

## 1. Identification

#### **GHS PRODUCT IDENTIFIER**

Product name: Cap Mix A (THF/Pyridine/Acetic Anhydride 8:1:1)

#### OTHER MEANS OF IDENTIFICATION

Item number:	4110
Catalogue number(s):	4110-YZZZ, where Y=letters A-Z, ZZZ=numbers 000-999
Brand:	Not applicable.

#### RECOMMENDED USE OF THE CHEMICAL AND RESTRICTIONS ON USE

For laboratory and manufacturing use. Not for drug, household or other use.

#### SUPPLIER'S DETAILS

Link Technologies Ltd	Tel:	+44 (0) 1698 849911
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#### **EMERGENCY PHONE NUMBER**

+44 (0) 1698 849911 (Monday to Friday 8 am to 6 pm)

## 2. Hazard identification

#### GHS CLASSIFICATION OF THE SUBSTANCE/MIXTURE

Classification according to Regulation (EC) No. 1272/2008Flammable Liquid:Category 2Acute ToxicityCategory 4Skin CorrosionCategory 1BEye Irritation:Category 2CarcinogenicityCategory 2Specific Target Organ Toxicity Single exposure –Respiratory systemCategory 3

Classification according to EU Directives 67/548/EEC or 1999/45/EC [Pertinent to tetrahydrofuran and pyridine; no data available for acetic anhydride]



R19	May form explosive peroxides.
F; R11	Highly flammable.
Xi; R36/37	Irritating to eyes and respiratory system.
Carc. Cat. 3; R40	Limited evidence of a carcinogenic effect.
Xn; R20/21/22	Harmful by inhalation, in contact with skin and if swallowed
Xi; R36/38	Irritating to eyes and skin.

#### GHS LABEL ELEMENTS, INCLUDING PRECAUTIONARY STATEMENTS

#### Pictogram



Signal word: Danger

#### **Hazard Statements**

H225:	Highly flammable liquid and vapour.
H302:	Harmful if swallowed.
H314	Causes severe skin burns and eye damage
H318:	Causes serious eye damage.
H332:	Harmful if inhaled.
H335:	May cause respiratory irritation.
H351:	Suspected of causing cancer.

#### **Precautionary Statements**

# PreventionP210Keep away from heat, hot surfaces, sparks, open flames and other ignition sources. No smoking.P233Keep container tightly closed.P243Take actions to prevent static discharges.P260Do not breathe vapours.P264Wash hands thoroughly after handlingP280Wear protective gloves/protective clothing/eye protection/face protection.P403Store in a well-ventilated place.

#### **Response**

P301 + P330 + P331	IF SWALLOWED: Rinse mouth. Do NOT induce vomiting.
P303 + P361 + P353	IF ON SKIN: Take off immediately all contaminated clothing. Rinse skin with water/shower.
P304 + P340	IF INHALED: Remove person to fresh air and keep comfortable for breathing.



P305 + P351 + P338	IF IN EYES: Rinse cautiously with water for several minutes. Remove contact lenses, if present		
	and easy to do. Continue rinsing.		
P310	Immediately call a POISON CENTRE/doctor.		
P337 + P313	If eye irritation persists: Get medical advice/attention.		
P370 + P378	In case of fire: Use $CO_2$ to extinguish.		
P501	Dispose of contents/container to in accordance with local/regional/national /international		
	regulations.		

#### **OTHER HAZARDS**

EUH019 May form explosive peroxides.

Additionally, the classification provided by companies to ECHA in REACH registrations identifies that acetic anhydride is fatal if inhaled.



## 3. Composition/information on ingredients

#### MIXTURES

#### Hazardous ingredients according to Regulation (EC) No 1272/2008

Component		Classification	Concentration
Tetrahydrofuran			
CAS No	109-99-9	Flammable Liquid 2; Acute Tox. 4; Eye	80%
EC No	203-726-8	Irritation 2A; STOT SE 3; Carcinogen 2.	
Index No	603-025-00-0	H225, H302, H319, H335, H351.	
Registration No	01-2119444314-46-XXXX	EUH019.	
Acetic Anhydride			
CAS No	108-24-7	Flammable Liquid 3; Acute Tox. 4;	10%
EC No	203-564-8	Skin Corr. 1B; Eye Dam. 1.	
Index No	607-008-00-9	H226, H332, H314, H318.	
Registration No	01-2119433003-58-XXXX		
Pyridine			
CAS No	110-86-1	Flammable Liquid 2; Acute Tox. 4;	10%
EC No	203-809-9	Skin Irrit. 2; Eye Irrit. 2.	
Index No	613-002-00-7	H225, H302, H312, H332, H315, H320.	
Registration No	01-2119493105-40-XXXX		



