

## SOME AROMATIC AMINES AND RELATED COMPOUNDS

**VOLUME 127** 



This publication represents the views and expert opinions of an IARC Working Group on the Identification of Carcinogenic Hazards to Humans, which met remotely, 25 May-12 June 2020

LYON, FRANCE - 2021

IARC MONOGRAPHS
ON THE IDENTIFICATION
OF CARCINOGENIC HAZARDS
TO HUMANS

Table S1.9 Review of exposure assessment quality in cohort studies on exposure to aniline

| Reference                  | What was the study design?  | What methods were used for the assessment?   | What was the definition of the exposure?  | Was the exposure defined well?               | What<br>route of<br>exposure<br>was<br>assessed? | Could there be exposed subjects in the reference group?     | Was intensity<br>assessed<br>well? | Was the<br>duration<br>of<br>exposure<br>assessed<br>well? | Was<br>cumulative<br>exposure<br>assessed? | Was<br>exposure<br>assessed<br>before<br>outcome<br>being<br>ascertained? | What was<br>the timing<br>of exposure<br>relative to<br>the<br>outcome? | Are there any known carcinogen co-exposures in the industry?  |
|----------------------------|---|--|---|--|--|---|------------------------------------|--|--|---|---|---|
| Case et al. (1954)         | Cohort study of<br>United Kingdom<br>workers<br>manufacturing or<br>using dyestuff<br>intermediates                           | Use of<br>substance in<br>processing   | Companies reported all workers who were known to have had any contact with aniline, benzidine, 1-naphthylamine or 2-naphthylamine | No, very<br>unclear                          | Not<br>defined                                   | No, general<br>population<br>used as<br>reference           | No                                 | Not<br>assessed  | No, only<br>ever/never                     | No, not in<br>most recent<br>analysis                                     | Exposure<br>preceded<br>outcome<br>(death)                              | Yes. Co-exposure with magenta is reported in 9 of 13 cases who were exposed to aniline but were not exposed to α- or β-naphthylamine [1- or 2-naphthylamine] or benzidine |
| Ott &<br>Langner<br>(1983) | Cohort study of<br>US workers<br>manufacturing<br>organic dyes  | Process<br>review of raw<br>products and<br>intermediates<br>produced,<br>departments<br>with aniline<br>identified in<br>two<br>departments | Ever/never<br>worked in area<br>with potential<br>exposure to<br>aniline<br>(acetanilide and<br>indigo areas)                     | Moderately<br>well-<br>defined as<br>to area | Not<br>defined                                   | No, USA<br>white male<br>population<br>used as<br>reference | Not assessed                       | Yes,<br>total years<br>in exposed<br>department            | By<br>duration of<br>exposure<br>only      | No, after<br>cancers had<br>occurred                                      | Exposure<br>preceded<br>outcome   | Yes: asbestos,<br>arsenic and vinyl<br>chloride. Workers<br>may have been co-<br>exposed to <i>ortho</i> -<br>toluidine   |
| Sorahan<br>(2008)          | Cohort study,<br>extended follow-<br>up in United<br>Kingdom workers<br>manufacturing<br>chemicals for the<br>rubber industry | JEM based on<br>potential for<br>exposure to<br>aniline in<br>each<br>department   | Ever/never<br>worked in any of<br>6 departments<br>with potential<br>exposure to<br>aniline                                       | Moderately<br>well-<br>defined as<br>to area | Not<br>defined                                   | No, general<br>population<br>used as<br>reference           | Not assessed                       | Yes,<br>total years<br>in exposed<br>department            | By<br>duration of<br>exposure<br>only      | No, not in<br>most recent<br>analysis                                     | Exposure<br>preceded<br>outcome   | Yes. Co-exposure<br>to <i>ortho</i> -toluidine,<br>PBN, MBT   |

