Annexure-VI

Decision for Cleaning Validation

Ref. Doc No. CVMP/ "Cleaning Validation"

A. If the evaluation identifies the molecule as Genotoxicity and Carcinogenicity

Parameter				
	Relatively safe	**Low hazard**	**Moderate hazard**	**Highly hazard**
Genotoxicity	Not identified in any study	Positive in in-vitro studies	Possible / probable	Known in Human
Carcinogenecity	Not identified in any study	Not identified in any study	Possible / probable	Known in Human
Requirements for cleaning validation and routine control	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	a. All molecules will be considered for cleaning validation b. Control procedures to be implemented during routine manufacturing based on risk assessment	Molecule require dedicated facility

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B. If the evaluation identifies the molecule as Reproductive and development toxicity

Parameter				
	Relatively safe	**Low hazard**	**Moderate hazard**	**Highly hazard**
Reproductive and developmental toxicity	Not identified in any study	a. More than 3 times recommended daily dose b. Identified only in animal studies beyond 12 months	a. Precautionary indications in patient information leaflet b. Clinical ≤ 50 mg / day to > 10 mg / day c. Animal studies ≤ 30 mg / day > 1 mg / day	Clinical : ≤ 10 mg / day Animal studies: ≤ 1 mg / kg / day
Requirements for cleaning validation and routine control	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	All molecules will be considered for cleaning validation along with control procedure during routine manufacturing	Molecule require dedicated area / equipment

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C. If the evaluation identifies the molecule as Highly pharmacological potency

Parameter			
	Low hazard	**Moderate hazard**	**Highly hazard**
Highly pharmacological potency	**Recommended daily dose >10 mg**	**Recommended daily dose ≤ 10 mg and > 1 mg**	**Recommended daily dose ≤ 1 mg**
Requirements for cleanin validation and routine control	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	All molecules will be considered for cleaning validation if the solubility is very slightly soluble and above

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D. If the evaluation identifies the molecule as sensitizing potential

Parameter			
	Low hazard	**Moderate hazard**	**Highly hazard**
Sensitizing potential	Not identified in any study	Positive for skin and Respiratory	Highly sensitizing potential for skin and Respiratory
Requirements for cleaning validation and routine control	Molecule to be evaluated as worst based on Solubility / Potency and cleanability	All molecules will be considered for cleaning validation along with control procedure during routine manufacturing and risk assessment	Molecule require dedicated area