#### Hazardous ingredients according to Directive 1999/45/EC

Component		Classification	Concentration
Tetrahydrofuran			
CAS No	109-99-9	R19 May form explosive peroxides.	80%
EC No	203-726-8	F; R11 Highly flammable.	
Index No	603-025-00-0	Xi; R36/37 Irritating to eyes and	
Registration No	01-2119444314-46-XXXX	respiratory system.	
		Carc. Cat. 3; R40 Limited evidence of a	
		carcinogenic effect.	
Acetic Anhydride			
CAS No	108-24-7	No data available.	10%
EC No	203-564-8		
Index No	607-008-00-9		
Registration No	01-2119433003-58-XXXX		
Pyridine			
CAS No	110-86-1	F; R11 Highly flammable.	10%
EC No	203-809-9	Xn; R20/21/22 Harmful by inhalation, in	
Index No	613-002-00-7	contact with skin and if swallowed.	
Registration No	01-2119493105-40-XXXX	Xi; R36/38 Irritating to eyes and skin.	

## 4. First aid measures

#### DESCRIPTION OF NECESSARY MEASURES

#### **General advice**

Consult a physician. Show this safety data sheet to the doctor in attendance.

#### If inhaled

If breathed in, move person into fresh air. If not breathing, give artificial respiration.

#### In case of skin contact

Remove any contaminated clothing. Wash off with soap and plenty of water.

#### In case of eye contact

Flush eyes copiously with water for at least 15 minutes. Use a sterile eye wash if available.

#### If swallowed

Do NOT induce vomiting. Keep person calm and immobile. Rinse mouth with water if conscious. Never give anything by mouth to an unconscious person.



#### MOST IMPORTANT SYMPTOMS/EFFECTS, ACUTE AND DELAYED

To the best of our knowledge, the chemical, physical, and toxicological properties have not been thoroughly investigated. See section 11.

#### INDICATION OF IMMEDIATE MEDICAL ATTENTION AND SPECIAL TREATMENT NEEDED, IF NECESSARY

No data available.

## 5. Fire-fighting measures

#### SUITABLE (AND UNSUITABLE) EXTINGUISHING MEDIA

Use water spray, alcohol-resistant foam, dry chemical or carbon dioxide.

# SPECIFIC HAZARDS ARISING FROM THE CHEMICAL (E.G. NATURE OF ANY HAZARDOUS COMBUSTION PRODUCTS)

Possibility of carbon oxides, nitrogen oxides (NOx).

#### SPECIAL PROTECTIVE EQUIPMENT AND PRECAUTIONS FOR FIRE-FIGHTERS

Wear mask and protective clothing to prevent contact with skin and eyes. Wear self-contained breathing apparatus if necessary.

#### 6. Accidental release measures

#### PERSONAL PRECAUTIONS, PROTECTIVE EQUIPMENT AND EMERGENCY PROCEDURES

Wear protective clothing, respirator, chemical safety goggles, rubber gloves and rubber boots. Avoid breathing vapours, mist or gas. Ensure adequate ventilation. Remove all sources of ignition. Evacuate personnel to safe areas. Be aware of vapours accumulating to form explosive concentrations. Vapours can accumulate in low areas.

#### **ENVIRONMENTAL PRECAUTIONS**

Prevent further leakage or spillage, if safe to do so. Prevent product from entering drains.

#### METHODS AND MATERIALS FOR CONTAINMENT AND CLEANING UP

Contain spills using absorbent barriers where available. Clean the contaminated area thoroughly with water taking care to avoid breathing fumes. Dispose of all cleaning materials with care (see section 13), where possible containing in sealed containers for appropriate disposal.



