**NOTE: For instructions on how to fill in the template see Word Doc with instructions.**

1. **Problem formulation**
2. **Hypothesised Mode of action Statement**
3. **Summary of data for use in Mode of Action Analysis**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| **Species** | **Route/Dose** | **Incidence** | **Comments** | **References** |
|  |  |  |  |  |
|  |  |  |  |  |

1. **Listing of key events identified for a specific Mode of Action**

|  |  |
| --- | --- |
| ***Key Event 1*** |  |
| ***Key Event 2*** |  |

1. **Bradford Hill Considerations for Weight of Evidence Analysis of available data/information for Mode of Action Analysis in experimental species**

**5a. Dose Response Relationships and Temporal Association**

**Species (name)**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
| ***Dose (units)*** | ***Key Event 1 (time observed and severity)*** | ***Key Event 2(time observed and severity)*** | ***Key Event 3(time observed and severity)*** | ***Key Event 4(time observed and severity)*** |
|  |  |  |  |  |
|  |  |  |  |  |

**5b. Consistency & Specificity – Biological Plausibility**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  | ***Key Event 1*** | ***Key Event 2*** | ***Key Event 3*** | ***Key Event 4*** |
| ***Consistency & Specificity*** |  |  |  |  |
| ***Biological Plausibility*** |  |  |  |  |

**5c. Qualitative and Quantitative human concordance**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  |  ***Qualitative Concordance*** |  |  |  |
| ***Key Event (name)*** | ***(Evidence in Experimental Species)*** | ***(Evidence in Humans)*** | ***Quantitative Species Concordance (experimental species and humans)*** | ***Quantitative Dose Response*** | ***Confidence/ Uncertainty*** |
| ***Key Event 1*** |  |  |  |  |  |
| ***Key Event 2*** |  |  |  |  |  |

1. **Other potential Modes of Action**

**Comparative Weight of Evidence Table**

|  |  |  |
| --- | --- | --- |
| ***Bradford Hill Considerations*** | ***Supporting Evidence*** | ***Potentially Inconsistent Evidence*** |
| ***Dose Response & Temporal Concordance*** |  |  |
| ***Consistency & Specificity*** |  |  |
| ***Biological Plausibility*** |  |  |

|  |  |  |
| --- | --- | --- |
| ***Bradford Hill Considerations*** | ***Supporting Evidence*** | ***Potentially Inconsistent Evidence*** |
| ***Dose Response & Temporal Concordance*** |  |  |
| ***Consistency & Specificity*** |  |  |
| ***Biological Plausibility*** |  |  |

1. **Uncertainties/Inconsistencies and Identification of Data Gaps**
2. **Conclusions in relation to problem formulation**