl dimethyl ammonium

carbonate and bicarbonate. Preferably, a weight ratio of didecyl dimethyl ammonium carbonate and bicarbonate to N-methyl-1,2-benzisothiazolin-3-one is from 1:227 to 1:0.179.

In one embodiment of the invention, the antimicrobial composition
5 comprises N-methyl-1,2-benzisothiazolin-3-one and Cu(II) 2-aminoethanolate. Preferably, a weight ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:375 to 1:0.004.

In one embodiment of the invention, the antimicrobial composition
comprises N-methyl-1,2-benzisothiazolin-3-one and glycerol monolaurate.
10 Preferably, a weight ratio of glycerol monolaurate to N-methyl-1,2-benzisothiazolin-3-one is from 1:0.143 to 1:0.0004.

In one embodiment of the invention, the antimicrobial composition
comprises N-methyl-1,2-benzisothiazolin-3-one and propylene glycol mono-
laurate. Preferably, a weight ratio of propylene glycol mono-laurate to N-methyl-
15 1,2-benzisothiazolin-3-one is from 1:0.286 to 1:0.0286.

In one embodiment of the invention, the antimicrobial composition
comprises N-methyl-1,2-benzisothiazolin-3-one and propylene glycol caprylate.
Preferably, a weight ratio of propylene glycol caprylate to N-methyl-1,2-
benzisothiazolin-3-one is from 1:0.442 to 1:0.0018.

20 The microbicides in the composition of this invention may be used "as is" or may first be formulated with a solvent or a solid carrier. Suitable solvents include, for example, water; glycols, such as ethylene glycol, propylene glycol, diethylene glycol, dipropylene glycol, polyethylene glycol, and polypropylene glycol; glycol ethers; alcohols, such as methanol, ethanol, propanol, phenethyl
25 alcohol and phenoxypropanol; ketones, such as acetone and methyl ethyl ketone; esters, such as ethyl acetate, butyl acetate, triacetyl citrate, and glycerol triacetate; carbonates, such as propylene carbonate and dimethyl carbonate; and mixtures thereof. It is preferred that the solvent is selected from water, glycols, glycol ethers, esters and mixtures thereof. Suitable solid carriers include, for
30 example, cyclodextrin, silicas, diatomaceous earth, waxes, cellulosic materials, alkali and alkaline earth (e.g., sodium, magnesium, potassium) metal salts (e.g., chloride, nitrate, bromide, sulfate) and charcoal.

When a microbicide component is formulated in a solvent, the formulation may optionally contain surfactants. When such formulations contain surfactants, they are generally in the form of emulsive concentrates, emulsions, microemulsive concentrates, or microemulsions. Emulsive concentrates form emulsions upon the addition of a sufficient amount of water. Microemulsive concentrates form microemulsions upon the addition of a sufficient amount of water. Such emulsive and microemulsive concentrates are generally well known in the art; it is preferred that such formulations are free of surfactants. U.S. Patent No. 5,444,078 may be consulted for further general and specific details on the preparation of various microemulsions and microemulsive concentrates.

A microbicide component also can be formulated in the form of a dispersion. The solvent component of the dispersion can be an organic solvent or water, preferably water. Such dispersions can contain adjuvants, for example, co-solvents, thickeners, anti-freeze agents, dispersants, fillers, pigments, surfactants, biodispersants, sulfosuccinates, terpenes, furanones, polycations, stabilizers, scale inhibitors and anti-corrosion additives.

When both microbicides are each first formulated with a solvent, the solvent used for the first microbicide may be the same as or different from the solvent used to formulate the other commercial microbicide, although water is preferred for most industrial biocide applications. It is preferred that the two solvents are miscible.

Those skilled in the art will recognize that the microbicide components of the present invention may be added to a locus sequentially, simultaneously, or may be combined before being added to the locus. It is preferred that the first microbicide and the second microbicide component be added to a locus simultaneously or sequentially. When the microbicides are added simultaneously or sequentially, each individual component may contain adjuvants, such as, for example, solvent, thickeners, anti-freeze agents, colorants, sequestrants (such as ethylenediamine-tetraacetic acid, ethylenediaminedisuccinic acid, iminodisuccinic acid and salts thereof), dispersants, surfactants, biodispersants, sulfosuccinates, terpenes, furanones, polycations, stabilizers, scale inhibitors and anti-corrosion additives.

The microbicidal compositions of the present invention can be used to inhibit the growth of microorganisms or higher forms of aquatic life (such as protozoans, invertebrates, bryozoans, dinoflagellates, crustaceans, mollusks, etc.) by introducing a microbically effective amount of the compositions onto, into, or at a locus subject to microbial attack. Suitable loci include, for example:

industrial process water; electrocoat deposition systems; cooling towers; air washers; gas scrubbers; mineral slurries; wastewater treatment; ornamental fountains; reverse osmosis filtration; ultrafiltration; ballast water; evaporative condensers; heat exchangers; pulp and paper processing fluids and additives; starch; plastics; emulsions; dispersions; paints; latices; coatings, such as varnishes; construction products, such as mastics, caulks, and sealants; construction adhesives, such as ceramic adhesives, carpet backing adhesives, and laminating adhesives; industrial or consumer adhesives; photographic chemicals; printing fluids; household products, such as bathroom and kitchen cleaners and sanitary wipes; cosmetics; toiletries; shampoos; soaps; detergents; industrial cleaners; floor polishes; laundry rinse water; metalworking fluids; conveyor lubricants; hydraulic fluids; leather and leather products; textiles; textile products; wood and wood products, such as plywood, chipboard, wallboard, flakeboard, laminated beams, oriented strandboard, hardboard, and particleboard; petroleum processing fluids; fuel; oilfield fluids, such as injection water, fracture fluids, and drilling muds; agriculture adjuvant preservation; surfactant preservation; medical devices; diagnostic reagent preservation; food preservation, such as plastic or paper food wrap; food, beverage, and industrial process pasteurizers; toilet bowls; recreational water; pools; and spas.

Preferably, the microbicidal compositions of the present invention are used to inhibit the growth of microorganisms at a locus selected from one or more of mineral slurries, pulp and paper processing fluids and additives, starch, emulsions, dispersions, paints, latices, coatings, construction adhesives, such as ceramic adhesives, carpet backing adhesives, photographic chemicals, printing fluids, household products such as bathroom and kitchen cleaners and sanitary wipes, cosmetics, toiletries, shampoos, soaps, detergents, industrial cleaners, floor polishes, laundry rinse water, metal working fluids, textile products, wood

and wood products, agriculture adjuvant preservation, surfactant preservation, diagnostic reagent preservation, food preservation, and food, beverage, and industrial process pasteurizers.

