Sub-statement on the potential risk(s) from exposure to microplastics: Inhalation route (Third draft)

This is a paper for discussion.

This does not represent the views of the Committee and should not be cited.

- 1. In 2019, as part of horizon scanning, the Committee on Toxicity of Chemicals in Food, Consumer Products and the Environment (COT) identified the potential risks from microplastics as a topic it should consider to inform Food Standards Agency (FSA) discussions on this (TOX/2019/08). Since then, several discussion papers have been presented to the COT (see Annex A1 and in 2021, the COT published an overarching statement on the potential risks from exposure to microplastics (COT Statement 2021/02). This document was first presented to the COT in May 2022 (COT Minutes May 2022) and a follow up draft in December 2022 (COT Minutes December 2022). This provided a high-level overview of the current state of knowledge, data gaps and research requirements with regards to this topic.
- 2. There is evidence for the presence of plastic particles in both indoor and outdoor air and thus inhalation is a possible route of exposure (Gasperi et al., 2018; Domenech & Marcos, 2021).
- 3. The purpose of this sub-statement is to provide supplementary material to the overarching statement (COT Statement 2021/02) and to consider in detail the potential toxicological risks of exposure from microplastics *via* inhalation. It is based on current available literature and on data from internal tools at the UK FSA (these internal tools include: a literature search application and signal prioritising dashboards). This would help inform overall exposure of microplastics and also any further toxicological information. As Members will recall, particles arising from tyre wear were considered outside the scope of this assessment, but

some information has been included to provide context.

4. When the Committee discussed this paper at the December 2022 meeting, it was suggested that the research priorities be ranked. It was also noted that the conclusions in this report are similar to that of the 2022 WHO review and the Chief Medical Officer annual report and these should be referenced in this document. The Committee decided that tyre wear should be included as a brief overview and to highlight the previous review from 2020. Colleagues at Committee on the Medical Effects of Air Pollution (COMEAP) have provided comments on this draft of the Statement and a Table has been added to give more weight to the study from Jenner et al. 2022 providing the lengths of the particles measured. Some restructuring of the paper has been carried out at the request of the COT and additional text has been highlighted in yellow.

Questions for the Committee

- 5. The Committee are asked to consider the following questions:
- a) Are the Committee content with the updated version of the sub-statement considering all the suggestions by Members?
- b) As requested by the Members, the research priorities have been ranked. Do the Committee agree with the ranking provided?
- c) Does the Committee have any further comments?
- d) Can the sub statement be cleared by Chairs action?

Secretariat

July 2023

Annex A to TOX/2023/38

Background

1. In 2019, as part of horizon scanning, the Committee on Toxicity of Chemicals in Food, Consumer Products and the Environment (COT) identified the potential risks from microplastics as a topic it should consider to inform UK Food Standards Agency (FSA) discussions on this area (TOX/2019/08). Since then,

several discussion papers have been presented to the COT (see <u>Annex A1</u>) and in 2021, the COT published an overarching statement on the potential risks from exposure to microplastics (COT Statement 2021/02).

- 2. This document provided a high-level overview of the current state of knowledge, data gaps and research requirements with regards to this topic. This was followed by a sub-statement considering oral exposure to microplastics in more detail.
- 3. The COT previously noted that there is little data on the effects of microplastics on mammals (including humans) whether taken in orally or via inhalation. The majority of microplastics (>90%) are excreted from the body but small amounts may remain in the gut (gastrointestinal tract; GIT) or move from the GIT into organs or tissues due to endocytosis by M cells or paracellular persorption. No epidemiological or controlled dose studies that evaluated the effects of orally ingested microplastics in humans were identified and there is a similar lack of information on inhaled microplastics.
- 4. Although exposure to airborne microplastics can arise from a wide range of environmental sources (see paragraph 77) there is still limited information regarding the concentrations of airborne microplastics.
- 5. In 2022, England's Chief Medical Officer Professor Chris Whitty published a report on indoor and outdoor air pollution which included comments on microplastics. In the report it is noted that microplastics are in the air unintentionally by stating:

"The airborne transport and inhalation of microplastics is an example showing how unintended air quality consequences might possibly arise far downstream from the public use of an originally safe synthetic product. Too great a focus on only meeting existing air quality standards and regulations, without considering how atmospheric composition may change with society and technology more broadly, may lead to problems that could have been intercepted earlier with greater non-targeted surveillance and horizon-scanning. A clear evidence gap exists between the extensive regulatory efforts placed on monitoring existing regulated air pollutants and research studies of emerging atmospheric composition, the latter being rarely systematic or long-term in nature" (CMO, 2022).

6. The fate and dispersion of microplastics in outdoor environments is dependent on several factors (see paragraph 79).

- 7. Atmospheric deposition of microplastic particles (MPPs) onto food prior to consumption must also be considered as a potential source of exposure. For example, Catarino et al., (2018) compared the potential exposure of humans to household dust fibres during a meal with the amounts of MPPs present in edible mussels from Scottish waters, showing that exposure was considerably higher from the household source (see paragraph 82).
- 8. An American study (Cox et al., 2019) estimated daily consumption and inhalation to be 142 MPPs and 170 MPPs in adult males, respectively. For adult females, the estimated values are 126 MPPs and 132 MPPs, respectively. Based on these values, an annual estimated exposure of ~120,00 and ~98,000 MPPs annually was calculated in male and female adults, respectively (see paragraph 83 below).
- 9. The deposition of inhaled microplastics within the lung is dependent on the particle's physicochemical properties, as well as the subject's physiology and lung anatomy shown in paragraph 64.
- 10. Inhalation of microplastics could result in toxicity due either to the particles (i.e. physical effect) or their leachates (i.e. chemical effect). The mechanisms of inhaled particle injury are covered in paragraph 85. With regard to the available inhalation studies in laboratory animals, Environment and Climate Change Canada and Health Canada (ECCC and HC) in their review of the scientific literature noted that no dose-response relationship had been observed in mortality, survival time, behaviour, clinical observations, or tumour incidence from inhalation exposures (ECCC and HC 2020).
- 11. The COT previously reviewed risk assessments carried out by various groups such as the European Tyre and Road Wear Platform; Tyre Industry Project (Jekel, 2019), Joint Research Centre (Grigoratos & Martini, 2014), Defra (AQEG, 2019), Health and Safety Executive (RUBIAC, 2007; HSE, 2011), Committee on Medical Effects of Air Pollutants (COMEAP, 2015; 2020), WHO (WHO, 2013), National Institute for Public Health and the Environment (Verschoor et al., 2016), and ECHA (ECHA, 2017).
- 12. The COT concluded that the literature data on exposure to particles from tyre wear would need separate consideration from microplastic exposure from food, since the particles are chemically quite different in their polymeric nature. The COT considered that inhalation was likely to be the most significant route of exposure to TRWPs (tyre and road wear particles). Risk assessments of such materials was considered outside of the scope of the current exercise;

however some information has been included to provide context. The COT was of the opinion that uptake by plants, with subsequent dietary exposure, was unlikely to be a major route of exposure but it would need further studies to review and confirm this.

