STATUS DECISION OF CONTROLLED AND NON-CONTROLLED SUBSTANCE(S)

Substance:
I – 8-Chloro-3a,4-dihydro-6-(2-fluorophenyl)-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazepine
II - 8-Chloro-6-(fluorophenyl)-3a,4,5,6-tetrahydro-1-methyl-3H-imidazo-[1,5a][1,4]-
benzodiazepine, dihydrochloride

Based on the current information available to the Office of Controlled Substances, it appears that the above substance is:

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<tbody>
<tr>
<td>Controlled</td>
<td>X</td>
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<tr>
<td>Not Controlled</td>
<td>□</td>
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under the schedules of the *Controlled Drugs and Substances Act* (CDSA) for the following reason(s):

- The substances are impurities of midazolam and captured under item 18 of Schedule IV to the CDSA.

Prepared by: ________________________________ Date: ________________

Evelyn C Soo

Verified by: ________________________________ Date: ________________

Mark Kozlowski

Approved by: ________________________________ Date: ________________

DIRECTOR, OFFICE OF CONTROLLED SUBSTANCES

This status was requested by: Nacer Silarbi, OCS
Drug Status Report

Drug:
I – 8-Chloro-3a,4-dihydro-6-(2-fluorophenyl)-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazepine
II - 8-Chloro-6-(fluorophenyl)-3a,4,5,6-tetrahydro-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazepine

Drug Name Status:
I - 8-Chloro-6-(2-fluorophenyl)-1-methyl-4H-benzo[f]imidazo[1,5-a][1,4]diazepine
II - 8-Chloro-6-(2-fluorophenyl)-1-methyl-5,6-dihydro-4H-benzo[f]imidazo[1,5-a][1,4]diazepine
are the common names.

Chemical Name:
I - 8-Chloro-6-(2-fluorophenyl)-1-methyl-4H-benzo[f]imidazo[1,5-a][1,4]diazepine
II - 8-Chloro-6-(2-fluorophenyl)-1-methyl-5,6-dihydro-4H-benzo[f]imidazo[1,5-a][1,4]diazepine

Other Names:
I – RMID
II –RRMID

Chemical structure:

Molecular Formula:  I – C_{18}H_{13}ClF_{3}, II – C_{18}H_{15}ClF_{3}

Pharmacological class / Application:  Pharmaceutical-related substances

CAS-RN: I, II – N/A

International status:
US:  8-Chloro-3a,4-dihydro-6-(2-fluorophenyl)-1-methyl-3H-imidazo-[1,5a][1,4]-
benzodiazepine and 8-chloro-6-(fluorophenyl)-3a,4,5,6-tetrahydro-1-methyl-3H-imidazo-
[1,5a][1,4]-benzodiazepine are not listed specifically in the Schedules to the US Controlled
Substances Act and are not mentioned anywhere on the DEA website.

United Nations:  The substances are not listed on the Yellow List - List of Narcotic Drugs under
International Control, the Green List - List of Psychotropic Substances under International
Canadian Status: 8-Chloro-3a,4-dihydro-6-(2-fluorophenyl)-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazipine and 8-chloro-6-(fluorophenyl)-3a,4,5,6-tetrahydro-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazipine are not listed specifically in the Schedules to the CDSA. The substances are impurities of midazolam which is currently listed as sub-item 18(22.1) under the heading “Midazolam (8-chloro-6-(o-fluorophenyl)-1-methyl-4H-imidazo[1,5a][1,4]benzodiazipine)” in Schedule IV to the CDSA. The substances also contain the core structure of the 1,4-benzodiazipines. Therefore they are considered captured under the heading “Benzodiazipines, their salts and derivatives including” of item 18 of Schedule IV to the CDSA.

Recommendation: 8-Chloro-3a,4-dihydro-6-(2-fluorophenyl)-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazipine and 8-chloro-6-(fluorophenyl)-3a,4,5,6-tetrahydro-1-methyl-3H-imidazo-[1,5a][1,4]-benzodiazipine are included under item 18 of Schedule IV to the CDSA and are controlled substances.

Date: November 1st, 2012.