



# Life cycle of tattooing pigments in the human body

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4th Joint Symposium on Nanotechnology, 30-31 May 2022

## The spectrum of body modifications and their perception in society

What is tattooing?

.... a mixture "for tattooing purposes" means injection or introduction of the mixture into a person's skin, mucous membrane or eyeball, by any process or procedure (including procedures commonly referred to as permanent make-up, cosmetic tattooing, micro-blading and micro-pigmentation), with the aim of making a mark or design on his or her body.



BfR Datenbank







https://www.flickr.com/photos/governmentofalberta/5815706525/



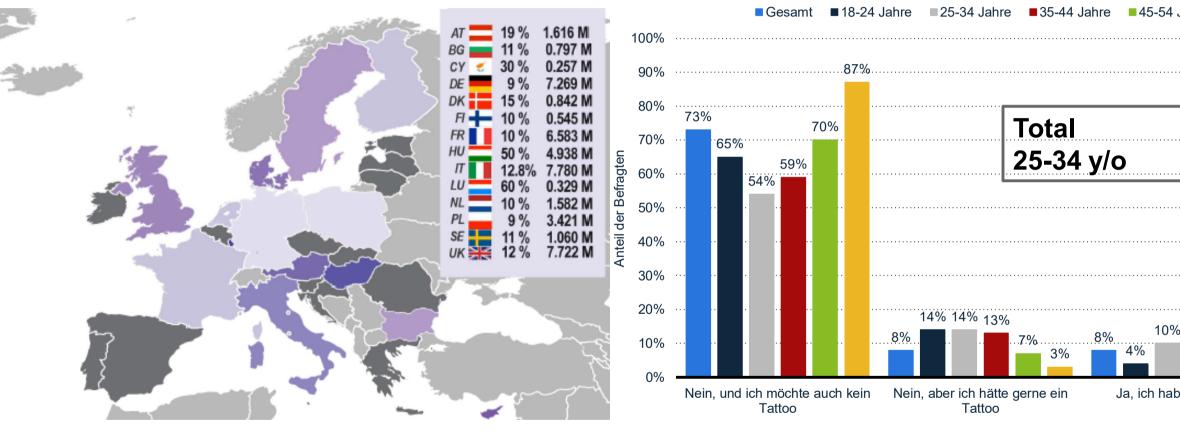
https://flickr.com/photos/47166475@N05/2001739496



### Are you wearing a tattoo?

**JRC 2016** 

### Survey in Germany on tattoos by



### YouGov; ID 1253983

https://de.statista.com/statistik/daten/studie/1253983/umfrage/umfrage-in-deutschland-zu-tattoos-nach-altersgruppen/