Scope and purpose

- 13. As there is evidence for the presence of plastic particles in both indoor and outdoor air, inhalation is a possible route of exposure (Gasperi et al., 2018; Domenech & Marcos, 2021).
- 14. The purpose of this sub-statement is to provide supplementary material to the overarching statement (<u>COT Statement 2021/02</u>) and to consider in detail the potential toxicological risks of exposure from microplastics *via* the inhalation route. It is based on the currently available literature and data from internal tools at the FSA (these include: a literature search application and signal prioritising dashboards).

Microplastics

- 15. Currently there is no internationally agreed definition of a microplastic, however, publications by Verschoor (2015) and Hartmann et al., (2015) have proposed criteria that could be included in the definition of microplastics. In Europe, the European Chemicals Agency (ECHA) has proposed a regulatory definition for a microplastic under the Registration, Evaluation, Authorisation and Restriction of Chemicals (REACH) Regulation (ECHA, 2019). In the US, the California Water Boards also recently published a proposed definition of microplastics in drinking water in March 2020.
- 16. Verschoor (2015) included 5 major properties that could be considered including chemical composition, physical state, particle size, solubility in water and degradability. Similarly, Hartmann et al., (2015) proposed seven criteria; chemical composition, solid state, solubility, size, shape and structure, colour and origin (i.e. primary or secondary particles; also known as pristine and aged), as discussed in the following paragraphs.
- 17. In Europe, the definition of a microplastic proposed by ECHA is a "material consisting of solid polymer-containing particles, to which additives or other substance(s) may have been added, and where $\geq 1\%$ w/w have (i) all dimensions $1 \text{ nm} \leq x \leq 5 \text{ mm}$ or (ii) for fibres, a length of $3 \text{ nm} \leq x \leq 15 \text{ mm}$ and length to diameter ratio of >3. Polymers that occur in nature that have not been

chemically modified (other than by hydrolysis) are excluded, as are polymers that are (bio)degradable." (ECHA, 2019).

- 18. The current definition of microplastics in drinking water adopted by the California Water Boards is: "Microplastics in drinking water are defined as solid polymeric materials to which chemical additives or other substances may have been added, which are particles which have at least two dimensions that are greater than 1 and less than 5,000 μ m. Polymers that are derived in nature that have not been chemically modified (other than by hydrolysis) are excluded." (California Water Boards, 2020).
- 19. As noted above, the definitions of microplastics are broad. Therefore, for the purposes of this document, the COT has adopted a working definition that microplastics are defined as synthetic particles or heavily modified natural particles with a high polymer content that are submicron in size (0.1 to 5,000 μ m or micrometres). Plastics that are below this size range are classed as nanoplastics (i.e. 1 nm to 0.1 μ m).
- 20. The Committee also noted that microplastic particles that are present in the environment are not stable in size, meaning that as the duration of the degradation process lengthens, the particle size continues to decrease due to fragmentation and erosion/weathering.

Nanoplastics

- Nanoplastics have been defined as a material with any external dimension in the nanoscale or having internal structure or surface structure in the nanoscale (1 nm to 0.1 μ m) (EFSA, 2016, European Commission, 2011). Nanoparticle is a general term based on the physical properties for a variety of chemical compositions. There is currently no further proposed definition.
- 22. A number of authoritative bodies have assessed the risks of nanomaterials and provided guidance on their assessment, which could also apply to nanoplastics. For example, the European Food Safety Authority (EFSA) Scientific Committee published an opinion on the potential risks arising from nanoscience and nanotechnologies on food and feed safety in 2009 (EFSA, 2009). This opinion did not provide any definitions; however, it was stated that the term nanoscale refers to a dimension of the order of 100 nm and below. Engineered nanomaterial was described as any material that is deliberately created such that it is composed of discrete functional and structural parts, either internally or at

the surface, many of which will have one or more dimensions of the order of 100 nm or less.

- 23. The EFSA Scientific Committee recommended that the addition of other metrics (i.e. specific surface area which is independent of the agglomeration status of particles) should be included into the current definition of nanoscale materials (EFSA, 2009).
- 24. In 2011, EFSA published a guidance document on how EFSA's Panels should assess potential risks related to certain food-related uses of nanotechnology. New guidance on assessing the safety for humans and animals of nanoscience and nanotechnology applications in the food and feed chain was published in 2018 (EFSA, 2018).
- 25. The EFSA 2018 guidance is applicable to:
 - a material that meets the criteria for an engineered nanomaterial, as outlined in Novel Food Regulation (EU) No 2015/2283 and Regulation (EU) No 1169/2011 (i.e. have particle sizes in the defined nanoscale; 1-100 nm),
 - a material that contains particles having a size above 100 nm which could retain properties that are characteristic of the nanoscale (not further elaborated),
 - a material that is not engineered as nanomaterial but contains a fraction of particles (50% in the number-size distribution) with one or more external dimensions in the size range 1-100 nm or less,
 - a nanomaterial having the same elemental composition but that occurs in different morphological shapes, sizes, crystalline forms and/or surface properties, and;
 - a nanoscale entity that is made of natural materials.
- 26. In July 2020, EFSA held a public consultation on its draft "Guidance on technical requirements for regulated food and feed product applications to establish the presence of small particles including nanoparticles". The draft guidance outlines appraisal criteria grouped in three sections, to confirm whether or not the conventional risk assessment should be complemented with nanospecific considerations.
- 27. The first group of criteria addresses solubility and dissolution rate as key physicochemical properties to assess whether consumers will be exposed to particles. The second group establishes the information requirements for assessing whether the conventional material consists of small particles or

contains a fraction thereof, and its characterisation. The third group describes the information to be presented for existing safety studies to demonstrate that the fraction of small particles, including those at the nanoscale, has been properly evaluated. Post-finalisation, this guidance was to completement the EFSA 2018 guidance (as described above) (EFSA, 2020).

28. The definitions of nanomaterials above (paragraphs 21-24) are based on EFSA Guidance, but their guidance for the risk assessment of nanomaterials could also apply to nanoplastics.

Types of microplastics

- 29. Microplastics can be divided into two major types. Firstly, those that are deliberately manufactured to be in the size range of 0.1 to 5,000 µm which are known as primary microplastics, and are intentionally used in personal care products (for example, microbeads) or for various industrial applications. Secondary microplastics can be formed in the environment due to fragmentation of larger pieces of plastic caused by a culmination of physical, biological and photochemical degradation. Secondary microplastics have been termed microplastic particles (MPPs). MPPs can be further degraded to form nanoplastics, as defined above.
- 30. Besides the types of microplastics mentioned above, there is some debate within the scientific field as to whether rubber tyre particles should be considered microplastics. Tyres were initially made of natural rubber from the Brazilian rubber tree (Hevea brasiliensis). Currently, tyres are produced from a mixture of natural and synthetic materials. Synthetic rubbers are made from petroleum products and are functionalised with the addition of sulfur (1-4%), zinc oxide (1%), carbon black/silica (22-40%) and oil (Kole et al., 2017).
- 31. Car tyres release wear particles through mechanical abrasion, resulting from contact between the road surface and the tyre. The amount and particle size are dependent on several factors such as climate (temperature), composition and structure of the tyre, tyre age, road surface, driving speed, vehicle characteristics and style, and nature of the contact. As such, tyre wear particles could be described as another environmental source of microplastics, depending on the presence of synthetic materials in their composition (Baensch-Baltruschat et al., 2020; Kole et al., 2017).
- 32. Tyre wear and microplastics were previously reviewed by the COT (
 Annex B1). The COT summarised that "tyres contain a wide range of chemicals,

the bulk of tyre tread is composed of a variety of rubbers, including natural rubber co-polymers, poly-butadiene rubber, styrene-butadiene rubber, nitride rubber, neoprene rubber, isoprene rubber, and polysulphide rubber. The interaction of tyres and pavement alters both the chemical composition and characteristics of particles generated compared to the original tyre tread due to heat and friction, as well as incorporation of materials such as environmental "dust", brakes, fuels and the atmosphere, as well as roadway particles.

