

MEETING ABSTRACT

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Two decades of biomonitoring polar bear health in Greenland: a review

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Summary

We present an overview of studies of anthropogenic pollutants in East Greenland polar bears over the period of 1999-2011. East Greenland polar bears are among the most polluted species, not just in the Arctic but globally, and represent an excellent biomonitoring species for levels and effects of global pollution in an apex predator. Therefore, an international multidisciplinary team joined to monitor and assess the patterns and concentrations of contaminants and their potential negative impact on polar bears. The review showed that East Greenland polar bears are exposed to a mix of chlorinated, brominated and fluorinated organic compounds as well as mercury which are all known to have endocrine, immune and organ-system toxic properties. For example, the concentrations of PCBs (polychlorinated biphenyls) in blubber ranged approximately 800-21,000 ng/g lw while mercury concentrations in liver and kidney ranged 0.1-50 µg/g ww. Regarding health endpoints, bone density seemed to decrease as a function of time and OHC (organohalogen compound) concentrations and further T-score for adult males indicated risk for osteoporosis. The size of sexual organs decreased with increasing OHC concentrations. In the lower brain stem, mercury-associated decreases in NMDA-receptor levels and DNA-methylation was found. The present review indicated that age was one of the major drivers for liver and renal lesions, although contaminants and infectious diseases may also play a role. Lesions in thyroid glands were most likely a result of infectious and genetic factors and probably, together with endocrine disrupting chemical (EDCs), the reason for disturbances/fluctuations in blood plasma thyroid hormone concentrations. Except for bone density reductions and neurological measures, all findings were supported by case-control studies of Greenland sledge dogs exposed long-term orally to similar combinations of contaminant concentrations. The studies of sledge dogs also indicated that the mixture of contaminants and fatty acids in the blubber of prey similar to that of polar bears induces cellular as well as humoral immune toxic changes. These controlled studies using model species for polar bears indicate that the correlative findings between health endpoint and contaminants in polar bears could be a cause-and-effect relationship. Physiologically based pharmacokinetic (PBPK) modelling showed that the risk quotients were ≥ 1 for Σ PCB, dieldrin and PFOS, which indicate an increased risk of prenatally reproductive pathology. In conclusion polar bears are susceptible to long-range transported chemicals that may have various adverse effects on multiple organ systems such as the reproductive and immune system.

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Background

The use of East Greenland polar bears (*Ursus maritimus*) as a biomonitoring key species for measuring exposure to OHCs (organohalogen compounds) and Hg (mercury) was initiated in Scoresby Sound in 1983 and is now the most comprehensive time trend studies of pollution on this species [1-7]. In addition, museum samples of skin and skulls collected since year 1892 have been included for Hg analysis and various patho-morphological and toxico-pathological density analyses [4,6-11].

The studies of adverse health effects of pollution were started in 1999 by the implementation of a biomonitoring health programme via AMAP (Arctic Monitoring and Assessment Programme) [12-14]. The reason for choosing polar bears, and for upgrading the research intensity, was because the East Greenland polar bear subpopulation is one of the most contaminated and that local Inuit people rely on this species as a food resource in addition to ringed seal (*Phoca hispida*) that plays a much greater role. Because polar bears reflect temporal trends and biological effects of contaminants they may also serve as a proxy for human health exposure and possible effects despite the fact that the physiology, metabolism, food and way of life of these two species differ fundamentally [e.g. [5,7,13-15]].

We present an overview of contaminant concentrations and potential adverse health effects from anthropogenic contaminants in polar bears during the period 1999-2010. The health effects include decrease in bone density, morphological changes in sexual organs, liver, kidney and thyroid glands, as well as potential neurological alterations and impacts on the immune and endocrine systems. Finally, some speculations about the synergistic effects of

environmental stressors (e.g. decrease in sea ice) on polar bear health are presented.

Levels of contaminants

Chlorinated legacy contaminants (PCBs and OC pesticides), brominated flame retardants (PBDEs), perfluoroalkyl contaminants (PFCs) and Hg have been analyzed in brain, adipose tissue, liver, kidney, blood and hair samples from East Greenland polar bears. The range of contaminant concentrations used in relation to health endpoints during the period 1999-2002 are seen in Table 1. Concentrations increase in the order: PBDEs<PFCs<PCBs<Hg and are in the concentration of having adverse health effects according to the international scientific literature [13,14,16,17].

