

Developer’s Handbook Version 2.5 (Release Notes)

These release notes for Developer’s Handbook version 2.5 outline significant changes to the Handbook relative to the previous version (2.0) released February 14, 2018. Line numbers listed below correspond to lines in the archived pdf copy of Developer’s Handbook version 2.5. The Developer’s Handbook is now accessible in its entirety from the AOP-Wiki (aopwiki.org). Guidance provided here is current with the most recent release of the AOP-Wiki (AOP-Wiki 2.5). This document will continue to be updated as additional features in the AOP-Wiki are implemented and feedback on the AOP framework is received.

Line	Description
1	Title changed to “Developer’s Handbook”
25	Contributing authors listed
103-124	A text-box added to define AOP-KB versus AOP-Wiki. The remainder of the text refers to AOP-Wiki only.
142-143	Figure 1 – generalized workflow of AOP development (organization of handbook sections) added
147-158	Developers are encouraged to review Annex 1 before starting development.
328-337	AOP title convention has been updated to “MIE leading to AO via distinctive KE” to provide greater specificity with growing numbers of closely related AOPs in the AOP-Wiki.
427-457	Status fields: No changes for release 2.5, but status fields will be re-vamped for the next edition of the handbook along with a new model for licensing of AOP-Wiki pages that will allow for all rights reserved AOP pages to address intellectual property concerns associated with open AOP development in the AOP-Wiki.
475-546	AOP Development Strategy – guidance for new information field implemented with release of AOP-Wiki v. 2.5. This field is intended to provide information on why an AOP was developed (context) and how the AOP was developed (strategy). This field replaces the former optional background section on the AOP page. All previous content from background sections will migrate to the “context” subheading under the AOP Development Strategy.
572-578	Network view figure (Figure 4) introduced.
581-602	Authors are no longer asked to identify stressors that may act on a KE as part of their Event descriptions. Prototypical Stressors, which are stressors for which responses at multiple key events other than the MIE have been well documented may be listed on the AOP page. Associated definition and desired contents of the prototypical stressors field is defined. Reduced emphasis on stressors responds to concern that AOP development was being conflated with stressor-specific hazard assessment.
694-782	Minor updates to text concerning biological domain of applicability of the KEs including a clearer distinction between empirical domain of applicability and plausible domain of applicability, which should both be addressed by the developer(s).
805-810	Added clarification that references should be cited in context, not just listed bibliography style.
866-871	Figure 6 was simplified. Removed an example that was creating confusion.
887-892	Figure 7 showing the “add relationship” dialog from the AOP-Wiki was added.
910-916	Figure 8, illustrating the ways that the domain of applicability of the upstream and downstream KEs linked via a KER are considered when defining the domain of applicability of each relationship was added.

997-1015	Guidance on how to fill the new “Evidence Collection Strategy” field on the relationship pages (new to AOP-Wiki 2.5) was added.
1082-1089	Added Figure 9 to illustrate dose concordance, temporal concordance, incidence concordance.
1101-1104	Added an example of an empirical evidence table (Table 5). The table also provides illustrative examples of dose, temporal, and incidence concordance at the KER level.
1159-1182	The section on Known Modulating Factors has been moved from a sub-section of the quantitative understanding of the KER to its own stand-alone section. Guidance on filling the new stand alone Known Modulating Factors section of the KER pages is provided.
1404-1408	The example of a table format (Table 7) for summarizing relative evidence supporting the essentiality of KEs was simplified compared to the previous handbook.
1503-1509	The previous example of an AOP-level concordance table was replaced with a more generic example illustrating the expected pattern (Table 8).
1524-1533	Explicit identification of known modulating factors was added to the Overall Assessment of the AOP. Added text provides guidance on how to fill the “Known Modulating Factors” section of the Overall Assessment of the AOP.
1569-1584	Added text on how to fill the “Consideration for Potential Applications of the AOP (optional) section. This was absent from the previous version.

Anticipated updates for version 2.6 and invitation for input:

Based on feedback received, we are aware of on-going confusion regarding the distinction between the assembly of weight of evidence on the key event relationship (KER) pages and weight of evidence evaluation (e.g., overall assessment of the AOP) on the AOP pages. Clarifications and improvements in the handbook related to these aspects of AOP development will be an emphasis for Developer’s Handbook version 2.6. If you have questions related to these topics, or ideas for how these sections could be improved, please contact aopwiki@googlegroups.com and/or post your comments, questions, or ideas in the [AOP Forum](#).