- 33. Human exposure to chemicals leached from tyres, shredded tyres, and tyre wear material can occur by dermal exposure from environmental sources and ingestion of contaminated materials, as well as inhalation of airborne particulate matter derived from tyre wear material.
- 34. Challenges associated with evaluating risk from exposure becomes complex when considering other factors such as the effects of weathering and ageing of tyre materials, the effects of temperature, pavement types and driving style. All in all, this will result in the generation of various chemicals with significantly different biological and toxicological effects and potencies".
- 35. In the recent CMO report it stated "In the longer term, measuring tyre pollution accurately is also needed to assess the full impact of tyres on air and microplastic pollution, and reduce toxicity" (CMO, 2022).

Analytical detection methodologies

- 36. From the literature, the detection methods described for microplastics include one or more of the following steps: sample collection and removal of biogenic matter, detection and quantification/enumeration and, the characterisation of the plastic (i.e. its chemical composition or polymer type) (Nguyen et al., 2019; Kwon, et al., 2020). It is important to note that during all these steps, precautions to avoid contamination from particles in the air, or with fibres from clothing, equipment or the reagents used, should be optimised (see Figure 1).
- 37. As seen in Figure 1, the majority of biological samples have been taken from aquatic species. The pre-separation method is dissection which recovers MPPs >500 μ m, followed by separation methods including density separation, digestion using enzymes and various chemical compounds and filtration techniques. The analytical method is split between three categories: visual microscopic analysis, vibration spectroscopy (e.g. FTIR and Raman spectroscopy) and mass spectroscopy, the last of which is also suitable for the characterisation,

quantification and identification of nanoplastics.

- 38. Personal communications with National Reference Laboratories in the UK revealed that organic contaminant analysis usually applies only to the edible portions of food and that contaminants adsorbed to microplastics are generally not accounted for when measuring residues in foodstuff. Although it was noted that this would be method dependent rather than an intrinsic property (Gilbert, personal communication, 2020).
- 39. For example, when analysing fish the head, digestive tract, offal and bones are removed before analysis. It is expected that the majority of microplastics will be in the GIT; therefore, any contaminants associated with them would not directly contribute to measured contaminant levels. However, depending on the nature of the microplastic contaminants, some may be desorbed in the stomach and may contribute to the measured concentration in the edible parts of the fish.
- Figure 1. Provides an overview of the methodologies utilised in the separation and analysis of microplastics and nanoplastics in complex environmental samples including: biological samples (fish, mussels and plankton), air samples (indoor and outdoor, synthetic textile fibres), wastewater samples (influent, effluent and sludge), water samples (drinking water, sea water and fresh water) and sediment samples (adapted from Nguyen et al., 2019). Abbreviations: H2O2 = hydrogen peroxide; HNO3 = nitric acid; KOH = potassium hydroxide; HCl = hydrochloric acid; SEM = Scanning electron microscope; TEM = Transmission electron microscopy; FT-IR = Fourier-transform infrared; TDS-GC-MS = Thermodesorption gas chromatography-mass spectrometry; py-GC-MS = Pyrolysis gas chromatography-mass spectrometry (Image taken from COT Microplastics Overarching Statement 2021).
- 40. Currently there are only limited analytical methods available to detect and quantify the presence NMPs in various matrices. These include FT-IR, Nile Red staining techniques, Micro-Raman spectroscopy, quantitative 1H nuclear magnetic resonance spectroscopy (qNMR) (Peez et al., 2019) and mass-spectroscopy; however, each of these methods has its own associated limitations (Nguyen et al., 2019).
- 41. Additionally, there are neither standardised testing protocols for different matrices (i.e. air, soil, food and water), nor standard refence materials for the analysis, characterization and quantification of micro and nanoplastics. No

single technique is suitable for all plastic types or for all particle sizes or shapes. Therefore, the use of either a suite of methods or generation of new techniques will be necessary.

- 42. Comparison and replication of studies can be difficult due to differences in sampling, extraction, purification and analytical methods for enumerating and characterising microplastics. These methods are not yet standardized and have not been subject to interlaboratory validation. Contamination with airborne microplastics or cross contamination of samples can also occur, so suitable control samples may be difficult to obtain.
- 43. Most studies have performed tests on pristine particles; however, this may not be representative of what is present in the environment (i.e. the particles have not been subject to environmental degradation). Therefore, it is important to consider the variability among samples and batches of pristine particles when comparing studies of the same polymer type.

Physicochemical properties

- 44. There are four morphological and chemical characteristics of microplastics, i.e. physicochemical properties, which influence their potential hazards. These are:
- i). Physical (e.g. bulk, fibres in the lung or those which could lead to gut blockage, as observed in aquatic and avian species);
- ii). Chemical composition (unbound monomers, additives, sorbed chemicals from the environment e.g. persistent organic pollutants and metals);
- iii). Metabolism or degradation to form monomers or other derivatives, some of which could be chemically reactive (e.g. isocyanates from polyurethane) and;
- iv). The presence of biofilms (attachment and colonisation of microorganisms on the plastics).
- 45. Due to the small size of some nano-/microplastics, (NMPs) uptake across the gastrointestinal tract (GIT) and uptake into internal tissues is possible and thus they may have both local and systemic effects. Particles 50 μ m in size can be absorbed from the gut via inter-cellular gaps and by phagocytic and endocytic pathways but only those of 1-2 μ m in size are able to cross the cell membranes of internal organs.

Physical properties

- 46. NMPs can differ in their physical chemical properties (shape, size, density, surface charge, etc). The consideration of physical properties during hazard and/or risk assessment of plastic particles is important because the interactions of NMPs with biological systems can vary with differences in their size and shape (Nel et al., 2009), even when they have the same chemical composition.
- 47. The physical properties and morphologies of tyre materials can also vary under different sampling conditions. Those collected from road runoff and shredded tyres have elongated shapes, whilst samples generated from road simulator systems in laboratories range between jagged, droplets, granules, warped, porous, irregular, and near spherical in shape (Wagner et al., 2018). A review by Kole et al., (2017) revealed that the size distribution range of tyre wear and tear particles, could be from 6-350,000 nm. This wide size distribution range was attributed to several factors including the use of difference size metrics (e.g. particle mass versus particle numbers), analytical difficulties in separating tyre from road particles, and the large variation in experimental conditions and analytical equipment.