According to Dietz et al. [3-5,7] the concentrations of legacy chlorinated contaminants (PCBs and OC pesticides) have decreased and stabilized since 1990, while newer contaminants like PFCs and contemporary threats like Hg have increased. So, despite international regulations on all these groups of contaminants they persist and biomagnify in the environment, which results in a cocktail of toxic chemicals in the tissues of East Greenland polar bears.

Skeletal system

Analyses on the skeletal system have exclusively focused on skulls. The reasons for this are because Natural History Museums have archived these since 1892 and because skulls are relatively easily obtained in connection with the subsistence hunt.

We examined skull bone mineral density (BMD) in 139 bears in the period 1892-2009 and after controlling for age

Table 1 Concentrations of various contaminants [Mean (Min-Max, n) divided on tissues in East Greenland polar bears sampled 1999-2002. All data are in ng/g lw except for nPCBs, PCDDs and PCDFs (pg/g lw), PFCs (ng/g ww) and mercury (µg/g ww). Reworked from Sonne (2010)]

	Subcutaneous adipose tissue	Liver	Kidney	Brain	Blood
Lipid-%	88	11	-	21	1.3
ΣPCB	6,543 (897-20,407, 92)	28,409 (12,836-67,664, 20)	-	148-2,186 (20)	538-15,692 (20)
HCB	102 (2.4-785, 92)	109 (20)	-	15 (20)	28 (20)
ΣHCH	194 (13-818, 92)	67 (20)	-	15 (20)	12-146 (20)
Dieldrin	204 (26-866, 92)	4,900 (20)	-	-	-
ΣChlordane	1,414 (243-7,465, 92)	37,400 (20)	-	62 (20)	531 (20)
ΣDDT	436 (73-1,580, 92)	<0.1-476 (20)	-	ND	12-1,769 (20)
ΣPBDE	70 (22-192, 92)	127-936 (20)	-	<0.5-36 (20)	38-146 (20)
ΣPFC	-	1,056-8,010 (29)	-	-	-
ΣnPCB	241 (125-442, 5)	124 (114-148, 5)	-	-	-
ΣPCDD	10 (7-12, 5)	20 (8-38, 5)	-	-	-
ΣPCDF	4 (3-5, 5)	14 (10-18, 5)	-	-	-
ΣOH/MeSO ₂ -PCB	90-3134 (20)	1,882-18,018 (20)	-	66-352 (20)	29,692-226,154 (20)
ΣOH/MeSO ₂ -PBDE	<0.3-39 (20)	<0.5 (20)	-	<0.5 (20)	<0.5-1,060 (20)
ΣMeSO ₂ -p,p'-DDE	9 (20)	482 (20)	-	-	769 (20)
Mercury	-	11 (1-36, 59)	14 (1-50, 57)	0.4 (0.1-0.9, 82)	-

and sex we found that BMD decreased significantly over time in subadults (Figure 1) and adult males but not in adult females (data not shown) [18]. That a similar decrease was not found in adult females was likely due to the fact that they increase bone density prior to denning in order to avoid demineralisation. Otherwise, the low mechano-transduction and limited food intake and high calcium flux via foetal development and lactation transfer during denning theoretically should lead to clinical osteoporosis [19].

When correlating BMD with individual body burdens of OHCs in adipose tissue for individuals sampled during 1999-2002, significant inverse relationships were found in subadults (Figure 2) and adult males but not females [18]. Similar relationships were found for baculum BMD [20]. Furthermore, a calculation of T-score in males showed their risk of developing osteoporosis [14] as neuro-signalling may be influenced due to disruption of calcium homeostasis [21]. The exact pathways for these relationships are unknown; however, OHCs may disrupt brain and endocrine organ hormones as well as their transport proteins in polar bears as shown by multiple laboratory experiments [13,14,18]. During the study period 1892-2002 the pack sea ice in East Greenland has decreased [22]. Although undocumented, Sonne et al. [18] and Sonne [14] assumed that availability of polar bear prey (in East Greenland a variety of marine mammal prey other than ringed seals are available to polar bears) decreased simultaneous with the decrease in offshore sea ice and speculated whether this apparent decrease in sea ice may have led to increased energy expenditure and therefore indirectly served as a synergistic factor leading to reduced BMD.

Reproductive organs

Studying impacts from OHCs optimally includes investigation of the reproductive tract as adverse effects on this

organ-system have a direct impact on the population size [14].