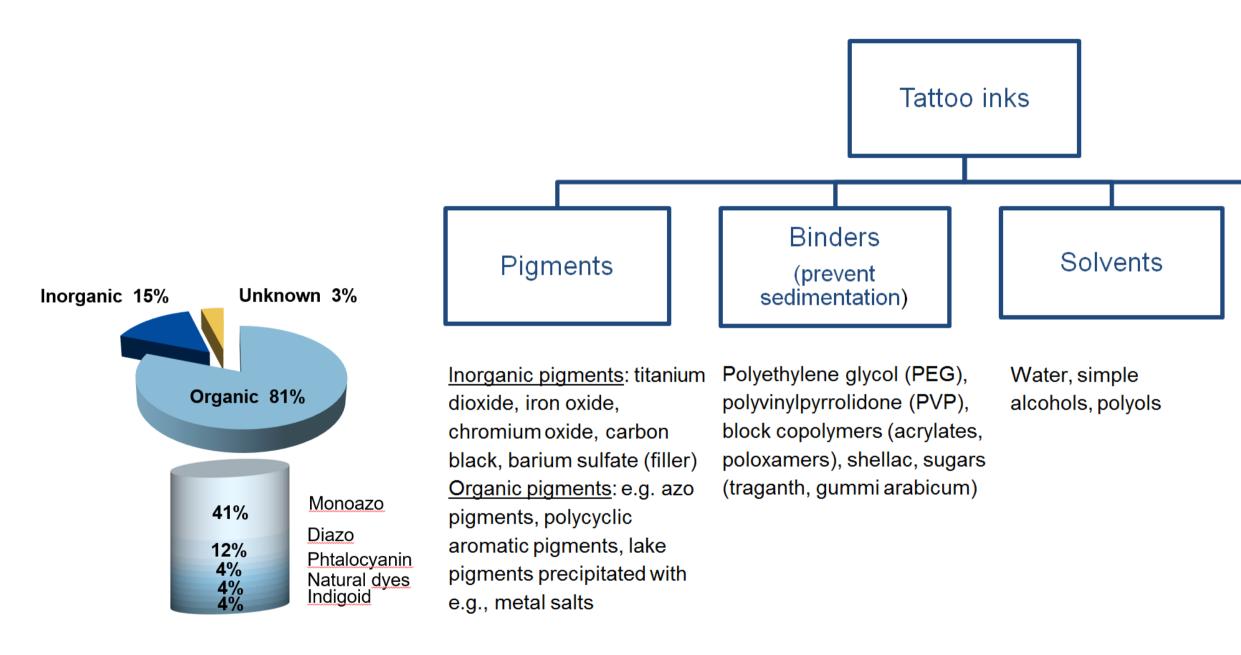
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/ age group 2	2021
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Jahre 55 Jahr	e und älter
470/	
17%	
26%	
6 11% <sub>10%</sub>	9% <sup>12%</sup> 14% 11%
5%	4%
be ein Tattoo	Ja. ich habe mehrere Tattoos



### Composition of tattoo inks



M. Giulbudagian et al., (2020) Arch Toxicol & JRC (2016)

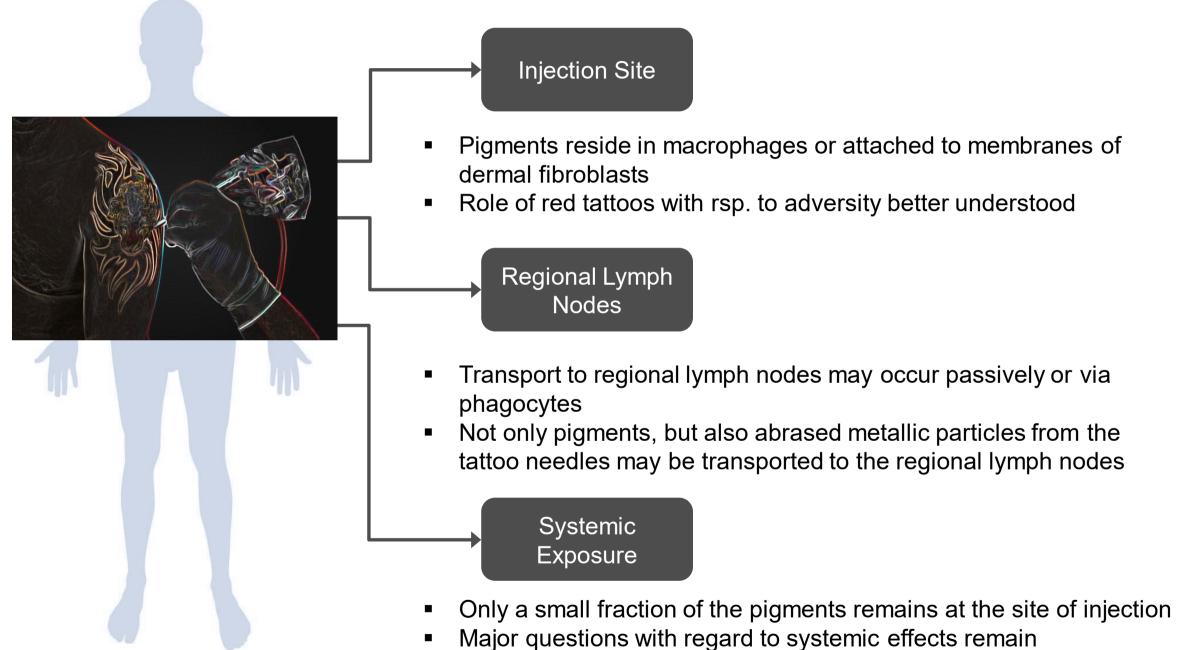
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### Additives