## 7. Handling and storage

#### PRECAUTIONS FOR SAFE HANDLING

Avoid contact with skin and eyes. Avoid inhalation of vapour or mist. Do not eat, drink or smoke when using this product. Keep away from sources of ignition. Take measures to prevent build-up of electrostatic charge.

#### CONDITIONS FOR SAFE STORAGE, INCLUDING ANY INCOMPATIBILITIES

Containers should be kept sealed and safely stored when not in use. Store in a cool, dry, well-ventilated area. Containers which are opened must be carefully resealed and kept upright to prevent leakage. Store under inert gas. Dry residue can be explosive. Test for peroxide formation periodically and before distillation.

## 8. Exposure controls/personal protection

#### CONTROL PARAMETERS (OCCUPATIONAL EXPOSURE LIMIT VALUES OR BIOLOGICAL LIMIT VALUES)

Component	CAS No	Value	Control Parameters	Basis
Tetrahydrofuran	109-99-9	STEL	100ppm 300mg/m <sup>3</sup>	Europe – Commission Directive 2000/39/EC
		TWA	50ppm 150mg/m <sup>3</sup>	UK – EH40/2005 – Workplace Exposure Limits
	Remarks	Identifies the	e possibility of sigr	nificant uptake through the skin.
Acetic Anhydride	108-24-7	STEL	2ppm 10mg/m <sup>3</sup>	UK – EH40/2005 – Workplace Exposure Limits
		TWA	0.5ppm 2.5mg/m <sup>3</sup>	
Pyridine	110-86-1	STEL	10ppm 33mg/m <sup>3</sup>	UK – EH40/2005 – Workplace Exposure Limits
		TWA	5ppm 16mg/m <sup>3</sup>	
	Remarks	Identifies the possibility of significant uptake through the skin.		

#### APPROPRIATE ENGINEERING CONTROLS

General good industrial laboratory hygiene and safety practice. Use product within air-extracted fume hood where possible. Wash hands before breaks and at the end of the workday.



#### INDIVIDUAL PROTECTION MEASURES, SUCH AS PERSONAL PROTECTIVE EQUIPMENT

#### Eye/face protection

Tightly fitting safety goggles. Face shield (8-inch minimum). Use equipment for eye protection tested and approved under appropriate government standards such as NIOSH (US) or EN 166(EU).

#### **Skin protection**

Handle with gloves. Gloves must be inspected prior to use. Use proper glove removal technique (without touching glove's outer surface) to avoid skin contact with this product. Dispose of contaminated gloves after use in accordance with applicable laws and good laboratory practices. Wash and dry hands. The selected protective gloves have to satisfy the specifications of EU Directive 89/686/EEC and the standard EN 374 derived from it.

Splash contact - 0.3mm butyl rubber

Breakthrough time: 10 min

#### **Body Protection**

Complete suit protecting against chemicals, flame-retardant antistatic protective clothing. The type of protective equipment must be selected according to the quantity and concentration of the dangerous substance at the specific workplace.

#### **Respiratory protection**

Where risk assessment shows air-purifying respirators are appropriate, use a full-face respirator with multi-purpose combination (US) or type ABEK (EN14387) respirator cartridges as a back-up to engineering controls. If the respirator is the sole means of protection, use a full-face supplied air respirator. Use respirators and components tested and approved under appropriate government standards such as NIOSH (US) or CEN (EU).

## 9. Physical and chemical properties

Appearance (physical state, colour etc.):	Colourless to yellow liquid.
Odour:	Pungent odour
Odour threshold:	No data available
pH:	No data available
Melting point/freezing point:	No data available
Initial boiling point and boiling range:	No data available
Flash point:	-14.4°C
Evaporation rate:	No data available
Flammability (solid, gas):	Highly flammable
Upper/lower flammability or explosive limits:	No data available
Vapor pressure:	No data available
Vapor density:	No data available
Relative density:	No data available
Water Solubility:	Fully soluble
Partition coefficient: n-octanol/water:	No data available
Auto-ignition temperature:	No data available
Decomposition temperature:	No data available
Viscosity:	No data available



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Empirical formula: Molecular weight (g/mol): Not applicable. Not applicable.

## **10. Stability and reactivity**

#### REACTIVITY

No data available.

#### CHEMICAL STABILITY

Stable under recommended storage conditions.

#### POSSIBILITY OF HAZARDOUS REACTIONS

No data available.