The specific amount of the composition of this invention necessary to
5 inhibit or control the growth of microorganisms and higher aquatic life forms in a locus depends upon the particular locus to be protected. Typically, the amount of the composition of the present invention to control the growth of microorganisms in a locus is sufficient if it provides from 0.1 to 1,000 ppm of the isothiazoline ingredient of the composition in the locus. It is preferred that the isothiazolone
10 ingredients of the composition be present in the locus in an amount of at least 0.5 ppm, more preferably at least 4 ppm and most preferably at least 10 ppm. It is preferred that the isothiazolone ingredients of the composition be present in the locus in an amount of no more than 1000 ppm, more preferably no more than 500 ppm, and most preferably no more than 200 ppm.

EXAMPLES

Materials and Methods

5 The synergism of the combination of the present invention was demonstrated by testing a wide range of concentrations and ratios of the compounds.

 One measure of synergism is the industrially accepted method described by Kull, F.C.; Eisman, P.C.; Sylwestrowicz, H.D. and Mayer, R.L., in Applied
10 Microbiology 9:538-541 (1961), using the ratio determined by the formula:

$$Q_a/Q_A + Q_b/Q_B = \text{Synergy Index ("SI")}$$

wherein:

Q_A = concentration of compound A (first component) in ppm, acting alone,
15 which produced an end point (MIC of Compound A).

Q_a = concentration of compound A in ppm, in the mixture, which produced an end point.

Q_B = concentration of compound B (second component) in ppm, acting alone, which produced an end point (MIC of Compound B).

20 Q_b = concentration of compound B in ppm, in the mixture, which produced an end point.

 When the sum of Q_a/Q_A and Q_b/Q_B is greater than one, antagonism is indicated. When the sum is equal to one, additivity is indicated, and when less
25 than one, synergism is demonstrated. The lower the SI, the greater the synergy shown by that particular mixture. The minimum inhibitory concentration (MIC) of a microbicide is the lowest concentration tested under a specific set of conditions that prevents the growth of added microorganisms.

 Synergy tests were conducted using standard microtiter plate assays with
30 media designed for optimal growth of the test microorganism. Minimal salt medium supplemented with 0.2% glucose and 0.1% yeast extract (M9GY medium) was used for testing bacteria; Potato Dextrose Broth (PDB medium) was used for testing yeast and mold. In this method, a wide range of

combinations of microbicides was tested by conducting high resolution MIC assays in the presence of various concentrations of MBIT. High resolution MICs were determined by adding varying amounts of microbicide to one column of a microtitre plate and doing subsequent ten-fold dilutions using an automated liquid handling system to obtain a series of endpoints ranging from 2 ppm to 10,000 ppm active ingredient.

The synergy of the combinations of the present invention was determined against several microorganisms, as described in the Tables below. The bacteria were used at a concentration of about 5×10^6 bacteria per mL and the yeast and mold at 5×10^5 fungi per mL. These microorganisms are representative of natural contaminants in many consumer and industrial applications. The plates were visually evaluated for microbial growth (turbidity) to determine the MIC after various incubation times at 25° C (yeast and mold) or 30° C (bacteria).

The test results for demonstration of synergy of the MBIT combinations of the present invention are shown below in Tables 1 through 8. In each test, Second Component (B) was MBIT and the First Component (A) was the other commercial microbicide. Each table shows the specific combinations of MBIT and the other component; results against the microorganisms tested with incubation times; the end-point activity in ppm measured by the MIC for MBIT alone (Q_B), for the other component alone (Q_A), for MBIT in the mixture (Q_b) and for the other component in the mixture (Q_a); the calculated SI value; and the range of synergistic ratios for each combination tested (other component/MBIT or A/B).

Table1

Ca: ppm AI of CYTO GUARD LA (ethyl lauroyl arginate hydrochloride)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

Ratio:Ca:Cb

5

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	110	-	-	-
		-	37.5	-	-
		0.2	18.8	0.50	1:94.0000
		2	18.8	0.52	1:9.4000
		11	18.8	0.60	1:1.7091
		20	18.8	0.68	1:0.9400
		30	18.8	0.77	1:0.6267
		42.5	18.8	0.89	1:0.4424
		52.5	18.8	0.98	1:0.3581
		65	18.8	1.09	1:0.2892
		11	9.4	0.35	1:0.8545
		20	9.4	0.43	1:0.4700
		30	9.4	0.52	1:0.3133
		42.5	9.4	0.64	1:0.2212
		52.5	9.4	0.73	1:0.1790
		65	9.4	0.84	1:0.1446
		87.5	9.4	1.05	1:0.1074
		42.5	4.7	0.51	1:0.1106
		52.5	4.7	0.60	1:0.0895
		65	4.7	0.72	1:0.0723
		87.5	4.7	0.92	1:0.0537
		110	4.7	1.13	1:0.0427
		42.5	2.4	0.45	1:0.0565
		52.5	2.4	0.54	1:0.0457
		65	2.4	0.65	1:0.0369
		87.5	2.4	0.86	1:0.0274
		110	2.4	1.06	1:0.0218
		42.5	1.2	0.42	1:0.0282
		65	1.2	0.62	1:0.0185
		87.5	1.2	0.83	1:0.0137
		110	1.2	1.03	1:0.0109
	7 days	110	-	-	-
		-	37.5	-	-
		2	18.8	0.52	1:9.4000
		11	18.8	0.6	1:1.7091
		20	18.8	0.68	1:0.9400
		30	18.8	0.77	1:0.6267
		42.5	18.8	0.89	1:0.4424
		52.5	18.8	0.98	1:0.3581
		65	18.8	1.09	1:0.2892
		20	9.4	0.43	1:0.4700
		30	9.4	0.52	1:0.3133
		42.5	9.4	0.64	1:0.2212
		52.5	9.4	0.73	1:0.1790
		65	9.4	0.84	1:0.1446