We studied gross morphology and histopathology in sexual organs from 55 male and 44 female polar bears sampled 1999-2002. When controlling for age, the analyses showed that size of testicles, baculum/penis, and ovaries/female reproductive tract decreased as a function of increasing OHC concentrations in adipose tissue (Figure 3) [20]. In addition to this we found enlarged clitoris (megaclitoris) in two adults but ascribed this to chronic inflammation rather than OHC exposure [14,23]. In adult males we found chronic orchitis, atrophy, and fibrosis independently of season, all of which could be partly ascribed to OHC exposure and infectious pathogens as shown by high-dose, short-term controlled laboratory experiments [20]. The pathways for these are suspected to be disruptions of brain and endocrine organ hormones as well as their transport proteins [13,20]. It is not possible to fully estimate the impact from these disruptions and pathological conditions, however, it is likely that semen quality and quantity could be influenced, which may have fatal consequences for fecundity and maintenance of population size.

Liver, kidneys and thyroid glands

Other key organs that are susceptible to endocrine disruption are the liver, kidneys, and various endocrine organs such as the thyroid glands and adrenals [13,14]. Liver (n=79), renal (n=75) and thyroid glands (n=20) were analyzed histologically in bears sampled 1999-2002. The evaluation, when not having a control group, is very difficult not least due to the effects from age and infectious diseases [14,24-28]. However, the various pathologies found in liver parenchyma and renal glomeruli, tubules and interstitium were similar to those found in controlled studies of Hg and OHCs in laboratory mammals and other OHC exposed wildlife [14,24-27]. The overall most

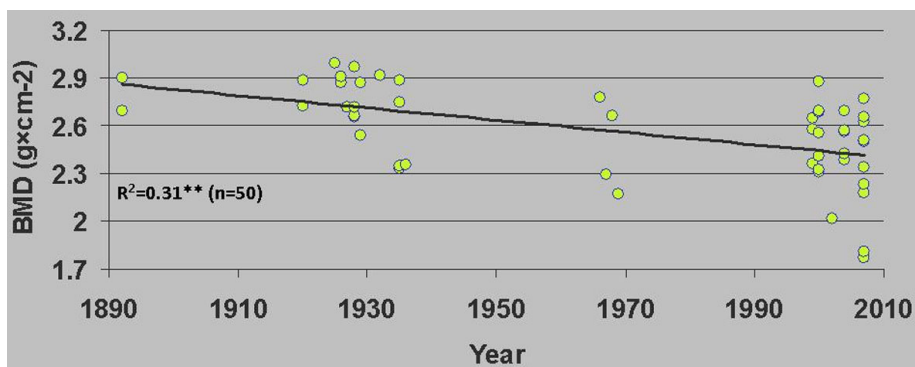


Figure 1 Bone mineral density ($\text{g}\times\text{cm}^{-2}$) as a function of sampling year in subadult East Greenland polar bears sampled 1892-2002. R^2 - and p-values from a full multiple regression model controlling for age. **: $p < 0.01$.

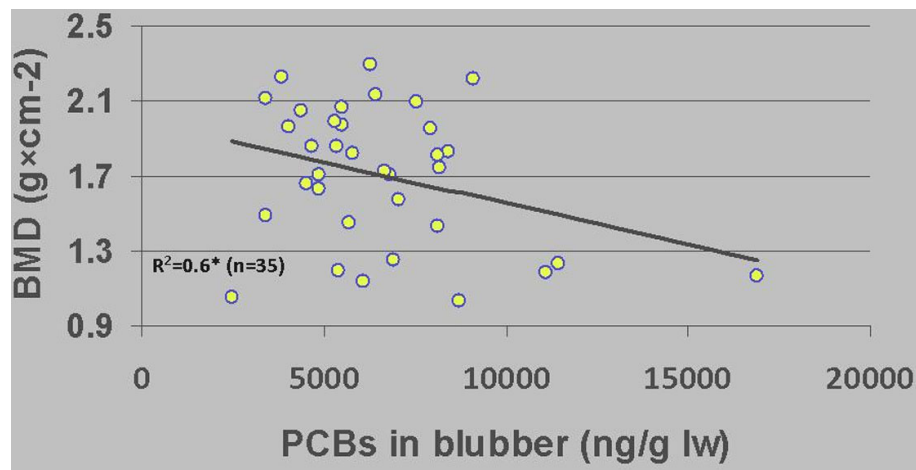


Figure 2 Bone mineral density ($\text{g}\times\text{cm}^{-2}$) as a function of adipose tissue PCBs (ng/g lw) in subadult East Greenland polar bears sampled 1999-2002. R^2 - and p -values from a full multiple regression model controlling for age. *: $p < 0.05$.

important parameters determining liver and renal pathology was age while also some statistical relationships were found between pathology and different chlorinated and brominated groups of contaminants as well as the liver and renal toxic Hg concentrations.