#### CONDITIONS TO AVOID (E.G. STATIC DISCHARGE, SHOCK OR VIBRATION)

Heat, flames and sparks. Extremes of temperature and direct sunlight.

#### **INCOMPATIBLE MATERIALS**

Oxidising agents, reducing agents, bases, acid chlorides, powdered metals, water, alcohols, acids, chloroformates.

#### HAZARDOUS DECOMPOSITION PRODUCTS

Possibility of carbon oxides or nitrogen oxides (NOx) on combustion.

## 11. Toxicological information

#### **TOXIC EFFECTS**

#### Tetrahydrofuran

*Acute oral toxicity* - Based on the results of acute oral toxicity testing, tetrahydrofuran is classified as Xn;R22 (Harmful if swallowed) under the EU DSD classification criteria (EU Directive 67/548/EEC). It is assigned an Acute Toxicity Category 4 rating (Harmful if swallowed) under the EU CLP classification criteria (EU Regulation 1272/2008). *Acute inhalation toxicity* - Based on the clear presence of CNS depression in animal studies, tetrahydrofuran is classified as R67 (Vapours may cause drowsiness and dizziness) under the EU DSD classification criteria. Based on the presence of upper respiratory tract irritation in animal studies, tetrahydrofuran would be classified as Xi;R37 (Irritating to respiratory system) under the EU DSD classification criteria. Tetrahydrofuran would be rated STOT SE 3, based on either CNS depression or upper respiratory tract irritation according to the EU CLP classification criteria. *Acute dermal toxicity* - Based on the results of the key study (rat dermal LD50 >2000 mg/kg), tetrahydrofuran would not be classified under the EU DSD classification criteria. Similarly, tetrahydrofuran would not receive classification under the EU CLP classification criteria.



*Skin irritation* - Based on the results of the key study (Primary Irritation Index of 1.93), tetrahydrofuran would not be rated a primary dermal irritant under the EU DSD criteria (EU Directive 67/548/EEC). Tetrahydrofuran would also not be classified for skin irritation under the EU CLP criteria (EU Regulation 1272/2008).

*Eye irritation* - Based on the results of the key study, tetrahydrofuran would be classified as Xi;R41 (Risk of serious damage to eyes) under the EU DSD criteria (EU Directive 67/548/EEC). This rating is based on severe and possibly irreversible effects on the eye. In the case of the EU CLP criteria (EU Regulation 1272/2008), tetrahydrofuran would be rated Category 1 (Serious eye damage/eye irritation).

*Genetic Toxicity* - THF was generally negative when tested for mutagenicity/cytogenicity in either in vitro or in vivo assays. Based on this evidence, THF would not be rated as a mutagen under either the EU DSD classification system (EU Directive 67/548/EEC) or the EU CLP classification system (EU Regulation 1272/2008).

#### Germ cell mutagenicity - No data available.

*Carcinogenicity* - Tetrahydrofuran has been tested for carcinogenicity in chronic studies on rats and mice by the National Toxicology Program. Critical target organs for neoplasia in these studies included the liver for mice, and kidney in male rats. In exposed male mice, the incidences of hepatocellular tumours were not significantly different from the chamber controls. The lower incidences of hepatocellular neoplasms in male mice at the 1800 ppm exposure concentration was attributed to the reduced survival of this group. The incidences of combined hepatocellular neoplasms in male mice exposed to 200 ppm and in the chamber controls exceeded the historical control range for inhalation studies. The incidence of combined hepatocellular neoplasms in male mice at the 600 ppm exposure concentration was at the upper limit of the historical control range. Female mice exposed at 1800 ppm had significantly greater incidences of hepatocellular neoplasms were increased with a positive trend. In addition, the incidences of multiple hepatocellular neoplasms were increased in female mice exposed at 1800 ppm. The incidences of combined hepatocellular neoplasms in female mice exposed to 1800 ppm exceeded the historical range for chamber controls in 2-year NTP inhalation studies. Considering that THF is non-genotoxic, and that the hepatocellular tumours occurred at high, liver enzyme inducing doses, the liver tumours observed in this strain of mice are unlikely to be relevant to human cancer risk.