10

		87.5	9.4	1.05	1:0.1074
		42.5	4.7	0.51	1:0.1106
		52.5	4.7	0.6	1:0.0895
		65	4.7	0.72	1:0.0723
		87.5	4.7	0.92	1:0.0537
		110	4.7	1.13	1:0.0427
		42.5	2.4	0.45	1:0.0565
		52.5	2.4	0.54	1:0.0457
		65	2.4	0.65	1:0.0369
		87.5	2.4	0.86	1:0.0274
		110	2.4	1.06	1:0.0218
		65	1.2	0.62	1:0.0185
		87.5	1.2	0.83	1:0.0137
		110	1.2	1.03	1:0.0109
<i>C.albicans</i> ATCC # 10231	48 hrs	87.5	-	-	-
		-	30	-	-
		6.5	15	0.57	1:2.3077
		11	15	0.63	1:1.3636
		20	15	0.73	1:0.7500
		30	15	0.84	1:0.5000
		42.5	15	0.99	1:0.3529
		6.5	7.5	0.32	1:1.1538
		11	7.5	0.38	1:0.6818
		20	7.5	0.48	1:0.3750
		30	7.5	0.59	1:0.2500
		42.5	7.5	0.74	1:0.1765
		52.5	7.5	0.85	1:0.1429
		65	7.5	0.99	1:0.1154
		8.75	3.75	0.23	1:0.4286
		11	3.75	0.25	1:0.3409
		20	3.75	0.35	1:0.1875
		30	3.75	0.47	1:0.1250
	72 hrs	42.5	3.75	0.61	1:0.0882
		52.5	3.75	0.73	1:0.0714
		65	3.75	0.87	1:0.0577
		87.5	3.75	1.13	1:0.0429
		20	1.86	0.29	1:0.0930
		30	1.86	0.40	1:0.0620
		42.5	1.86	0.55	1:0.0438
		52.5	1.86	0.66	1:0.0354
		65	1.86	0.80	1:0.0286
		87.5	1.86	1.06	1:0.0213
		87.5	0.94	1.03	1:0.0107
		87.5	-	-	-
		-	30	-	-
		8.75	15	0.60	1:1.7143
		11	15	0.63	1:1.3636
		20	15	0.73	1:0.7500
		30	15	0.84	1:0.5000
		42.5	15	0.99	1:0.3529
		8.75	7.5	0.35	1:0.8571
		11	7.5	0.38	1:0.6818

		20	7.5	0.48	1:0.3750
		30	7.5	0.59	1:0.2500
		42.5	7.5	0.74	1:0.1765
		52.5	7.5	0.85	1:0.1429
		65	7.5	0.99	1:0.1154
		11	3.75	0.25	1:0.3409
		20	3.75	0.35	1:0.1875
		30	3.75	0.47	1:0.1250
		42.5	3.75	0.61	1:0.0882
		52.5	3.75	0.73	1:0.0714
		65	3.75	0.87	1:0.0577
		87.5	3.75	1.13	1:0.0429
		42.5	1.86	0.55	1:0.0438
		52.5	1.86	0.66	1:0.0354
		65	1.86	0.8	1:0.0286
		87.5	1.86	1.06	1:0.0213
<i>Ps. aeruginosa</i> ATCC#9027	48hrs	42.5	-	-	-
		-	125	-	-
		30	18.75	0.86	1:0.6250
		42.5	18.75	1.15	1:0.4412
		30	9.4	0.78	1:0.3133
		42.5	9.4	1.08	1:0.2212
		30	4.6	0.74	1:0.1533
		42.5	4.6	1.04	1:0.1082
<i>S. aureus</i> ATCC#6538	24hrs	42.5	2.3	1.02	1:0.0541
		2.125	-	-	-
		-	15	-	-
		1	7.5	0.97	1:7.5000
		1.5	7.5	1.21	1:5.0000
		1	3.8	0.72	1:3.8000
		1.5	3.8	0.96	1:2.5333
		1.5	1.8	0.83	1:1.2000
	48hrs	2.125	1.8	1.12	1:0.8471
		1.5	0.94	0.77	1:0.6267
		2.125	0.94	1.06	1:0.4424
		2.625	-	-	-
		-	15	-	-
		1.5	7.5	1.07	1:5.0000
		1.5	3.8	0.82	1:2.5333
		2.125	3.8	1.06	1:1.7882
		2.125	1.8	0.93	1:0.8471
		2.625	1.8	1.12	1:0.6857
		2.125	0.94	0.87	1:0.4424
		2.625	0.94	0.94	1:0.3581

Table:2

Ca: ppm AI of MONTALINE C40 (cocamidopropyl-N-2-hydroxyethylcarbamoyl methyl dimethyl ammonium chloride)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

5 Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	650	-	-	-
		-	37.5	-	-
		20	18.8	0.53	1:0.9400
		30	18.8	0.55	1:0.6267
		42.5	18.8	0.57	1:0.4424
		87.5	18.8	0.64	1:0.2149
		110	18.8	0.67	1:0.1709
		200	18.8	0.81	1:0.0940
		300	18.8	0.96	1:0.0627
		425	18.8	1.16	1:0.0442
		87.5	9.4	0.39	1:0.1074
		110	9.4	0.42	1:0.0855
		200	9.4	0.56	1:0.0470
		300	9.4	0.71	1:0.0313
		425	9.4	0.90	1:0.0221
		525	9.4	1.06	1:0.0179
		425	4.7	0.78	1:0.0111
		525	4.7	0.93	1:0.0090
		650	4.7	1.13	1:0.0072
		425	2.4	0.72	1:0.0056
		525	2.4	0.87	1:0.0046
		650	2.4	1.06	1:0.0037
	7 days	650	-	-	-
		-	75	-	-
		0.2	37.5	0.50	1:187.5000
		2	37.5	0.50	1:18.7500
		20	37.5	0.53	1:1.8750
		42.5	37.5	0.57	1:0.8824
		65	37.5	0.60	1:0.5769
		87.5	37.5	0.63	1:0.4286
		110	37.5	0.67	1:0.3409
		200	37.5	0.81	1:0.1875
		300	37.5	0.96	1:0.1250
		425	37.5	1.15	1:0.0882
		110	18.8	0.42	1:0.1709
		200	18.8	0.56	1:0.0940
		300	18.8	0.71	1:0.0627
		425	18.8	0.90	1:0.0442
		525	18.8	1.06	1:0.0358
		200	9.4	0.43	1:0.0470
		300	9.4	0.59	1:0.0313
		425	9.4	0.78	1:0.0221
		525	9.4	0.93	1:0.0179
		650	9.4	1.13	1:0.0145