Chemical Properties

- 48. A particle's chemical properties such as charge or zeta potential (when particles are immersed in a conducting liquid such as water) are dependent on its chemical composition.
- 49. A particle's properties can also be influenced and changed by its surface chemistry. Each particle could have its own unique corona consisting of proteins adsorbed from plasma and/or intra/extracellular fluid, adsorbed chemicals from the environment or microbiological organisms (see Figure 2.).
- Figure 2. (a) Comparison of the protein, environmental, and ecocoronas. Formed within organisms at locations of high protein content, the term protein corona has been used to describe the binding of proteins to ENM (engineered nanomaterials) surfaces, but also incorporates lipids, metabolites (typically 1000 Da which are either reactants, intermediaries or products of enzymatic processes), and

other biomolecules. To date, the term environmental corona has described a corona formed in aquatic environments with high concentrations of NOM, including humic substances. By contrast, the eco-corona incorporates features of both the protein and environmental coronas, where the balance of proteins and other molecules varies. (b) The evolution of the protein corona concept, adapted from Hadjidemetriou and Kostarelos. Studies of protein adsorption to surfaces and particles dates back to at least the 1960s. The term protein corona was first coined in 2007. Protein corona studies developed with mass spectroscopy-based proteomics to aid identification of the proteins bound at the surface of ENMs and explore the role of surface curvature in altering protein structure and function relative to macroscale surfaces. Protein corona studies evolved in parallel with those on the environmental corona, but the characterization techniques and goals for each area remained separate, with environmental corona focusing mainly on the dispersion stabilization provided by NOM (natural organic material). The environmental dimensions of the protein corona began to appear later, as the concept of the eco-corona and its role in (nano)ecotoxicity emerged. Both the eco-corona and biomolecular corona embrace the diversity of molecules in solution with the goal of understanding and controlling downstream biological responses to nano-enabled technologies. (Image obtained from Wheeler et al. 2021).

50. The physicochemical properties of micro and nanoplastics can change over their life cycle and can also affect each other. For example, physical degradation resulting in the formation of nano-sized plastic particles and/or plastic particles with different shapes can generate a higher number of particles and thus gives rise to a larger total surface area and higher particle number which in turn affects the concentration. The weathering process can change the surface chemistry and size of microplastics, and chemical migration from the MPPs into the surrounding medium results in altered stability which in turn changes the physical degradation processes (Wheeler et al. 2021).

Toxicity

51. The COT have previously reviewed the human data on the toxicity of microplastics via inhalation (TOX/2019/62). The available toxicity data in humans was based on studies of occupational exposure. In 1975, Pimental et al., found that seven patients who had been exposed to synthetic fibres presented with different manifestations of bronchopulmonary disease. This was attributed in part

to the dose and concentration of fibre exposure and also the length of service in the industry.

- 52. In Hillerdal et al., (1988), a case study of three female workers in the synthetic textiles industry reported fibrotic areas of the lung with multiple foreign bodies believed to be inhaled fibres which caused inflammation. Although the size of the fibres was unreported, from other literature it was known that the length, biopersistence and width of inhaled fibres was related to their potential pathogenicity (Davis et al. 1986).
- 53. Pauly et al., (1998) reported a comparison study of inhaled cellulosic and plastic (polyester) fibres which were found in human lung tissue with samples being taken from different pulmonary sites, indicating the inhaled fibres were distributed throughout the lung. Authors suggested a possible link that there was a correlation between the inhalation of these plastic fibres with lung cancer, whereas the cellulosic fibres were relatively inert. Other studies have expanded on this. In general, there is no lung cancer risk.
- 54. In their discussion of <u>TOX/2019/62</u> the COT considered that the assessment of microplastics exposure via inhalation could be easier in comparison to the assessment of oral exposure given the availability of occupational data from the synthetic textile industry, however, the context should be considered. The Committee further noted that microplastic concentrations present in food and water were thought to be lower in comparison to airborne exposure.
- According to Panko et al., (2019), particulate matter (PM) from tyre abrasion may represent between 0.8 and 8.5 % mass fraction of PM10 and 1 to 10% of PM2.5 in the air. However, it is unknown what percentage of the PM2.5 burden consists of microplastics (Zhang et al., 2020).
- The toxicological properties of microplastics may differ to that of PM2.5 (or other pollutants) due to the additives present and their particular characteristics such as morphology and chemical composition (Zhang et al., 2020).
- 57. In 2020, the Committee on the Medical Effects of Air Pollutants (COMEAP) considered that the evidence on non-exhaust particles (road surface wear, re-suspended road dust, brake and tyre wear) from road transport and associated health effects should be re-evaluated. COMEAP concluded that, as a whole, the body of published work is small and did not provide a compelling

narrative of adverse health effects of exposure to non-exhaust particles. However, as there was strong evidence that exposure to particulate pollutants in ambient air is harmful to health, some health risk associated with exposure to non-exhaust particles was likely.

58. COMEAP concluded that the available evidence is not very informative about which components or sources of particulate air pollution are particularly harmful to health and that evidence relating to non-exhaust emissions from traffic, is limited (COMEAP, 2020; 2022)

Inhalation studies (2020-March 2023)

In vitro

- 59. When the <u>Overarching Statement</u> by the COT was published, it included studies up to 2020 and therefore this statement concentrates on those studies published from 2020 to the March 2023.
- 60. Dong et al., (2020) assessed the pulmonary toxicity of polystyrene microplastics *in vitro* using BEAS-2B lung cells to determine the cytotoxic and inflammatory effects. The polystyrene MPs decreased 1 α -antitrypsin levels and transepithelial electrical resistance by depleted zonula occludens proteins: these are a type of scaffolding protein. The study results indicated that low levels of polystyrene MPs cause disruption to the protective pulmonary barrier and high levels may have influenced an adverse effect on human lung health. This is based on testing in a single cell line which is not representative of in vivo exposure, however there are now 3D models available which can provide a more suitable model for examining the effects of airborne microplastics (Winkler et al. 2022).

In vivo

61. There is limited research on *in vivo* exposure to airborne microplastics. Lim et al., 2021 used a modified version of the OECD guideline (TG 412) 28-day inhalation toxicity study using a whole-body system. Sprague-Dawley rats were exposed to three different concentrations (0.75, 1.50 and 3.00 x 10^5 particles/cm 3) of polystyrene NMPs. There was no definitive link between concentration at 14 days exposure, and observed alterations to the physiological, serum biochemical and haematological parameters or markers of respiratory function. However, there was a concentration-dependent response associated with increased expression of TGF-β and TNF-α inflammatory proteins. These results suggest that

sustained exposure to higher concentrations of NMPs may result in alterations at the molecular level, thus a risk to health from inhalation of polystyrene micro/nanoplastics. However, caution must be given to the weighting of this study as rat lungs differ greatly from human lungs and therefore may not give a realistic representation of exposure in the human lungs. In addition, it is important to distinguish between normal acute particle clearance mechanisms and more persistence, potentially pathophysiological, responses.