Preservatives, thickening agents, surfactants



## What do we know about the distribution of tattoo pigments in the body?

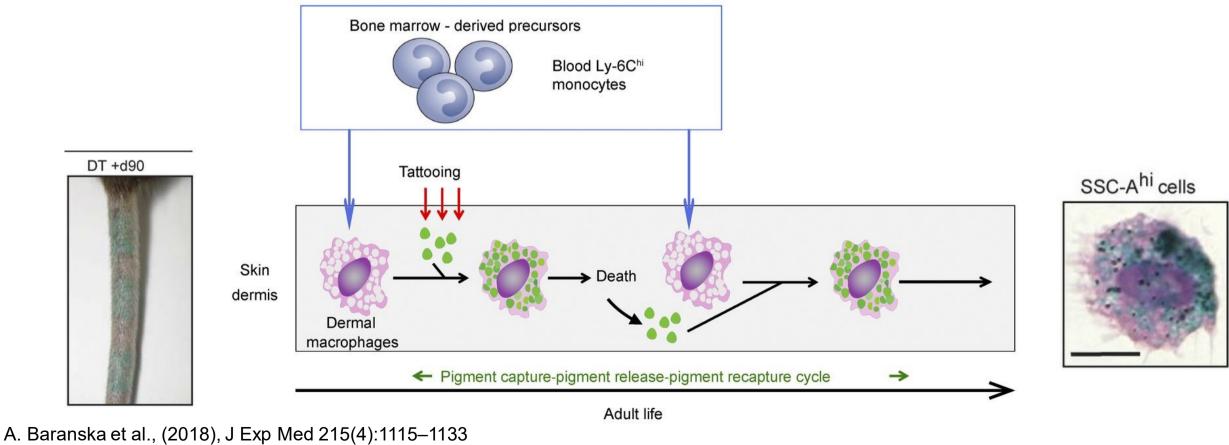






### What happens at the injection site?

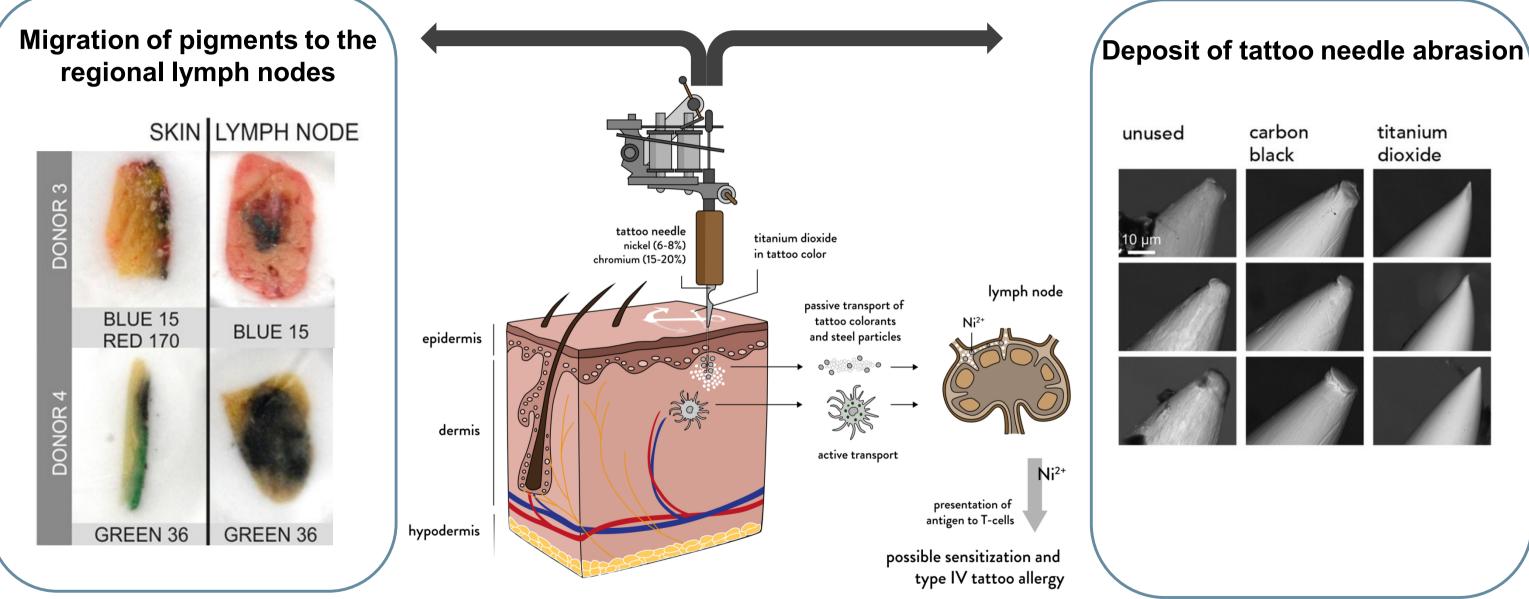
- Pigments injected into the dermis reside in cells and connective tissue.
- Tattoo pigments injected into the mouse tail remained mainly in the dermal resident macrophages after endocytosis.
- Free pigment particles are readily internalised by newly forming macrophages.