		425	4.7	0.72	1:0.0111
		525	4.7	0.87	1:0.0090
		650	4.7	1.06	1:0.0072
<i>C.albicans</i> ATCC # 10231	48 hrs	525	-	-	-
		-	30	-	-
		300	15	1.07	1:0.0500
		425	7.5	1.06	1:0.0176
		425	3.75	0.93	1:0.0088
		425	1.86	0.87	1:0.0044
		425	0.94	0.84	1:0.0022
	72 hrs	525	0.94	1.03	1:0.0018
		1100	-	-	-
		-	30	-	-
		87.5	30	0.67	1:0.3429
		110	30	0.71	1:0.2727
		200	30	0.88	1:0.1500
		300	30	1.07	1:0.1000
		300	15	0.82	1:0.0500
		425	15	1.06	1:0.0353
		425	7.5	0.93	1:0.0176
		525	7.5	1.13	1:0.0143
		425	3.75	0.87	1:0.0088
		525	3.75	1.06	1:0.0071
		425	1.86	0.84	1:0.0044
		525	1.86	1.03	1:0.0035
		525	0.94	1.02	1:0.0018
<i>Ps. aeruginosa</i> ATCC#9027	24hrs	300	-	-	-
		-	125	-	-
		87.5	75	0.89	1:0.8571
		110	75	0.97	1:0.6818
		200	37.5	0.97	1:0.1875
		300	37.5	1.3	1:0.1250
		200	18.75	0.82	1:0.0938
		300	18.75	1.15	1:0.0625
		200	9.4	0.74	1:0.0470
		300	9.4	1.08	1:0.0313
		300	4.6	1.04	1:0.0153
		300	2.3	1.02	1:0.0077
	48hrs	300	-	-	-
		-	125	-	-
		87.5	75	0.89	1:0.8571
		110	75	0.97	1:0.6818
		200	37.5	0.97	1:0.1875
		300	37.5	1.3	1:0.1250
		200	18.75	0.82	1:0.0938
		300	18.75	1.15	1:0.0625
		200	9.4	0.74	1:0.0470
		300	9.4	1.08	1:0.0313
<i>S.aureus</i> ATCC#6538	24hrs	5	-	-	-
		-	15	-	-

		2.188	7.5	0.94	1:3.4278
		2.75	3.8	0.80	1:1.3818
		5	3.8	1.25	1:0.7600
		5	1.8	1.12	1:0.3600
		5	0.94	1.06	1:0.1880
	48hrs	5	-	-	-
		-	15	-	-
		2.188	7.5	0.94	1:3.4278
		2.75	3.8	0.80	1:1.3818
		5	3.8	1.25	1:0.7600
		5	1.8	1.12	1:0.3600
		5	0.94	1.06	1:0.1880

Table:3

TM

Ca: ppm AI of ACQ type D (Cu(II) 2-aminoethanolate & didecyldimethyl ammonium chloride) (ppm CuO)

5 Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	42.5	-	-	-
		-	37.5	-	-
		2	18.8	0.55	1:9.4000
		5.25	18.8	0.62	1:3.5810
		6.5	18.8	0.65	1:2.8923
		11	18.8	0.76	1:1.7091
		20	18.8	0.97	1:0.9400
		20	9.4	0.72	1:0.4700
		30	9.4	0.96	1:0.3133
		20	4.7	0.6	1:0.2350
		30	4.7	0.83	1:0.1567
		42.5	4.7	1.13	1:0.1106
		20	2.4	0.53	1:0.1200
		30	2.4	0.77	1:0.0800
	7 days	42.5	2.4	1.06	1:0.0565
		42.5	-	-	-
		-	75	-	-
		2	37.5	0.55	1:18.7500
		4.25	37.5	0.60	1:8.8235
		6.5	37.5	0.65	1:5.7692
		11	37.5	0.76	1:3.4091
		20	37.5	0.97	1:1.8750
		6.5	18.8	0.4	1:2.8923
		11	18.8	0.51	1:1.7091
		20	18.8	0.72	1:0.9400
		30	18.8	0.96	1:0.6267
<i>C. albicans</i> ATCC # 10231	48 hrs	42.5	4.7	1.06	1:0.1106
		42.5	1.2	1.02	1:0.0282

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	72 hrs	0.525	15	0.56	1:28.5714
		1.1	15	0.63	1:13.6364
		2	15	0.73	1:7.5000
		4.25	15	0.99	1:3.5294
		4.25	7.5	0.74	1:1.7647
		6.5	7.5	0.99	1:1.1538
		5.25	3.75	0.73	1:0.7143
		6.5	3.75	0.87	1:0.5769
		8.75	3.75	1.13	1:0.4286
		5.25	1.86	0.66	1:0.3543
		6.5	1.86	0.80	1:0.2862
		8.75	1.86	1.06	1:0.2126
		6.5	0.94	0.77	1:0.1446
		8.75	0.94	1.03	1:0.1074
		11	-	-	-
		-	60	-	-
		0.3	30	0.53	1:100.0000
		1.1	30	0.6	1:27.2727
		2	30	0.68	1:15.0000
		4.25	30	0.89	1:7.0588
		5.25	30	0.98	1:5.7143
		6.5	30	1.09	1:4.6154
		2	15	0.43	1:7.5000
		2	15	0.43	1:7.5000
		4.25	15	0.64	1:3.5294
		5.25	15	0.73	1:2.8571
		6.5	15	0.84	1:2.3077
		8.75	15	1.05	1:1.7143
		4.25	7.5	0.51	1:1.7647
		5.25	7.5	0.6	1:1.4286
		6.5	7.5	0.72	1:1.1538
		8.75	7.5	0.92	1:0.8571
		11	7.5	1.13	1:0.6818
		5.25	3.75	0.54	1:0.7143
		6.5	3.75	0.65	1:0.5769
		8.75	3.75	0.86	1:0.4286
		11	3.75	1.06	1:0.3409
		6.5	1.86	0.62	1:0.2862
		8.75	1.86	0.83	1:0.2126
		11	1.86	1.03	1:0.1691
		6.5	0.94	0.61	1:0.1446
		8.75	0.94	0.81	1:0.1074
		11	0.94	1.02	1:0.0855
<i>Ps. aeruginosa</i> ATCC#9027	24hrs	20	-	-	-
		-	75	-	-
		2	37.5	0.60	1:18.7500
		3	37.5	0.65	1:12.5000
		4.25	37.5	0.71	1:8.8235
		5.25	37.5	0.76	1:7.1429
		6.5	37.5	0.83	1:5.7692
		8.75	37.5	0.94	1:4.2857
		11	37.5	1.05	1:3.4091