Microbiological colonisation of micro- and nanoplastics and their significance to the food chain

- 62. The UK FSA has completed a critical literature review on the microbiological colonisation of micro- and nanoplastics and their significance to the food chain (FS307021); the project was completed and published in early 2022 (Walker et al. 2022). The critical review presented an overview of NMPs in the environment, the interaction of NMPs and micro-organisms, the identification of key pathways by which these microbiologically contaminated materials could enter the food chain from environmental sources (e.g. water, soil, and air), and the risk(s) that these might pose to the consumer (FSA, 2020). This critical review found that the amount and quality of evidence did not enable any conclusions on possible human health impacts to be reached:
 - The scale of NMP in the environment has been shown to be a large and growing problem with the majority of studies focusing on marine and aquatic systems.
 - Association between NMP and pathogenic organisms (such as Vibrio spp.,
 Pseudoalteromonas spp. and Aeromonas salmonicida) has been
 demonstrated in research but the amount of data is low and there is a lack of
 data comparing association with similarly sized, non-plastic particles (such
 as sediments).
 - There is a general lack of papers on potential pathways by which NMP can enter the food chain. From limited data available, it can be inferred that pathogens in association with NMP could potentially reach the human food chain. However, the consequence of this and comparison to non-plastic vectors have been insufficiently studied to create a clear picture of the risk posed to public health.
 - Research quantifying the dietary NMP intake is not available for most food and drink categories. From the identified studies involving foods, most have been carried out on fish and shellfish, with a very few considering food at retail sale or point of consumption.

• Currently, it is difficult to assess the impact of microbiological contaminated NMPs in the food chain due to significant data gaps and lack of research in this area.

Toxicokinetics

- 63. As discussed in <u>COT Statement Number 2021/02</u>, the toxicity of microplastics is dependent on a number of factors including size, morphology, chemical composition, additive leaching density and surface functionalization.
- 64. The deposition of inhaled microplastics in the lung is dependent on the particle's physicochemical properties, as well as the subject's physiology and lung anatomy. Deposition in the upper airways occurs by impaction, while in the small airways it occurs by sedimentation. Fibres have higher potential than spherical particles for penetration due to their high aspect ratio (Donaldson & Tran, 2002). Clearance relies on mechanical processes (e.g. mucociliary clearance where the mucous progresses towards the pharynx caused by the beating of cilia), alveolar macrophage phagocytosis and migration, and by lymphatic transport which can result in deposition in the GIT.
- 65. Surface functional groups may affect the adsorption of organic contaminants and heavy metals leading to different mobility and toxicity (Kim et al., 2017; Sun et al., 2020; Yao et al., 2022). However, the MP vector effect is usually framed as 'complex', 'under debate' or 'controversial' (Koelmans et al., 2022b).
- 66. Deposition of respirable fibres occurs in the lung as a function of its aerodynamic diameter of the particle, whereas non-respirable fibres are often inhaled through the nasal passage, but are then caught in mucus and swallowed thus creating a secondary exposure via the gastrointestinal tract. There are four main mechanisms of deposition in the lung: impaction, sedimentation, diffusion and interception (Darquenne, 2006).
- 67. Inertial impaction occurs for particles with a diameter $> 5 \mu m$ or for those with excessive momentum. As a particle travels through the airways, it remains on the same trajectory. If the air flow changes direction, the particle will remain on its existing pathway, deviating from the changed airflow and impacting on to the surface of the airways. Inertial impaction occurs in both the upper respiratory tract and the conducting zone only.

- 68. For particles with a diameter between 0.5 to 5 μ m, the main deposition mechanism is sedimentation. This occurs mainly in the bronchi and bronchioles; when air resistance and gravity overcome the buoyancy of the particle causing it to settle on the surface of the lung.
- 69. Airborne particles are in constant random Brownian motion, due to collisions with the gas molecules. This results in random, omnidirectional particle movement, known as diffusion. This occurs only with smaller particles, typically 0.5 μ m. Occasionally particles will collide with the cell surface, causing them to settle there. Diffusion occurs mainly in the small airways and alveoli, although the particles can also deposit in the upper airways by this mechanism particularly when their diameter is 0.01 μ m (Tsuda et al., 2013).
- 70. Interception occurs when fibres with a large ratio between their length and diameter travel so close to the surface of the lung that they make contact. Deposition by interception increases with the length of the fibre. The area of deposition in the lung is dependent on the aspect ratio of the fibre, but can also arise due to changes in airflow.
- 71. Clearance mechanisms for inhaled MPs $> 5 \mu m$, are likely to occur via the mucociliary escalator, where the particles are either expelled from the body via sneezing or coughing, or they are swallowed and taken into the gastrointestinal tract.
- 72. Some particles bypass the mucociliary clearance and travel deeper into the lung where phagocytosis occurs. Macrophage phagocytosis can break down particles 20 μ m by either dissolution or degradation but this is dependent on the particle biopersistence. If a particle is > 20 μ m in length, macrophages will not be able to fully engulf the particle, resulting in frustrated phagocytosis (Donaldson et al., 2010). This state causes an increased recruitment of macrophages, which can result in the phenomenon known as an "oxidative burst" occurring where inflammatory mediators and oxidants are released in high concentration, potentially leading towards the onset of lung inflammation and fibrosis (Donaldson et al., 2010; Gasperi et al., 2018). Inflammation induces cell proliferation and secondary genotoxicity due to the continuous formation of reactive oxygen species (ROS), resulting in oxidative stress, but this is not always the case.
- 73. It is believed that once particles reach the pleura then they may reach the pleural space, however it is currently unknown how this particle migration occurs. Once particles reach the pleura, they may then travel to the lymphatic

system which also helps clear phagocytic cells (Donaldson, et al., 2010; Enyoh et al., 2019). Translocation to secondary tissues and organs may then occur (Fournier et al., 2020; Wright and Kelly, 2017).

Exposure

- 74. Microplastics are present in the indoor and outdoor environment. The MPs can result from textiles, furniture, toys, electric cables and cleaning agents, construction material and litter.
- 75. The data available on outdoor exposure is limited. However, studies have shown than when comparing inhalation and ingestion routes indoors, microplastic exposure via ingestion is minimal in comparison to inhalation.

