IN VITRO METHODS FOR PREDICTING SKIN IRRITATION AND CORROSION

START here if you suspect your test substance is corrosive **TOP-DOWN APPROACH** OECD TG 431 or 435 STOP testing: Test substance Test substance in in vitro skin should be labelled as corrosive irritation assay: OECD TG 439

STOP testing: Test substance should STOP testing: Test substance is be labelled as an irritant not classified as an irritant

START here if you suspect your test substance is not corrosive **BOTTOM-UP APPROACH OECD TG 439** Test substance in *in vitro* skin STOP testing: Test substance is not corrosion assay: OECD TG 431 or 435 classified as an irritant STOP testing: Test substance should STOP testing: Test substance should be labelled as corrosive be labelled as an irritant

METHOD	PRINCIPLE OF THE TEST	RHE MODEL (IF APPLICABLE)	APPLICABILITY DOMAIN	GHS CATEGORISATION
OECD TG 439: In Vitro Skin Irritation: Reconstructed Human Epidermis Test Method	Test substance is applied to a 3-dimensional reconstructed human epidermis (RhE) model. Following exposure and a post-exposure incubation period, the vital dye MTT is added and cell viability determined. Optional histology may also be conducted to gain further information.	 EpiSkin™ (L'Oréal, France) EpiDerm™ (MatTek, US) SkinEthic™ (L'Oréal, France) LabCyte EPI-MODEL24 (J-TEC, Japan) epiCS® (Phenion, Germany) Skin+™ (Sterlab, France) 	Applicable to solids, semi-solids, liquids, waxes, and mixtures	Discriminates skin irritants (Cat 2) from substances not classified for skin irritation (No Cat). Materials that test positive should be tested for skin corrosion (bottom-up approach).
OECD TG 431: In Vitro Skin Corrosion: Reconstructed Human Epidermis (RhE) Test Method	Test substance is applied topically to a 3-dimensional RhE model. Corrosive chemicals are able to penetrate the tissue and are cytotoxic to cells in the underlying layers. Cell viability is measured using the vital dye MTT.	• EpiSkin™ (L'Oréal, France) • EpiDerm™ (MatTek, US) • SkinEthic™ (L'Oréal, France) • epiCS® (Phenion, Germany) • LabCyte EPI-MODEL24 (J-TEC, Japan)	Applicable to solids, semi-solids, liquids, waxes, and mixtures	Discriminates non-corrosive substances from corrosive substances (Cat 1) and allows subcategorisation into 1A or 1B and 1C together. Materials that test negative should be tested for skin irritation (top-down approach).
OECD TG 435: In Vitro Membrane Barrier Test Method for Skin Corrosion	Test substance is applied to the surface of an artificial membrane barrier designed to respond in a manner similar to skin <i>in vivo</i> . The time taken for the test substance to penetrate the barrier predicts corrosivity.	N/A	Applicable to solids, liquids, and emulsions. Aqueous chemicals with a pH in the range of 4.5 to 8.5 may not qualify for testing	Discriminates non-corrosive substances from corrosive substances and allows full subcategorisation into 1A, 1B, and 1C.

References

- OECD. New guidance document on an integrated approach on testing and assessment (IATA) for skin corrosion and irritation. Series on Testing and Assessment, No 203. July 2014 (available at www.oecd.org).
- European Chemicals Agency, 2017. Guidance on information requirements and chemical safety assessment. Chapter R.7a: Endpoint specific guidance. Version 6.0. See R.7.2.
- PETA Science Consortium International. In vitro methods for skin irritation and corrosion testing, Training tool (available at ThePSCI.eu/skin-in-vitro/).