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### What happens at the regional lymph nodes?



I. Schreiver et al., Scientific Reports (2017), 7 (1), 11395 I. Schreiver et al., (2019) Part Fibre Toxicol 16(1):33

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## What is known about the systemic distribution of tattoo inks?

- Direct contact of tattooing agents with blood and lymph fluids takes place.
- A rapid systemic distribution of the soluble ingredients can be assumed.
- Studies suggest that over 80% of the injected pigments are eliminated.
- In animal studies, pigments could be detected in the liver of mice one year after administration.
- No correlation could be found between tattoos and cancers or other systemic complications.





### Lerche et al., (2015) Photoimmunol Photomed 31(5):261-268



### Minimum Testing Requirements

### **Specifications for Ingredients of Tattoo Inks**

Purity

- Stability
- Additional information on pigment particles:
  - i. Number-based particle size distribution
  - ii. Morphology/shape/crystalline structure
  - iii. Surface properties
  - iv. Dispersibility
  - v. Density

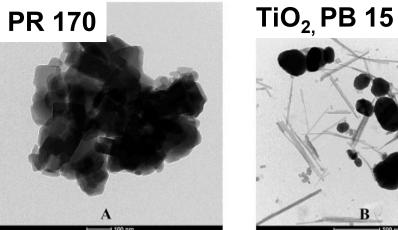
### Minimum toxicological requirements - in vitro/in chemico testing • Consider compatibility of tests with the physico-chemical properties of pigments Test according to OECD Test Guidelines and GLP Eye irritation & Skin irritation Skin sensitization **Phototoxicity** Mutagenicity/genotoxicity damage & corrosion incl. photogenotoxicity

### Tattoo pigments that meet the minimum toxicological requirements reduce possible risks according to the current state of science and technology.

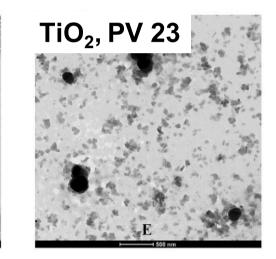
https://www.bfr.bund.de/cm/349/tattoo-inks-minimum-requirements-and-test-methods.pdf

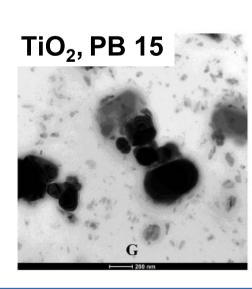
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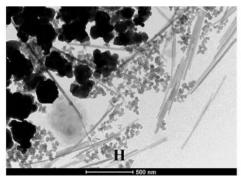