No histological lesions were found in any of the 50 adrenals examined while pathology was found in 8 out of 20 examined thyroid glands [28]. None of the thyroid gland lesions including c-cell hyperplasia, interstitial fibrosis, and nodular hyperplasia were associated with age or gender, so environmental factors such as energetic stress and autoimmunity/genetic could be co-factors as well as OHCs. Such lesions may interfere with the hypothalamic-pituitary-thyroid (HPT) axis leading to endocrine disruptions having an impact on fecundity and foetal and neonatal development in East Greenland polar bears.

Neuro-endocrine system

During 1999-2002 we sampled the medulla oblongata from 82 specimens in order to analyze the concentrations of Hg, a known neuro-toxicant. The analyses showed that

the concentrations of Hg were relatively low compared to liver and kidney burdens, as well as other species, probably due to high demethylation capacity of the liver, formation of Hg-selenium complexes, and fur as an efficient excretion route [29]. Despite the low Hg concentrations some inverse correlations of statistical significance were found. First, like in several other wildlife species [30,31], Hg-associated decreases in the levels of NMDA receptor were found. The NMDA-receptor facilitates the neurotransmission of glutamate and is important for learning and Hg (Figure 4). Second, DNA methylation seemed to decrease with increasing Hg concentrations indicating potential epigenetic alterations in gene expression [32].

Villanger et al. [33] analyzed the circulating full blood concentrations of TT3 and TT4 in 62 East Greenland polar bears. The reason for analyzing thyroid hormones is the fact that this endocrine system is known to be extremely susceptible to EDC exposure including bio-transformed metabolites such as OH-PCBs [13,14,33]. The analyses showed that various biological parameters such as age, size, and condition affected the circulating

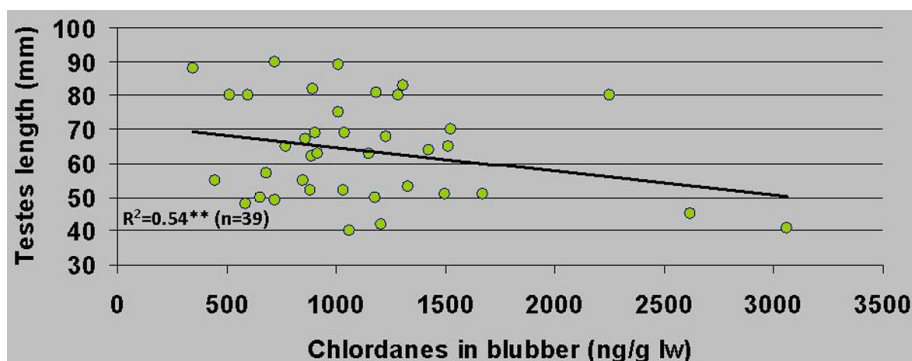


Figure 3 Testes length (mm) as a function of adipose tissue chlordane concentrations (ng/g lw) in 39 East Greenland polar bears sampled 1999-2002. R^2 - and p -values from a full multiple regression model controlling for age. **: $p < 0.01$.

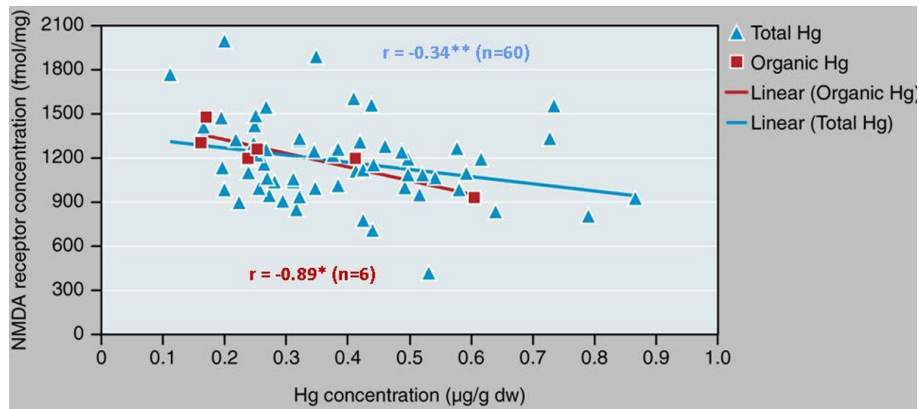


Figure 4 Correlations between NMDA-glutamate receptor levels and (A) Total Hg (blue) and (B) Organic Hg (MeHg; red) in the brain stem of 82 East Greenland polar bears sampled 1999-2002. Modified from Basu et al. [29]. *. $p < 0.05$.

concentrations while different organochlorine contaminants and bromated flame retardants had negative as well as positive effects on the concentrations of both TT3 and TT4. The conclusions were that these correlations indicate biological effects on the HPT axis from EDC exposure, which is also supported by various in vivo studies of laboratory mammals as well as in vitro studies of polar bears [13,14,33].