Table S1.9 Review of exposure assessment quality in cohort studies on exposure to aniline

| Reference  | What was the study design?   | What methods were used for the assessment?   | What was the definition of the exposure?                                  | Was the exposure defined well?  | What route of exposure was assessed?                          | Could<br>there be<br>exposed<br>subjects in<br>the<br>reference<br>group? | Was intensity<br>assessed<br>well?   | Was the<br>duration<br>of<br>exposure<br>assessed<br>well? | Was<br>cumulative<br>exposure<br>assessed?                             | Was<br>exposure<br>assessed<br>before<br>outcome<br>being<br>ascertained? | What was<br>the timing<br>of exposure<br>relative to<br>the<br>outcome? | Are there any known carcinogen co-exposures in the industry?   |
|--|--|--|---|---|---|---|--|--|--|---|---|--|
| Carreón et<br>al. (2014),<br>Hanley et<br>al. (2012) | Cohort study,<br>extended follow-<br>up in USA<br>workers<br>manufacturing<br>chemicals for the<br>rubber industry | Review of<br>work<br>histories,<br>process<br>records,<br>hygiene<br>measurements<br>to create a<br>JEM. Details<br>in Hanley et<br>al. (2012) | Four exposure categories for combined exposure and duration of employment | Well- defined as exposure to combined three agents, not well- defined for aniline alone | Personal<br>breathing<br>zone and<br>biological<br>monitoring | No, general<br>population<br>used as<br>reference                         | Combined exposure assessed as: PNE (probably not exposed); PEI (probably exposed low and irregularly /occasionally); PER (probably exposed low and regularly); DER (definitely exposed moderate/high and regularly). A ranking system was also used (see text) | Yes, days<br>in each job                                   | Yes, sum of all jobs with level of exposure by days worked in that job | No, not in most recent analysis   | Exposure preceded outcome   | Yes. Exposure was to any of ortho-toluidine, aniline, and nitrobenzene combined. No individual data for aniline provided. Patterns of exposure were very similar for both ortho- toluidine and aniline and co- exposures occurred in all departments |

JEM, job-exposure matrix; MBT, 2-mercaptobenzothiazole; PBN, phenyl-2-naphthylamine.

4

## References

- Alguacil J, Kauppinen T, Porta M, Partanen T, Malats N, Kogevinas M, et al.; PANKRAS II Study Group (2000). Risk of pancreatic cancer and occupational exposures in Spain. Ann Occup Hyg. 44(5):391–403. https://doi.org/10.1016/S0003-4878(99)00119-2 PMID:10930502
- Carreón T, Hein MJ, Hanley KW, Viet SM, Ruder AM (2014). Bladder cancer incidence among workers exposed to *o*-toluidine, aniline and nitrobenzene at a rubber chemical manufacturing plant. Occup Environ Med. 71(3):175–82. https://doi.org/10.1136/oemed-2013-101873 PMID:24368697
- Case RA, Hosker ME, McDonald DB, Pearson JT (1954). Tumours of the urinary bladder in workmen engaged in the manufacture and use of certain dyestuff intermediates in the British chemical industry. I. The role of aniline, benzidine, alpha-naphthylamine, and beta-naphthylamine. Br J Ind Med. 11(2):75–104. https://doi.org/10.1136/oem.11.2.75 PMID:13149741
- Feingold L, Savitz DA, John EM (1992). Use of a job-exposure matrix to evaluate parental occupation and childhood cancer. Cancer Causes Control. 3(2):161–9. https://doi.org/10.1007/BF00051656 PMID:1562706
- Hanley KW, Viet SM, Hein MJ, Carreón T, Ruder AM (2012). Exposure to *o*-toluidine, aniline, and nitrobenzene in a rubber chemical manufacturing plant: a retrospective exposure assessment update. J Occup Environ Hyg. 9(8):478–90. https://doi.org/10.1080/15459624.2012.693836 PMID:22708702
- Nizamova RS (1991). [Occupational hazards and bladder cancer]. Urol Nefrol (Mosk). (5):35–8. PMID:1836689 [Russian]
- Ott MG, Langner RR (1983). A mortality survey of men engaged in the manufacture of organic dyes. J Occup Med. 25(10):763–8. https://doi.org/10.1097/00043764-198310000-00018 PMID:6631562
- Preti G, Labows JN, Kostelc JG, Aldinger S, Daniele R (1988). Analysis of lung air from patients with bronchogenic carcinoma and controls using gas chromatography-mass spectrometry. J Chromatogr A. 432:1–11. https://doi.org/10.1016/S0378-4347(00)80627-1 PMID:3220881
- Sorahan T (2008). Bladder cancer risks in workers manufacturing chemicals for the rubber industry. Occup Med (Lond). 58(7):496–501. https://doi.org/10.1093/occmed/kqn104 PMID:18725381