	48 hrs	5.25	18.75	0.51	1:3.5714
		6.5	18.75	0.58	1:2.8846
		8.75	18.75	0.69	1:2.1429
		11	18.75	0.80	1:1.7045
		20	18.75	1.25	1:0.9375
		8.75	9.4	0.56	1:1.0743
		11	9.4	0.68	1:0.8545
		20	9.4	1.13	1:0.4700
		11	4.6	0.61	1:0.4182
		20	4.6	1.06	1:0.2300
		20	2.3	1.03	1:0.1150
		30	-	-	-
		-	125	-	-
		0.425	75	0.61	1:176.4706
		0.525	75	0.62	1:142.8571
		1.1	75	0.64	1:68.1818
		2	75	0.67	1:37.5000
		4.25	75	0.74	1:17.6471
		6.5	75	0.82	1:11.5385
		11	75	0.97	1:6.8182
		3	37.5	0.40	1:12.5000
		4.25	37.5	0.44	1:8.8235
		6.5	37.5	0.52	1:5.7692
		11	37.5	0.67	1:3.4091
		20	37.5	0.97	1:1.8750
		8.75	18.75	0.44	1:2.1429
		11	18.75	0.52	1:1.7045
		20	18.75	0.82	1:0.9375
		30	18.75	1.15	1:0.6250
		20	9.4	0.74	1:0.4700
<i>S.aureus</i> ATCC#6538	24hrs	1.1	-	-	-
		-	15	-	-
		0.425	7.5	0.89	1:17.6471
		0.525	7.5	0.98	1:14.2857
		0.875	3.8	1.05	1:4.3429
		0.875	1.8	0.92	1:2.0571
	48hrs	0.875	0.94	0.86	1:1.0743
		1.1	0.94	1.06	1:0.8545
		1.1	-	-	-
		-	15	-	-
		0.525	7.5	0.98	1:14.2857
		0.875	3.8	1.05	1:4.3429
		1.1	0.94	1.06	1:0.8545

Table:4

Ca: ppm AI of CARBOQUAT WP-50(didecyl dimethyl ammonium carbonate and bicarbonate)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

5 Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	15	-	-	-
		-	37.5	-	-
		1	18.8	0.57	1:18.8000
		2.125	18.8	0.64	1:8.8471
		3.25	18.8	0.72	1:5.7846
		4.375	18.8	0.79	1:4.2971
		5.5	18.8	0.87	1:3.4182
		10	18.8	1.17	1:1.8800
		4.375	9.4	0.54	1:2.1486
		5.5	9.4	0.62	1:1.7091
		10	9.4	0.92	1:0.9400
	7 days	32.5	-	-	-
		-	37.5	-	-
		1	18.8	0.53	1:18.8000
		2.125	18.8	0.57	1:8.8471
		3.25	18.8	0.60	1:5.7846
		4.375	18.8	0.64	1:4.2971
		5.5	18.8	0.67	1:3.4182
		10	18.8	0.81	1:1.8800
		15	18.8	0.96	1:1.2533
		26.25	9.4	1.06	1:0.3581
		21.25	4.7	0.78	1:0.2212
		26.25	4.7	0.93	1:0.1790
		32.5	4.7	1.13	1:0.1446
		32.5	2.4	1.06	1:0.0738
<i>C.albicans</i> ATCC # 10231	48 hrs	3.25	-	-	-
		-	30	-	-
		2.125	15	1.15	1:7.0588
		2.125	7.5	0.90	1:3.5294
		3.25	7.5	1.25	1:2.3077
		2.125	3.75	0.78	1:1.7647
		3.25	3.75	1.13	1:1.1538
		2.125	1.86	0.72	1:0.8753
		3.25	1.86	1.06	1:0.5723
		3.25	0.94	1.03	1:0.2892
	72 hrs	4.375	-	-	-
		-	60	-	-
		0.2625	30	0.56	1:114.2857
		0.55	30	0.63	1:54.5455
		1	30	0.73	1:30.0000
		1.5	30	0.84	1:20.0000
		2.125	30	0.99	1:14.1176
		2.125	15	0.74	1:7.0588

		3.25	15	0.99	1:4.6154
		2.625	7.5	0.73	1:2.8571
		3.25	7.5	0.87	1:2.3077
		4.375	7.5	1.13	1:1.7143
		2.625	3.75	0.66	1:1.4286
		3.25	3.75	0.81	1:1.1538
		4.375	3.75	1.06	1:0.8571
		2.625	1.86	0.63	1:0.7086
		3.25	1.86	0.77	1:0.5723
		4.375	1.86	1.03	1:0.4251
		3.25	0.94	0.76	1:0.2892
		4.375	0.94	1.02	1:0.2149
<i>Ps. aeruginosa</i> ATCC#9027	24hrs	10	-	-	-
		-	125	-	-
		0.4375	75	0.64	1:171.4286
		1	75	0.70	1:75.0000
		1.5	75	0.75	1:50.0000
		2.625	75	0.86	1:28.5714
		3.25	75	0.93	1:23.0769
		4.375	75	1.04	1:17.1429
	48 hrs	10	4.6	1.04	1:0.4600
		10	2.3	1.02	1:0.2300
		10	-	-	-
		-	125	-	-
		2.125	75	0.81	1:35.2941
		2.625	75	0.86	1:28.5714
		3.25	75	0.93	1:23.0769
		4.375	75	1.04	1:17.1429
		10	4.6	1.04	1:0.4600
		10	2.3	1.02	1:0.2300
<i>S. aureus</i> ATCC#6538	24hrs	0.2125	-	-	-
		-	15	-	-
		0.033	7.5	0.66	1:227.2727
		0.055	7.5	0.76	1:136.3636
		0.1	7.5	0.97	1:75.0000
		0.15	3.8	0.96	1:25.3333
		0.2125	0.94	1.06	1:4.4235
		0.2125	-	-	-
		-	15	-	-
		0.2125	0.94	1.06	1:4.4235

Table:5

TM

Ca: ppm AI of ACQ C2 (Cu(II) 2-aminoethanolate) (ppm CuO)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