Outdoor exposure from air

- 76. Microplastics found outdoors are more likely to fracture due to weathering in comparison to the microplastics present indoors. However, data on the levels and types of microplastics in the air compared to other media are limited (Ageel et al., 2021).
- 77. Environmental exposure to airborne microplastics occurs from a wide range of sources with synthetic textiles and the erosion of synthetic rubber tyres being the most frequently reported in the literature. Resuspended city dust which contains a fraction of settled synthetic fibres/rubber tyre wear is a secondary source of airborne microplastics. Wind transfer is estimated to be responsible for 7% of the ocean's contamination (Boucher & Friot, 2017).
- 78. There is limited information regarding the concentrations of airborne microplastics, however, the Dris et al., (2016, 2017) studies carried out in Greater Paris found average outdoor deposition rates of 53 and 110 particles/m²/day. Data for Central London on outdoor deposition rates of microplastics have also been reported, and these range from 575-1,008 fibres/m²/day (Wright et al., 2020). These numbers are affected by climate conditions and seasonality and are also affected by the sampling and analytical methodologies used.
- **79**. The fate and dispersion of microplastics in outdoor environments are dependent on several factors. These include the vertical pollution concentration gradient where there are higher concentrations near the ground due to deposition and settling, wind speed and direction, land topography, precipitation and temperature. Concentrations of airborne microplastics in outdoor air are expected

to be low, due to dilution. O'Brien et al. (2023) noted in their review that the concentration of microplastics in outdoor ambient air ranges between 1 and >1000 microplastics/m³, while the outdoor deposition concentrations ranged between 0.5 and 1357 microplastics/m²/day.

- 80. The indoor behaviour of airborne microplastics is dependent on factors including room partition, ventilation and airflow.
- 81. Atmospheric deposition of MPPs onto food prior to consumption should also be considered as a potential source of exposure.
- As stated previously, Catarino et al., (2018) compared the potential 82. exposure of humans to household dust fibres during a meal using information on the amounts of MPPs present in edible mussels from Scottish waters collected throughout 2015. The mean number of MPPs in Modiolus modiolus was 0.086/gram wet weight (n=6). In Mytilus spp. the mean number of MPPs/gram wet weight was 3.0 (n=36). Fibres were the most common shape morphology of MPPs detected, utilising FT-IR and Nile Red staining techniques; PET was estimated to be the most common plastic type. The authors estimated that microplastic ingestion by humans via consumption of mussels was 123 MPPs/person/year in the UK, however, the risk of plastic ingestion via mussel consumption was minimal when compared to fibre exposure during an evening meal via dust fallout in a household at ~14,000-68,000 MPPs/person/year. These values were based on the following assumptions; deposition of 1 particle per 20 minutes for an area of 4.32 cm², this value was extrapolated to a 12.5 cm radius plate, resulting in 114 particles (assuming constant exposure rate), equating to approximately 42,000 MPPs consumption/person/year, for 20 minutes during consumption of an evening meal. During a cooking period of 20 minutes, a constant fibre fallout of 5 MPPs per 4.32 cm² was assumed, the potential human ingestion increases to \sim 207,000 MPPs/person/year. These values were then corrected by 33% which was reported to be the proportion of petrochemical based fibres found in dust by Dris et al., (2017).
- 83. An American study (Cox et al., 2019) has proposed an estimated daily consumption and inhalation of 142 MPPs and 170 MPPs in adult male, respectively. For adult females, the estimated values are 126 MPPs and 132 MPPs, respectively for the same exposure routes. Based on these values, a total annual estimated exposure of ~120,00 and ~98,000 MPPs annually was calculated in male and female adults, respectively. These exposure estimates were based on reported microplastic concentrations in salt, alcohol (beer), seafood (fish, shellfish and crustaceans), added sugars (sugar and honey), water (bottled and tap), and

in air. Note that the estimated annual exposure values did not take into account atmospheric deposition of microplastics during food preparation and consumption. The authors are of the view that "these estimates are subject to large amounts of variation; however, given methodological and data limitations, these values are likely underestimates."

- 84. Inhalation of microplastics can result in toxicity due either to the particles (a physical effect) or their leachates (a chemical effect). The author concluded the response in humans depends on differences in individual metabolism and susceptibility (Amato Lourenço et al., 2020). It is not yet known how the toxicity of synthetic fibres compares with that of organic/natural fibres (Donaldson & Tran, 2002). However, it is known that fibres from synthetic textiles are quite flexible (Bunsell (ed), 2018) and hence do not possess the characteristic long thin morphology of asbestos fibres, which is responsible for their toxicity and carcinogenicity.
- 85. In general, the mechanisms of inhaled particle injury include dust overload where high surface area particles induce high chemotactic gradients that prevent macrophage migration, oxidative stress (production of reactive oxygen species, which induces cell injury and release of inflammatory mediators), cytotoxicity (free intracellular particles may damage cellular structures), and translocation (injury of secondary sites and vascular occlusion by particles or increased coagulability). Depending on the nature of the particle and the extent of exposure, such mechanisms might lead to adverse endpoints such as fibrosis, which can develop as a result of chronic cytotoxicity and inflammation.

Indoor exposure

- 86. Dris et al., (2017) investigated indoor (two apartments and one office) air samples in the city centre of Paris. Indoor concentrations of microplastics ranged between 1.0 and 60 fibres/m3. The fibres that were measured indoors consisted of 67% of natural materials, primarily made of cellulosic materials and the remaining 33% contained petrochemicals, predominantly polypropylene.
- 87. Zhang et al., (2020) collated data from 46 studies and calculated the annual intake of indoor and outdoor microplastics using an inhalation rate of 14.3 m³ per day as 1.9×10^3 - 1.0×10^5 and 0-3.0 x 10^7 particles respectively, confirming that there is increased exposure to microplastics in the indoor environment. Whereas Fang et al., (2022), calculated the annual atmospheric deposition of MPs as 3.5×10^5 2.2×10^7 items (Figure 3).

- Figure 3. Diagram of microplastics (MPs) via the inhalation and ingestion routes of exposure showing that ingestion of microplastics is minimal in comparison to the inhalation route, whereas microplastics that have deposited on food and then ingested was of a similar magnitude to the microplastics via the inhalation route(Taken from Fang et al., 2022).
- 88. A recent study conducted in Hull, UK sampled 20 households each month for a 6 month period for atmospheric fallout detecting an average of 1414 microplastics/ m^2 per day with particles in the size range of 2-250 μ m contributing 90% of the particles found. Polyethylene terephthalate (PET), polyamide(PA) and polypropylene(PP) were the most abundant materials in the samples collected (Jenner et al., 2021).
- 89. Microplastics have been identified in all areas of the lung from tissue samples obtained following surgical resection for cancer or lung reduction surgery. Table 1 below contains the information obtained from the tissue samples, including the length and width of the particles found. Data was not normally distributed (p = 0.013) and a Kruskal-Wallis test showed that the number of NMPs in the lower region were significantly higher than the middle/lingular (p = 0.038) and the upper region (p = 0.026). Within the upper region (n = 6, total mass = 33.66 g), 11 NMPs were identified; PE (polyethylene) (18%), PP (18%), PES (polyester) (9%), PS (polystyrene) (9%), resin (9%), SEBS (styrene-ethylene-butylene co-polymer) (9%), TPE (thermoplastic elastomer) (9%). Within the middle/lingular region (n -3, total tissue mass = 12.19 g), 7 NMPs were identified; PET (29%), resin (29%), PE (14%), PMMA (polymethylmethacrylate) (14%), PUR (polyurethane)(14%). Within the lower region (n = 4, total tissue mass = 9.56 g), 21 NMPs were identified; PP (33%), PTFE (polytetrafluoroethylene) (19%), PET (14%), Resin (14%), PS (10%), PAN (polyacrylonitrile)(5%), PE (5%) (Jenner et al. 2022) (see Figure 4).
- Figure 4. Diagram showing the difference polymer types discovered in the lung and the area of the lung Particle number (total MPs detected with no account taken for MPs found in controls) and polymer type of MPs identified within human lung tissue samples, assigned to their lung region. (Taken from Jenner et al., 2022).
- Table 1. Data obtained in the Jenner et al. 2022 study including the length and width of particles and region of the lung particles are found (taken from Jenner et al. 2022)

90. The concentration of microplastics in indoor air is dependent on what occurs in the environment, for example, whether it is a home or occupational setting (discussed below in para. 95-96).