Immune system

Very little has been published on the East Greenland polar bear immune system [13,14,34]. However,

investigations on Svalbard bears have indicated immune toxic effects at OHC exposure concentrations similar to the East Greenland polar bear's [13,14,35,36] and multiple studies in the laboratory and field show similar biological effects [13,14,37-39].

Future challenges

As shown above, contaminant exposure is suggested to have potential health effects on various organ-systems in East Greenland polar bears. However, contaminants are not the only environmental stressor in East Greenland (Figure 5). Also global warming leading to decreased

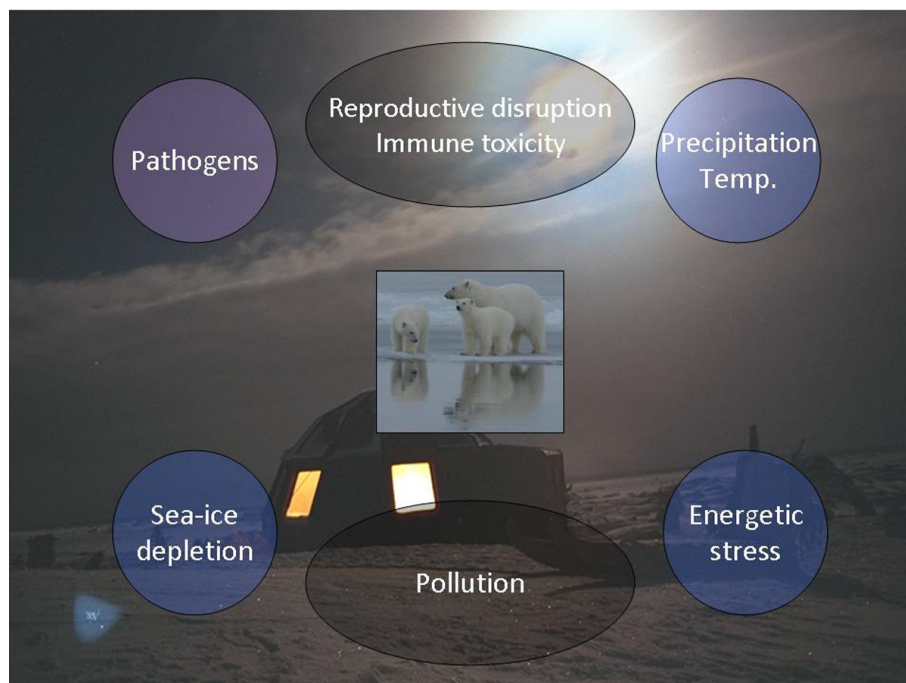


Figure 5 The multiple stressors in the Arctic that influences polar bear health. Oil and mineral activities, vessel shipment and subsistent hunt are not included.

food access and negative energy balance may influence bear health via (sub)clinical impacts on immune functioning and reduced fecundity having an impact on populating size [13,14,40-42]. In addition to this global warming may also increase the infectious stress due to invasive micro pathogen and parasitic diseases [13,14]. The main challenge in the future is therefore to integrate the cumulative impact from these multiple stressors across temporal and spatial gradients by integrating empirical data and laboratory studies. The East Greenland polar bear seems an excellent biomonitoring organism for such.

Conclusions

East Greenland polar bears are among the most contaminated species on our globe in spite of their remote Arctic habitat. This sub-population inhabits Arctic biotopes being constantly under pressure from global warming and associated environmental changes. Anthropogenic environmental contaminants seem to be a co-factor in various organ-system lesions in East Greenland polar bears. This includes reduced bone density and sexual and reproductive organ size, thereby having potential impacts on individual health and population maintenance. On top of this, also global warming seems to affect polar bears via negative energy balances, which may have consequences for fecundity and immune resistance. The main challenge in the future is to integrate the cumulative impact from these multiple stressors across temporal and spatial gradients by integrating empirical data and laboratory studies.

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