Ratio:Ca:Cb

5

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	425	-	-	-
		-	37.5	-	-
		11	18.8	0.53	1:1.7091
		20	18.8	0.55	1:0.9400
		30	18.8	0.57	1:0.6267
		42.5	18.8	0.60	1:0.4424
		65	18.8	0.65	1:0.2892
		87.5	18.8	0.71	1:0.2149
		110	18.8	0.76	1:0.1709
		200	18.8	0.97	1:0.0940
		30	9.4	0.32	1:0.3133
		42.5	9.4	0.35	1:0.2212
		65	9.4	0.40	1:0.1446
		87.5	9.4	0.46	1:0.1074
		110	9.4	0.51	1:0.0855
		200	9.4	0.72	1:0.0470
		300	9.4	0.96	1:0.0313
		87.5	4.7	0.33	1:0.0537
		110	4.7	0.38	1:0.0427
		200	4.7	0.60	1:0.0235
		300	4.7	0.83	1:0.0157
		425	4.7	1.13	1:0.0111
		300	1.2	0.74	1:0.0040
		425	1.2	1.03	1:0.0028
	7 days	650	-	-	-
		-	37.5	-	-
		42.5	18.8	0.57	1:0.4424
		65	18.8	0.60	1:0.2892
		87.5	18.8	0.64	1:0.2149
		110	18.8	0.67	1:0.1709
		200	18.8	0.81	1:0.0940
		300	18.8	0.96	1:0.0627
		425	18.8	1.16	1:0.0442
		42.5	9.4	0.32	1:0.2212
		65	9.4	0.35	1:0.1446
		87.5	9.4	0.39	1:0.1074
		110	9.4	0.42	1:0.0855
		200	9.4	0.56	1:0.0470
		300	9.4	0.71	1:0.0313
		425	9.4	0.90	1:0.0221
		87.5	4.7	0.26	1:0.0537
		110	4.7	0.29	1:0.0427
		200	4.7	0.43	1:0.0235
		300	4.7	0.59	1:0.0157
		425	4.7	0.78	1:0.0111

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		525	4.7	0.93	1:0.0090
		650	4.7	1.13	1:0.0072
		200	2.4	0.37	1:0.0120
		300	2.4	0.53	1:0.0080
		425	2.4	0.72	1:0.0056
		525	2.4	0.87	1:0.0046
		650	2.4	1.06	1:0.0037
<i>C.albicans</i> ATCC # 10231	48 hrs	300	-	-	-
		-	60	-	-
		3	30	0.51	1:10.0000
		11	30	0.54	1:2.7273
		20	30	0.57	1:1.5000
		42.5	30	0.64	1:0.7059
		87.5	30	0.79	1:0.3429
		110	30	0.87	1:0.2727
		200	30	1.17	1:0.1500
		0.525	15	0.25	1:28.5714
		3	15	0.26	1:5.0000
		11	15	0.29	1:1.3636
		20	15	0.32	1:0.7500
		42.5	15	0.39	1:0.3529
		87.5	15	0.54	1:0.1714
		110	15	0.62	1:0.1364
		200	15	0.92	1:0.0750
		6.5	7.5	0.15	1:1.1538
		11	7.5	0.16	1:0.6818
		20	7.5	0.19	1:0.3750
	72 hrs	42.5	7.5	0.27	1:0.1765
		87.5	7.5	0.42	1:0.0857
		110	7.5	0.49	1:0.0682
		200	7.5	0.79	1:0.0375
		300	7.5	1.13	1:0.0250
		11	3.75	0.10	1:0.3409
		20	3.75	0.13	1:0.1875
		42.5	3.75	0.20	1:0.0882
		87.5	3.75	0.35	1:0.0429
		110	3.75	0.43	1:0.0341
		200	3.75	0.73	1:0.0188
		300	3.75	1.06	1:0.0125
		300	1.86	1.03	1:0.0062
		300	0.94	1.02	1:0.0031
		300	-	-	-
		-	60	-	-
		3	30	0.51	1:10.0000
		11	30	0.54	1:2.7273
		20	30	0.57	1:1.5000
		42.5	30	0.64	1:0.7059
		87.5	30	0.79	1:0.3429
		110	30	0.87	1:0.2727
		200	30	1.17	1:0.1500
		0.525	15	0.25	1:28.5714
		3	15	0.26	1:5.0000

		11	15	0.29	1:1.3636
		20	15	0.32	1:0.7500
		42.5	15	0.39	1:0.3529
		87.5	15	0.54	1:0.1714
		110	15	0.62	1:0.1364
		200	15	0.92	1:0.0750
		6.5	7.5	0.15	1:1.1538
		11	7.5	0.16	1:0.6818
		20	7.5	0.19	1:0.3750
		42.5	7.5	0.27	1:0.1765
		87.5	7.5	0.42	1:0.0857
		110	7.5	0.49	1:0.0682
		200	7.5	0.79	1:0.0375
		300	7.5	1.13	1:0.0250
		11	3.75	0.10	1:0.3409
		20	3.75	0.13	1:0.1875
		42.5	3.75	0.20	1:0.0882
		87.5	3.75	0.35	1:0.0429
		110	3.75	0.43	1:0.0341
		200	3.75	0.73	1:0.0188
		300	3.75	1.06	1:0.0125
		300	1.86	1.03	1:0.0062
		300	0.94	1.02	1:0.0031
<i>Ps. aeruginosa</i> ATCC#9027	24hrs	110	-	-	-
		-	75	-	-
		0.875	37.5	0.51	1:42.8571
		2	37.5	0.52	1:18.7500
		4.25	37.5	0.54	1:8.8235
		8.75	37.5	0.58	1:4.2857
		11	37.5	0.60	1:3.4091
		20	37.5	0.68	1:1.8750
		30	37.5	0.77	1:1.2500
		42.5	37.5	0.89	1:0.8824
		52.5	37.5	0.98	1:0.7143
		65	37.5	1.09	1:0.5769
		4.25	18.75	0.29	1:4.4118
		8.75	18.75	0.33	1:2.1429
		11	18.75	0.35	1:1.7045
		20	18.75	0.43	1:0.9375
		30	18.75	0.52	1:0.6250
		42.5	18.75	0.64	1:0.4412
		52.5	18.75	0.73	1:0.3571
		65	18.75	0.84	1:0.2885
		87.5	18.75	1.05	1:0.2143
		30	9.4	0.40	1:0.3133
		42.5	9.4	0.51	1:0.2212
		52.5	9.4	0.60	1:0.1790
		65	9.4	0.72	1:0.1446
		87.5	9.4	0.92	1:0.1074
		110	9.4	1.13	1:0.0855
		42.5	4.6	0.45	1:0.1082
		52.5	4.6	0.54	1:0.0876