Occupational exposure

- 91. Certain occupations for example working in the synthetic and flock material industries can result in higher exposure to microplastics increasing the risk of developing occupational diseases. Burkhart et al., (1999) reported that in the nylon flocking area of a manufacturing plant, where the nylon particles are applied to a product, the highest concentration of airborne particles reached 7 mg/m^3 .
- 92. Characterisation of MPs from nail salons found the predominant sizes to be 50 μ m and the estimated average annual indoor exposure of MPs was 67,567 \pm 81,782 MPs/year. The polymers were predominantly acrylic (27%), rubber (21%) and polyurethane (13%), however the characteristics and polymer composition differed between nail salons. Factors that influenced the concentrations of MPs included the type of nail treatment being conducted, the use of air conditioning units and number of people present (Chen et al., 2022). Therefore, concentrations of MPs are such that they are more likely to affect human health from occupational exposure than from indoor exposure at home and the concentration is dependent on the atmospheric conditions (Prata, 2018).

Risk assessment tools

- 93. To date there are no standardised characterisation, collection and analytical methods for airborne microplastics or comprehensive risk assessment of NMPS. However, studies are beginning (Koelmans 2022a) to suggest ways in which this could be done.
- 94. It has also been suggested that an adverse outcome pathway (AOP) framework including the mechanisms of adverse effects, and new approach methodologies (NAMs) can be used in improving the decision making process with regard to microplastic hazard assessment. The use of read-across, computational models and "omics" cannot only reduce the number of animals used and the traditional testing methodologies, but might also provide a more robust scientific basis for decision-making (Halappanavar and Mallach 2021).

COT evaluation

- 95. NMPs are widespread, they are either intentionally added to products or occur as a result of plastics being fragmented down into smaller sizes by natural processes such as wear, weathering and corrosion. There is no internationally agreed definition of what a microplastic is, however, the most widely used size range is from $0.1\text{-}5,000~\mu m$. Plastic particles that are smaller than the lower range are considered nanoplastics (*i.e.* 1 nm $0.1~\mu m$).
- 96. Microplastics can have a wide range of physicochemical properties, depending on the primary purpose of the plastic; however, these properties may not be the same in secondary microplastics, where fragmentation has occurred as a result of natural processes (and as such the MPPs are not considered pristine). Additionally, analytical methodology is limited to Fourier-transform infrared spectroscopy(FT-IR), Nile Red, quantitative nuclear magnetic resonance(qNMR), Micro-Raman spectroscopy and mass-spectroscopy.
- 97. There are no standardised testing methods for different matrices such as air, soil, food and water; the available methods have their own associated limitations, and suitable reference materials are not currently available. Furthermore, no single technique is suitable for all plastic types and/or for all particle sizes or shapes. Using a suite of methods or generation of new techniques may be necessary to fully assess microplastics.
- 98. In terms of the toxicity of NMPs, there is no identified No Observed Adverse Effect Level (NOAEL) for any polymer type (with the possible exception of PET powder at 2,500 mg/kg bw/day in rats for oral exposure as reported by Merski et al., 2008, however, this study has several limitations and was conducted using the oral exposure route). Available data from the European Chemical Agency registration, evaluation, authorisation and restriction of chemicals (ECHA REACH) database relates only to the starting materials *i.e.* the monomers. Furthermore, variability in exposure routes must also be considered.
- 99. Comparing studies using different methodology and analytical techniques can be challenging as there is currently no standardization for characterizing and testing microplastics.
- 100. Contamination with airborne microplastics or cross-contamination of samples may also affect the interpretation of studies, so suitable control samples may be difficult to obtain.

- 101. The COT also considered whether particles arising from tyre wear were considered to be microplastics (TOX/2020/15 Annex B1). It was concluded that human exposure can occur from airborne particulate matter derived from tyre wear material. However, challenges arise in evaluating the risk of this particulate matter due to other variables such as aging and weathering of tyres, temperature effects, the types of road surfaces they are used on and driving style. These variables result in the generation of a variety of chemicals which have significantly different biological and toxicological potencies and effects.
- 102. Most toxicity studies have been performed with pristine particles, mostly polystyrene; however, these may not be representative of what is present in the environment as the particles have not undergone degradative processes or contain any additional pollutants that attach to the microplastic. There are no specific reference materials that can be used and batch to batch variation can also occur.
- 103. Currently a full risk assessment on the potential toxic effect(s) of NMPs could not be carried out due to several data gaps including:
 - The unavailability of harmonised methodologies to characterise, quantify and identify NMPs;
 - The lack of toxicokinetic and toxicity data in general. There is no identified NOAEL for the different polymer types except possibly for PET powder by the oral route at 2,500 mg/kg bw/day in rats, (see paragraph 98), which had a number of limitations (e.g. particle size and count were not determined/reported);
 - The paucity of currently available data for microplastics and airborne exposure and;
 - The difficulty of performing an accurate exposure assessment.
- 104. For the reasons above, a case-by-case approach to risk assessments may need to be considered. This aligns with the conclusions reached by other authoritative bodies (WHO, 2022; Environment and Climate Change and Health Canada(ECC and HC), 2020; EU Science Advice for Policy by European Academies (SAPEA), 2019; EU Group of Chief Scientific Advisors; Scientific Advice Mechanism (SAM), 2019, as described in the COT overarching statement on the potential risks from exposure to microplastics; COT Statement 2021/02, paragraphs 101-129).