In rats, Incidences of renal tubule epithelial adenoma were marginally increased in 600 and 1800 ppm males. Female rats did not exhibit renal neoplasia. The nature of these tumours was reviewed and concluded to be associated with age-related chronic progressive nephropathy, a physiological condition common in male rats and unlikely to be of relevance to human cancer risk.

Based on these findings, THF should not be rated for carcinogenicity. However there is a harmonised classification for this substance as a Category 2 carcinogen under the EU CLP classification system (EC No 1272/2008), and this will be applied to the substance.

*Reproductive toxicity* - Tetrahydrofuran is not a selective developmental toxicant based on studies conducted in Wistar rats and in Swiss CD-1 mice. In mice, embryotoxicity was observed at an inhalation exposure concentration (1800 ppm) also causing significant maternal effects. Similarly in rats, no selective effect on the developing foetus is observed. Based on the evidence presented, tetrahydrofuran should not be rated as a reproductive hazard (Substances which cause concern for human fertility) under the EU DSD classification system (EU Directive 647/548/EEC). Similarly, tetrahydrofuran would not be rated under the EU CLP classification system (EU Regulation 1272/2008). *STOT-single exposure* - Tetrahydrofuran would be rated STOT SE 3, based on either CNS depression or upper respiratory tract irritation according to the EU CLP classification criteria.



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*STOT-repeated exposure* - Repeated-dose drinking water toxicity studies in rats indicate tetrahydrofuran is of low toxicity. The subacute and subchronic toxicity of tetrahydrofuran in rats and mice is generally moderate to low following repeated-dose inhalation exposures. Mice are more sensitive than rats following repeated inhalation exposures. *Aspiration hazard* - No data available

#### Acetic Anhydride

Acute toxicity:	No data available
Skin corrosion/irritation:	No data available
Serious eye damage/irritation:	No data available
Respiratory or skin sensitization:	No data available
Germ cell mutagenicity:	No data available
Carcinogenicity:	No data available
Reproductive toxicity:	No data available
STOT-single exposure:	No data available
STOT-repeated exposure:	No data available
Aspiration hazard:	No data available

#### Pyridine

*Acute oral toxicity* - Toxicity Category IV. The oral LD50 of pyridine is between 800 and 1600 mg/kg bw in rats. *Acute inhalation toxicity* - Toxicity Category IV. Pyridine was tested in a nose-only exposure protocol in a guideline acute inhalation test. The LC50 in male rats after 4 hours was 4900 ppm or 15852 mg/m3.

*Acute toxicity dermal* - Toxicity Category IV. Pyridine was testing in albino rabbits in an acute dermal toxicity test at 500, 1000 and 2000 mg/kg bw. No deaths occurred at 1000 mg/kg or lower, but all animals died at 2000 mg/kg bw. The LD50 is between 1000 and 2000 mg/kg bw.

*Skin irritation* – Corrosive. Pyridine, applied undiluted to intact and abraded rabbit skin, was corrosive at 24 and 72 hours.

*Eye irritation* – Moderately irritating. Pyridine was evaluated as a moderate eye irritant with corneal damage in rabbits. *Genetic toxicity* - Cytotoxicity and mutagenic activity was not observed with pyridine in the presence and absence of metabolic activation, according to OECD 471 bacterial reverse mutation assay guidelines. *Germ cell mutagenicity* - No data available.

*Carcinogenicity* - A 2 year carcinogenicity toxicity study in F344 rats was undertaken with pyridine at doses from 100-400 ppm in the drinking water (7 to 33 mg/kg bw/d). Consumption of water in males and females was increased over control levels, but body weights were significantly lower in the 400 ppm dose group compared with controls. Doses of 400 ppm resulted in a statistically significant increase in renal adenomas and renal tubule hyperplasia, but not renal carcinomas, in males. Females (but not males) displayed a statistically significant increase in mononuclear cell leukemia. Animals of both sexes displayed an increased incidence of non-neoplastic hepatocellular injury and glandular stomach mineralization. The NOAEL was 100 ppm (7 mg/kg bw/d). The NTP concluded that there was some evidence of carcinogenic activity in male F344 rats based on renal adenomas, and equivocal evidence in female F344 rats of carcinogenic activity of pyridine based on mononuclear cell leukemia.