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	48 hrs	65	4.6	0.65	1:0.0708
		87.5	4.6	0.86	1:0.0526
		110	4.6	1.06	1:0.0418
		200	-	-	-
		-	125	-	-
		0.2	75	0.60	1:375.0000
		2	75	0.61	1:37.5000
		4.25	75	0.62	1:17.6471
		8.75	75	0.64	1:8.5714
		11	75	0.66	1:6.8182
		20	75	0.70	1:3.7500
		30	75	0.75	1:2.5000
		42.5	75	0.81	1:1.7647
		52.5	75	0.86	1:1.4286
		65	75	0.93	1:1.1538
		87.5	75	1.04	1:0.8571
		4.25	37.5	0.32	1:8.8235
		8.75	37.5	0.34	1:4.2857
		11	37.5	0.36	1:3.4091
		20	37.5	0.40	1:1.8750
		30	37.5	0.45	1:1.2500
		42.5	37.5	0.51	1:0.8824
		52.5	37.5	0.56	1:0.7143
		65	37.5	0.63	1:0.5769
		87.5	37.5	0.74	1:0.4286
		110	37.5	0.85	1:0.3409
		200	37.5	1.30	1:0.1875
		42.5	18.75	0.36	1:0.4412
		52.5	18.75	0.41	1:0.3571
		65	18.75	0.48	1:0.2885
		87.5	18.75	0.59	1:0.2143
		110	18.75	0.70	1:0.1705
		200	18.75	1.15	1:0.0938
		42.5	9.4	0.29	1:0.2212
		52.5	9.4	0.34	1:0.1790
		65	9.4	0.40	1:0.1446
		87.5	9.4	0.51	1:0.1074
		110	9.4	0.63	1:0.0855
		200	9.4	1.08	1:0.0470
		87.5	4.6	0.47	1:0.0526
		110	4.6	0.59	1:0.0418
		200	4.6	1.04	1:0.0230
		110	2.3	0.57	1:0.0209
		200	2.3	1.02	1:0.0115
<i>S.aureus</i> ATCC#6538	24hrs	52.5	-	-	-
		-	15	-	-
		0.875	7.5	0.52	1:8.5714
		1.1	7.5	0.52	1:6.8182
		2	7.5	0.54	1:3.7500
		4.25	7.5	0.58	1:1.7647
		6.5	7.5	0.62	1:1.1538
		11	7.5	0.71	1:0.6818

		20	7.5	0.88	1:0.3750
		30	7.5	1.07	1:0.2500
		11	3.8	0.46	1:0.3455
		20	3.8	0.63	1:0.1900
		30	3.8	0.82	1:0.1267
		42.5	3.8	1.06	1:0.0894
		30	1.8	0.69	1:0.0600
		42.5	1.8	0.93	1:0.0424
		52.5	1.8	1.12	1:0.0343
		42.5	0.94	0.87	1:0.0221
		52.5	0.94	1.06	1:0.0179
	48hrs	65	-	-	-
		-	15	-	-
		2	7.5	0.53	1:3.7500
		4.25	7.5	0.57	1:1.7647
		6.5	7.5	0.6	1:1.1538
		8.75	7.5	0.63	1:0.8571
		11	7.5	0.67	1:0.6818
		20	7.5	0.81	1:0.3750
		30	7.5	0.96	1:0.2500
		42.5	7.5	1.15	1:0.1765
		20	3.8	0.56	1:0.1900
		30	3.8	0.71	1:0.1267
		42.5	3.8	0.91	1:0.0894
		52.5	3.8	1.06	1:0.0724
		65	1.8	1.12	1:0.0277
		65	0.94	1.06	1:0.0145

Table:6

TM

Ca: ppm AI of CAPMUL GML(glycerol monolaurate)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

5 Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	11000	-	-	-
		-	37.5	-	-
		200	18.8	0.52	1:0.0940
		425	18.8	0.54	1:0.0442
		650	18.8	0.56	1:0.0289
		1100	18.8	0.60	1:0.0171
		2000	18.8	0.68	1:0.0094
		3000	18.8	0.77	1:0.0063
		4250	18.8	0.89	1:0.0044
		5250	18.8	0.98	1:0.0036
		6500	18.8	1.09	1:0.0029
		1100	9.4	0.35	1:0.0085
		2000	9.4	0.43	1:0.0047
		3000	9.4	0.52	1:0.0031
		4250	9.4	0.64	1:0.0022
		5250	9.4	0.73	1:0.0018

	7 days	6500	9.4	0.84	1:0.0014
		11000	4.7	1.13	1:0.0004
		11000	2.4	1.06	1:0.0002
		11000	-	-	-
		-	37.5	-	-
		2000	18.8	0.68	1:0.0094
		3000	18.8	0.77	1:0.0063
		4250	18.8	0.89	1:0.0044
		5250	18.8	0.98	1:0.0036
		6500	18.8	1.09	1:0.0029
		1100	18.8	0.60	1:0.0171
		11000	9.4	1.25	1:0.0009
		11000	4.7	1.13	1:0.0004
		11000	2.4	1.06	1:0.0002
<i>C.albicans</i> ATCC # 10231	48 hrs	11000	-	-	-
		-	30	-	-
		525	15	0.55	1:0.0286
		875	15	0.58	1:0.0171
		1100	15	0.60	1:0.0136
		2000	15	0.68	1:0.0075
		3000	15	0.77	1:0.0050
		4250	15	0.89	1:0.0035
		5250	15	0.98	1:0.0029
		6500	15	1.09	1:0.0023
		4250	7.5	0.64	1:0.0018
		5250	7.5	0.73	1:0.0014
		6500	7.5	0.84	1:0.0012
		8750	7.5	1.05	1:0.0009
		5250	3.75	0.60	1:0.0007
		6500	3.75	0.72	1:0.0006
		8750	3.75	0.92	1:0.0004
		11000	3.75	1.13	1:0.0003
		5250	1.86	0.54	1:0.0004
		6500	1.86	0.65	1:0.0003
		8750	1.86	0.86	1:0.0002
		11000	1.86	1.06	1:0.0002
		11000	0.94	1.03	1:0.0001
	72 hrs	11000	-	-	-
		-	30	-	-
		1100	15	0.60	1:0.0136
		2000	15	0.68	1:0.0075
		3000	15	0.77	1:0.0050
		4250	15	0.89	1:0.0035
		5250	15	0.98	1:0.0029
		6500	15	1.09	1:0.0023
		5250	7.5	0.73	1:0.0014
		6500	7.5	0.84	1:0.0012
		8750	7.5	1.05	1:0.0009
		11000	1.86	1.06	1:0.0002
		11000	0.94	1.03	1:0.0001
<i>Ps. aeruginosa</i> ATCC#9027	24hrs	11000	-	-	-
		-	125	-	-