Research priorities for risk assessment

Data gaps

- 105. For the inhalation route the significant data gaps include the lack of:
 - Harmonised analytical methods for detection of different NMPs during sample collection;
 - Methods of detection of NMPs in tissues and their systemic effects;
 - Understanding the contribution and effects of different exposure scenarios (e.g. indoor and outdoor environments);
 - Understanding how different pre-existing lung and other lung disease states may be involved in the observed effects from microplastic exposure and;
 - How available occupational data and on other particle types should be extrapolated to the general population.
 - Data on inhalation exposure to MPs that are resuspended in an indoor environment

Research priorities

- 106. The COT recommends the following research priorities in order of importance for addressing the data gaps in the potential toxicity of NMPs in humans and suggest a call should be put out to researchers. Information in these areas will assist in the future risk assessment of these particles by inhalation and other routes of exposure.
- 1. Development of reference standards and materials for use in wet lab/experimental settings. Also, development of appropriate fit-for-purpose quantification and detection methodologies for micro and nanoplastics in different matrices.
- 2. Studies (*in silico*, *in vitro* and/or *in vivo*) to explore the effect(s) of the same type of NMP on different tissues, and of different types of NMP (e.g. polymer type, size, shape) on the same target tissue. Physicochemical properties
- 3. Studies on the persistence and potential accumulation of NMPs in the human body, and on the extent to which NMPs are digestible. Kinetics
- 4. Investigation of the extent to which NMPs with a range of sizes and compositions are assimilated into human tissues and the development of techniques capable of identifying the presence of microplastics in the human body (e.g. in biopsies, samples from tissue banks, if possible, histopathology sections). Biological effects

- 5. From studies of particles at the nanoscale, it is known that nanoplastics can deposit lower down in the lung and have been shown to translocate across the pulmonary cellular barrier to secondary organs (Fournier et al., 2020). Therefore, more studies looking into the potential effects of nanoplastics are needed to understand size related effects. Physicochemical properties
- 6. Comprehensive assessment of MPs and their associated contaminant concentrations and matrices (such as air, soil, food and water) and the impact of the effect of cooking on the desorption and subsequent bioavailability of contaminants/leachates in relevant matrices is required. Chemical properties/migration /degradation.
- 7. Assessment of the degradation of novel/emerging plastic-based materials on the market such as biobased plastics (for example bamboo ware, polylactic acid, chitin) and other advanced polymer matrix composite materials during their use and end-of-life for their possible contribution to NMPs. It is unclear whether, and by how much, they already contribute to the burden of NMPs or similar particles. Migration and degradation
- 8. Microplastic concentrations in the environment are expected to increase in the future. In addition, increased and widespread use of single-use plastic personal protective equipment due to the COVID-19 pandemic may also be a significant contributing source of plastic pollution (Silva et al., 2021; Ma et al., 2021; Devereux et al. 2023). However, the quantification methods for microplastic particulate matter is currently limited and can only be estimated, thus improved technology is required. Therefore, there will be a need to regularly assess the levels of NMPs in relevant food stuffs, water and the air, such as by establishing a monitoring programme. This data would ideally be shared by collaboration among academia, researchers and government bodies at a national and international level. Migration/degradation and analytical methodology/detection.

COT conclusions

107. The COT noted that there are limited data regarding the toxicokinetic fate of inhaled microplastics in mammalian species. The extent to which retention in the lung is of concern is not yet clear. No epidemiological or controlled dose studies that evaluated the effects of inhaled microplastics in humans were identified.

- 108. As such, the COT concludes that based on the available data, it is not yet possible to perform a complete assessment for the potential risks from exposure to micro and nanoplastics *via* the inhalation routes; however, they concur with the conclusions reached by other authoritative bodies (EFSA (2016, 2020, 2021), WHO (2019, 2022), ECCC (2020) and HC, SAPEA (2019), SAM (2019), as described in the COT overarching statement on the potential risks from exposure to microplastics: COT Statement 2021/02, paragraphs 101-129).
- 109. The COT concluded that the literature data on exposure to particles from tyre wear would need separate consideration from microplastic exposure from food, since the particles are chemically quite different in their polymeric nature. Risk assessment of such material was considered potentially outside the scope of the current exercise.
- 110. The most significant data gaps are the lack of appropriate and harmonised analytical methods for the detection of micro- and nanoplastics (together with suitable reference standards), as well as information on their toxicokinetic and toxicity profiles in/relevant for humans.
- 111. The COT highlighted that additional information will be needed from all exposure sources, which include indoor and outdoor air, dust and soil before a risk assessment can be completed. The presence of MPs in food and water needs to be put into perspective with other sources of MPs such as atmospheric fallout.
- 112. Current studies typically focus on only one type of particle/tissue interaction, as such, further research is necessary to explore the effects of the range of particle types in different tissues *in vitro* and/or *in vivo*. This range of particle types should also take account of emerging/novel plastic-based materials such as bioplastics.

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Abbreviations

1 α-antitrypsin

Alpha-1 antitrypsin

ABS

Acrylonitrile butadiene styrene

COT Committee on Toxicity of Chemicals in Food, Consumer

Products and the Environment

COMEAP Committee on Medical Effects of Air Pollution

CONTAM Contaminants in the Food Chain

Defra

Department for Environment, Food and Rural Affairs

EC European Commission

ECC Environment and Climate Change

ECHA European Chemicals Agency

EFSA European Food Safety Authority

FT-IR Fourier-transform infrared spectroscopy

FSA Food Standards Agency

GIT Gastrointestinal tract

HC Health Canada

ILSI International Life Sciences Institute

MILC Mothers' information on lactation and collection

MOE Margin of exposure

MPPs Microplastic particles

NEE Non-exhaust emission

NMPs Nano- and microplastics

NOAEL No observed adverse effect level

OECD Organisation for Economic Co-operation and

Development

PAHs Polyaromatic hydrocarbons

PCBs Polychlorinated biphenyls

PE Polyethylene

PET Polyethylene terephthalate

PM10

Particulate matter (10 μm)

PP

Polypropylene

py-GC-MS

Pyrolysis coupled with gas chromatography and mass

spectroscopy

qNMR

Quantitative Nuclear Magnetic Resonance

RAC

Committee for Risk Assessment

REACH

Registration Evaluation Authorisation and Restriction of

Chemicals

ROS

Reactive oxygen species

RUBIAC

Rubber Industry Advisory Committee

SAM

EU Group of Chief Scientific Advisors; Scientific Advice

Mechanism

SAPEA

EU Science Advice for Policy by European Academies

SEAC

Committee for Socio-economic Analysis

TDS-GC-MS

Thermodesorption gas chromatography with mass

spectrometric detection

TWPs

Tyre wear particles

TRWPs Tyre and road wear particles

UK United Kingdom

UKWIR United Kingdom Water Industry Research

US United States

VOCs Volatile organic compounds

WHO World Health Organisation

Zonula Occludens Scaffolding proteins

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Annex B to TOX/2022/38

TOX/2019/62 (22/10/2019)

Paper 1: Scoping paper on the potential risks from exposure to microplastics.

TOX/2020/15 (11/03/2020)

Paper 2: Potential risks from exposure to microplastics: First draft overarching statement (Cover page).

Annex A1

First draft overarching statement on the potential risks from exposure to microplastics.

Annex B1

Paper for information: Background on tyre wear.

Annex C1

Paper for information: Update on literature.

TOX/2020/40 (15/09/2020)

Follow-up to Paper 2: Overarching statement on the potential risks from exposure to microplastics (Cover page).

Annex A2

Second draft overarching statement on the potential risks from exposure to microplastics.

TOX/2020/58 (01/12/2020)

Follow-up to September 2020 meeting: Overarching statement on the potential risks from exposure to microplastics: Third draft (Cover page).

<u>Annex A</u>3 Third draft overarching statement on the potential risks to microplastics.

COT Statement Number 2021/02

Follow-up to December 2020 meeting: Overarching statement on the potential risks from exposure to microplastics.

COT Statement Number 2021/05

Sub-statement on the potential risk(s) from exposure to microplastics: Oral Route.