### PO 16, PR 210 🌌





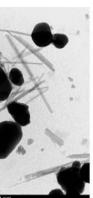
### Carbon Black, **Iron Oxides**

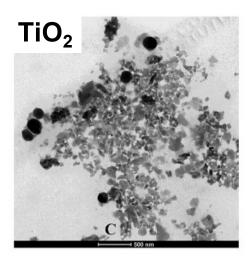


## Pigment particles in tattoo inks

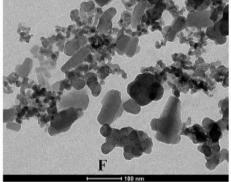
Bocca et al (2017) Journal of Analytical Atomic Spectrometry 32(3):616-628 Battistini et al (2020) Chemosphere 245:125667

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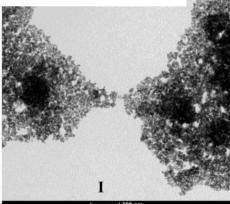








### **Carbon Black**







### Asymmetric Flow Field Fractionation (AF4) and Multi-Angle Light Scattering (MALS)-ICP-MS

Ink number (name)	TEM, <sup>a</sup> nm	DLS, <sup>c</sup> nm	AF4-MALS, <sup>d</sup> nm	SP nn
2 (Ice blue)	170 $\pm$ 64 (127–227)	421 (110–980)	42 (16–65) 269 (60–490)	10 18 44
5 (Deep violet)	$39 \pm 10~(3148)^b$	31 (19–52) 137 (60–281)	56 (11–100) 271 (100–490)	42
6 (Black outlining)	$19 \pm 11 \; (1229)$	152 (49–450)	137 (21–330)	10
7 (Grasshopper green)	$162\pm51~(126 extrm{}208)$	277 (81–1350)	46 (18–56) 405 (50–1200)	11 22 46

<sup>*a*</sup> Diameter, median  $\pm$  median absolute deviation (first-third quartile). <sup>*b*</sup> Larger particles (100–450 nm) were also observed. <sup>*c*</sup>  $D_{\rm h}$ , hydrodynamic diameter, mean (range). <sup>*d*</sup>  $R_{\rm g}$ , radius of gyration (=diameter of gyration ( $D_{\rm g}$ )/2), mean (range). <sup>*e*</sup> Diameter, mean  $\pm$  standard deviation.

Ink	Peak	Rt (min)	Rg (nm)	< 100 nm
Ice blue	1	17-28	42 (16-65)	53%
	2	28-56	269 (60-490)	
	1	15-32	56 (11-100)	78%
Deep Violet	2	32-54	271 (100-490)	
Black Outlining	1	17-50	137 (21-330)	77%
	1	16-27	46 (18-56)	32%
Grasshopper Green	2	27-56	405 (50-1200)	
				page 11

Bocca et al (2017) Journal of Analytical Atomic Spectrometry 32(3):616-628 Battistini et al (2020) Chemosphere 245:125667