*Reproductive toxicity* - There were no significant adverse findings in male fertility in F344 rats (sperm parameters and testis weights) after 14 weeks of exposure in drinking water to 3-methylpyridine, a category member with pyridine and 2- and 4-methylpyridine. There were no structural effects of this treatment in females, although there was a suggestion of



altered estrus cycling at doses of 325 mg/L and higher. This finding may be the result of other complications of the study, and is not highly relevant for human risk assessment. The NOAEL is 156 mg/L, equivalent to 22-23 mg/kg bw/d. *STOT-single exposure* - Toxicity Category IV.

STOT-repeated exposure – see Carcinogenicity above.

Aspiration hazard - No data available

#### INFORMATION ON THE LIKELY ROUTES OF EXPOSURE

Exposure via inhalation, ingestion, skin and eye contact – full data on the health effects of such exposure is not available.

#### SYMPTOMS RELATED TO THE PHYSICAL, CHEMICAL AND TOXICOLOGICAL CHARACTERISTICS

No data available. To the best of our knowledge the chemical, physical and toxicological properties of this substance have not been investigated. Handle with care to avoid all unnecessary exposure and release into the atmosphere. If you feel unwell after using this product then immediately seek medical attention.

# DELAYED AND IMMEDIATE EFFECTS AND ALSO CHRONIC EFFECTS FROM SHORT AND LONG TERM EXPOSURE

Components can cause central nervous system depression, cough, chest pain, difficulty in breathing. Exposure to high airborne concentrations may cause an anaesthetic effect.

#### NUMERICAL MEASURES OF TOXICITY (SUCH AS ACUTE TOXICITY ESTIMATES)

No data available.

## 12. Ecological information

#### TOXICITY

#### Tetrahydrofuran

Based on the short-term toxicity to fish test (which provided the lowest LC50 of all three trophic levels), tetrahydrofuran is not considered classified as hazardous to the environment according to the following criteria: Directive 67/548/EEC; UN GHS; Regulation 1272/2008 (CLP GHS).

Acetic Anhydride

No data available.

#### Pyridine

No data available.

#### PERSISTENCE AND DEGRADABILITY

No data available.



#### **BIOACCUMULATIVE POTENTIAL**

No data available.

MOBILITY IN THE SOIL

No data available.

#### **OTHER ADVERSE EFFECTS**

No data available.

## **13. Disposal Considerations**

#### **DISPOSAL METHODS**

For the safety of persons conducting disposal, recycling or reclamation activities, please refer to the information in section 8 of the SDS. Dispose by incineration at high temperature in an approved incinerator fitted with appropriate environmental protection equipment taking extra care in igniting, as this material is highly flammable. Contaminated packaging should be treated as product. Dispose of in accordance with all applicable Local, National, State and Federal regulations. Labels should not be removed from containers until they have been thoroughly cleaned in an appropriate manner. Containers should not be treated as domestic waste and disposed of appropriately. Always use an approved disposal company. Do not dispose to drains.

## 14. Transport information

UN number			
ADR/RID: 29	924	IMDG: 2924	IATA: 2924
UN proper s	hipping nam	ne	
ADR/RID:	FLAMMABL	E LIQUID, N.O.S. (Tetral	ydrofuran, Acetic Anhydride)
IMDG:	FLAMMABL	E LIQUID, N.O.S. (Tetral	ydrofuran, Acetic Anhydride)
IATA:	Flammable	liquid, N.O.S. (Tetrahydro	furan, Acetic Anhydride)
Transport hazard class(es)			
ADR/RID: 3	(8)	IMDG: 3 (8)	IATA: 3 (8)
Packing gro	oup		
ADR/RID: II		IMDG: II	IATA: II
Environmer	ntal hazards		
IMDG Marine	e Pollutant:	No	
ADR/RID:		No	
IATA: No		No	
Special precautions for the user			
No data available.			



## 15. Regulatory information

This safety datasheet references: the Globally Harmonized System of Classification and Labelling of Chemicals (GHS), Fourth revised edition, 2011; European Commission Directive 2000/39/EC; UK Health and Safety Executive EH40/2005 Workplace Exposure Limits; and the European Chemicals Agency (ECHA) <u>https://echa.europa.eu/</u>. No further safety, health and environmental regulations specific for the product in question are available.

## 16. Other information

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Approved: 02 April 2018

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