		525	75	0.65	1:0.1429
		650	75	0.66	1:0.1154
		875	75	0.68	1:0.0857
		1100	75	0.70	1:0.0682
		2000	75	0.78	1:0.0375
		3000	75	0.87	1:0.0250
		4250	75	0.99	1:0.0176
<i>S. aureus</i> ATCC#6538	24hrs	650	-	-	-
		-	15	-	-
		300	7.5	0.96	1:0.0250
		400	7.5	1.12	1:0.0188
	48hrs	2000	-	-	-
		-	15	-	-
		300	7.5	0.65	1:0.0250
		650	7.5	0.83	1:0.0115
		875	7.5	0.94	1:0.0086
		1100	7.5	1.05	1:0.0068

Table:7

Ca: ppm AI of CAPMULTM PG12 (propylene glycol mono-laurate)

Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)

5 Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	1100	-	-	-
		-	37.5	-	-
		110	18.8	0.60	1:0.1709
		200	18.8	0.68	1:0.0940
		300	18.8	0.77	1:0.0627
		425	18.8	0.89	1:0.0442
		525	18.8	0.98	1:0.0358
		650	18.8	1.09	1:0.0289
	7 days	1100	2.4	1.06	1:0.0022
		1100	-	-	-
		-	75	-	-
		300	37.5	0.77	1:0.1250
		425	37.5	0.89	1:0.0882
		525	37.5	0.98	1:0.0714
		1100	2.4	1.03	1:0.0022
<i>C. albicans</i> ATCC # 10231	48 hrs	1100	-	-	-
		-	30	-	-
		52.5	15	0.55	1:0.2857
		110	15	0.60	1:0.1364
		200	15	0.68	1:0.0750
		300	15	0.77	1:0.0500
		425	15	0.89	1:0.0353
		525	15	0.98	1:0.0286
		650	15	1.09	1:0.0231
		1100	0.94	1.03	1:0.0009
	72 hrs	1100	-	-	-

		-	30	-	-
		52.5	15	0.55	1:0.2857
		110	15	0.60	1:0.1364
		200	15	0.68	1:0.0750
		300	15	0.77	1:0.0500
		425	15	0.89	1:0.0353
		525	15	0.98	1:0.0286
		650	15	1.09	1:0.0231
		1100	0.94	1.03	1:0.0009

Table:8

- 5 Ca: ppm AI of CAPMUL PG8 (propylene glycol caprylate)
 Cb: ppm AI of MBIT(N-methyl-1,2-benzisothiazolin-3-one)
 Ratio:Ca:Cb

Test Organisms	Contact Time	Ca	Cb	S.I.	Ca:Cb
<i>A. niger</i> ATCC # 16404	3 days	875	-	-	-
		-	37.5	-	-
		42.5	18.8	0.55	1:0.4424
		65	18.8	0.58	1:0.2892
		87.5	18.8	0.60	1:0.2149
		110	18.8	0.63	1:0.1709
		200	18.8	0.73	1:0.0940
		300	18.8	0.84	1:0.0627
		425	18.8	0.99	1:0.0442
		425	9.4	0.74	1:0.0221
		525	9.4	0.85	1:0.0179
		650	9.4	0.99	1:0.0145
		525	4.7	0.73	1:0.0090
		650	4.7	0.87	1:0.0072
		875	4.7	1.13	1:0.0054
	7 days	650	2.4	0.81	1:0.0037
		875	2.4	1.06	1:0.0027
		650	1.2	0.77	1:0.0018
		875	1.2	1.03	1:0.0014
		1100	-	-	-
		-	75	-	-
		525	37.5	0.98	1:0.0714
		650	37.5	1.09	1:0.0577
		650	18.8	0.84	1:0.0289
		875	18.8	1.05	1:0.0215
		875	9.4	0.92	1:0.0107
		1100	9.4	1.13	1:0.0085
		875	4.7	0.86	1:0.0054
		1100	4.7	1.06	1:0.0043
		875	2.4	0.83	1:0.0027
		1100	2.4	1.03	1:0.0022
		1100	1.2	1.02	1:0.0011

<i>C.albicans</i> ATCC # 10231	48 hrs	200	-	-	-
		-	30	-	-
		65	15	0.83	1:0.2308
		87.5	15	0.94	1:0.1714
		110	15	1.05	1:0.1364
		110	7.5	0.80	1:0.0682
		200	3.75	1.13	1:0.0188
		200	1.86	1.06	1:0.0093
	72 hrs	200	0.94	1.03	1:0.0047
		200	-	-	-
		-	30	-	-
		65	15	0.83	1:0.2308
		87.5	15	0.94	1:0.1714
		110	15	1.05	1:0.1364
		200	0.94	1.03	1:0.0047

CLAIMS:

1. A microbicidal composition comprising:
 - (a) N-methyl-1,2-benzisothiazolin-3-one; and
 - (b) at least one microbicide selected from the group consisting of (i) Cu(II) 2-aminoethanolate and (ii) Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride.
2. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a weight ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:375 to 1:0.004.
3. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride, and a weight ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:176 to 1:0.08.
4. The microbicidal composition of claim 2 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:375 to 1:0.8824.
5. The microbicidal composition of claim 2 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:1.7091 to 1:0.0157.

6. The microbicidal composition of claim 2 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:0.0427 to 1:0.0090.

7. The microbicidal composition of claim 2 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is from 1:0.0056 to 1:0.0046.

8. The microbicidal composition of claim 2 in which said at least one microbicide is Cu(II) 2-aminoethanolate, and a ratio of Cu(II) 2-aminoethanolate to N-methyl-1,2-benzisothiazolin-3-one is about 1:0.0040.

9. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride, and a ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:176.4706 to 1:0.9375.

10. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride, and a ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:9.400 to 1:0.1200.

11. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride, and a ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is from 1:0.1446 to 1:0.1074.

12. The microbicidal composition of claim 1 in which said at least one microbicide is Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride, and a ratio of Cu(II) 2-aminoethanolate and didecyl dimethyl ammonium chloride to N-methyl-1,2-benzisothiazolin-3-one is about 1:0.080.