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SP-ICP-MS,<sup>e</sup> m

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109 \pm 17 (Cu-phthalocyanine)
183 \pm 11 (Al_2O_3)
441 \pm 161 \, (TiO_2)
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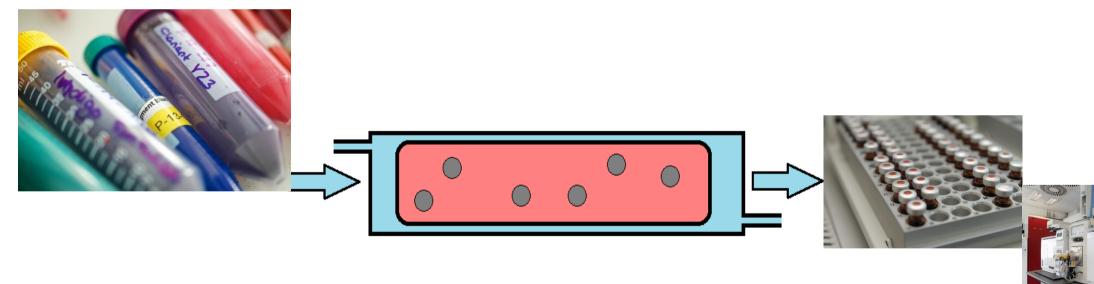
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427 \pm 97 (TiO_2)
```

 $109 \pm 19$  (CuO)

 $110 \pm 17$  (Cu-phthalocyanine)  $226 \pm 14 (Al_2O_3)$  $466 \pm 179 \, (TiO_2)$ 

### In-vitro approach for investigation of pigment dissolution

- Long-term dynamic dissolution testing of tattoo pigments in skin simulant
- External factors like UV or temperature
- Fractionated sampling and analysis of pigments and/or metabolites
- Identification of pigments that release harmful substances under physiological conditions



### Consideration of DIN EN ISO 10993-16:2018-02, Biological evaluation of medical devices.

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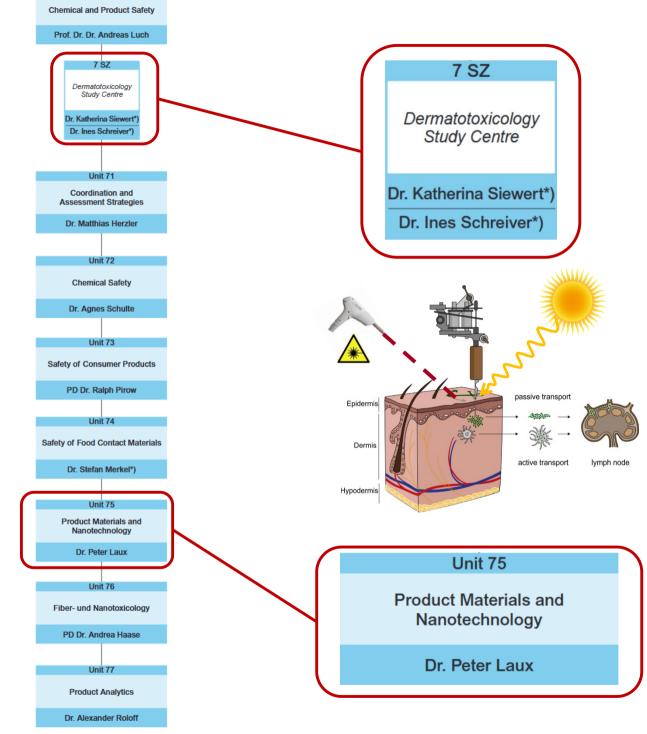


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Department 7

www	/.bfr.bund.de
Infe	ction Risks through Tattooing
www.bfr.bund.de	
FAQ about tattoo ink	S
Updated BfR FAQ, 16 Se	eptember 2019
the future. In the age gro too inks may consist of m their potential to be harm ments, capable of achiev	DOI 10.17590/20201006-102053 Tattoo inks: risk assessment for I BfR Opinion No 039/2020 issued 8 Sep
	To date, there is no binding regulation g European level. The EU Commission a
www.bfr.bund.de	
DOI 10.17590/20211021-11	5214
Tattoo inks: minimu	Im requirements and test me
Opinion No 031/2021 c	of the BfR of 14 October 2021
Tattoo inks contain pig	ments and additives. According to

the provisions of the German Food, Consumer Goods and Feed Code (Lebensmittel-, Bedarfsgegenstände- und Futtermittelgesetzbuch, LFGB), tattoo inks may not be used if there is any doubt as to their safety to health. Substances or mixtures for tattooing purposes are regulated in the REACH Regulation [entry 75 of Annex XVII of the REACH Regulation (Regulation (EC) No 1907/2006)].





Bundesinstitut für Risikobewertung



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### Pigment Blue 15:3 and Pigment Green 7

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# Thank you for your attention

https://www.philara.de/en/exhibitions/2021-02/timm-ulrichs

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### Identify Risks -**Protect Health**