

Ecological Soil Screening Levels for Nickel

Interim Final

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1.0 INTRODUCTION

Ecological Soil Screening Levels (Eco-SSLs) are concentrations of contaminants in soil that are protective of ecological receptors that commonly come into contact with and/or consume biota that live in or on soil. Eco-SSLs are derived separately for four groups of ecological receptors: plants, soil invertebrates, birds, and mammals. As such, these values are presumed to provide adequate protection of terrestrial ecosystems. Eco-SSLs are derived to be protective of the conservative end of the exposure and effects species distribution, and are intended to be applied at the screening stage of an ecological risk assessment. These screening levels should be used to identify the contaminants of potential concern (COPCs) that require further evaluation in the site-specific baseline ecological risk assessment that is completed according to specific guidance (U.S. EPA, 1997, 1998, and 1999). The Eco-SSLs are not designed to be used as cleanup levels and the United States (U.S.) Environmental Protection Agency (EPA) emphasizes that it would be inappropriate to adopt or modify the intended use of these Eco-SSLs as national cleanup standards.

The detailed procedures used to derive Eco-SSL values are described in separate documentation (U.S. EPA, 2003, 2005). The derivation procedures represent the collaborative effort of a multi-stakeholder group consisting of federal, state, consulting, industry, and academic participants led by what is now the U.S. EPA Office of Solid Waste and Emergency Response (OSWER).

This document provides the Eco-SSL values for nickel and the documentation for their derivation. This document provides guidance and is designed to communicate national policy on identifying nickel concentrations in soil that may present an unacceptable ecological risk to terrestrial receptors. The document does not, however, substitute for EPA's statutes or regulations, nor is it a regulation itself. Thus, it does not impose legally-binding requirements on EPA, states, or the regulated community, and may not apply to a particular situation based upon the circumstances of the site. EPA may change this guidance in the future, as appropriate. EPA and state personnel may use and accept other technically sound approaches, either on their own initiative, or at the suggestion of potentially responsible parties, or other interested parties. Therefore, interested parties are free to raise questions and objections about the substance of this document and the appropriateness of the application of this document to a particular situation. EPA welcomes public comments on this document at any time and may consider such comments in future revisions of this document.

2.0 SUMMARY OF ECO-SSLs FOR NICKEL

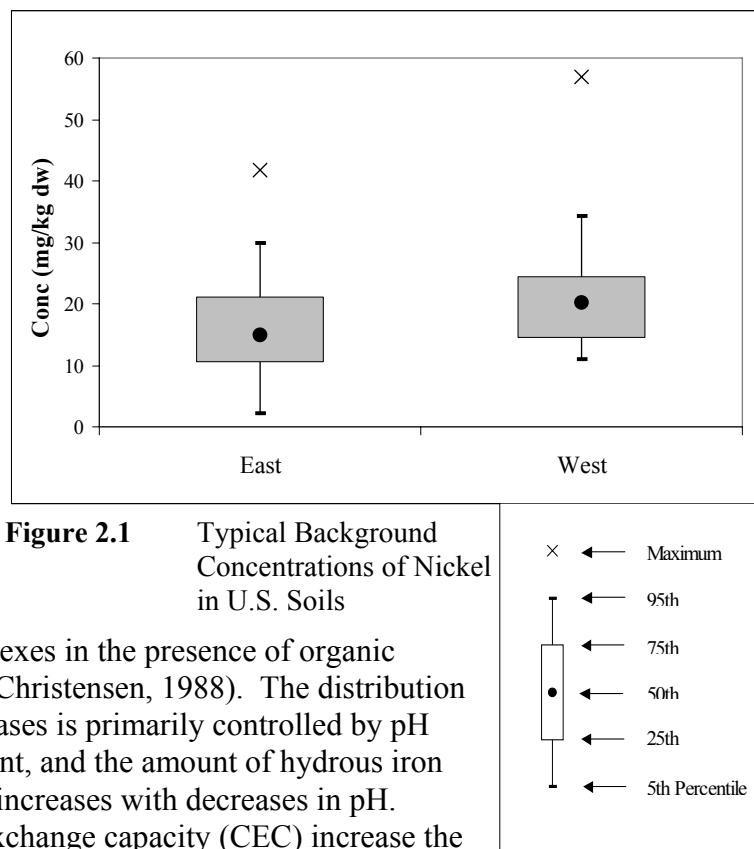
Nickel is a naturally occurring element which can be found in all environmental media: air, soil, sediment, and water. In the metal state, nickel is silvery white, hard, malleable, and ductile. It is somewhat ferromagnetic, and a fair conductor of heat and electricity. Nickel occurs in numerous minerals as sulfides, arsenides, antimonides and oxides or silicates. Primary sources include chalcopyrite, pyrrhotite, pentlandite, and garnierite (Budavari, 1996; HSDB).

Nickel is released to the environment through the extraction, processing and use of nickel compounds (HSDB). The single largest use of nickel is in the manufacture of stainless steels (Alloway, 1990). Nickel is also used in the production of alloys with other metals such as iron, copper, chromium, and zinc (ATSDR, 1988; HSDB). Other major uses are in electroplating alloys, nickel-cadmium batteries, electronic components, fuel cells, specialty ceramics, magnets, specialty chemicals, filters for gases, hydrogenation of fats, petroleum products, preparation of colored pigments and for color stabilization of color copy paper (ATSDR, 1988; Alloway, 1990). Nickel may also be released from natural sources, such as volcanoes, windblown dusts, the weathering of rocks, forest fires, and decaying vegetation (Davies, 1974; HSDB).

In the atmosphere, nickel is expected to exist in the particulate phase and is released to soils through wet and dry deposition. The species of nickel present in deposition include soil minerals, oxides and sulphates (Alloway, 1990). The largest anthropogenic sources of nickel to the atmosphere result from the burning of fuel and residual oils followed by diesel exhaust, the combustion of coal and nickel mining and smelting (Alloway, 1990).

Background concentrations reported for many metals in U.S. soils are described in Attachment 1-4 of the Eco-SSL guidance (U.S. EPA, 2003). Figure 2.1 provides a plot of the typical background concentrations of nickel in U.S. soils in the eastern and western portions of the country.

In soils, nickel may be present as soluble compounds including chlorides and nitrates, and insoluble compounds such as oxides and sulfides. Soluble nickel compounds tend to exhibit greater mobility than insoluble nickel compounds (Dean, 1985; HSDB). The degree of mobility is influenced by the formation of complexes in the presence of organic substances and sulfates (Anderson and Christensen, 1988). The distribution of nickel between solid and solution phases is primarily controlled by pH with secondary factors being clay content, and the amount of hydrous iron and manganese oxides. Soluble nickel increases with decreases in pH. Increases in metal loading and cation exchange capacity (CEC) increase the amount of metal adsorbed by soil (Alloway, 1990). Due to low vapor pressures, most nickel compounds are not expected to volatilize from moist or dry soil surfaces, with one notable exception being nickel carbonyl (Ohe, 1976; HSDB). The concentration of nickel in plants generally reflects the concentration in soil although the relationship is more related to soluble and exchanged forms of nickel. Factors that increase solubility and



exchangeability of nickel in soils also result in an increase of the element in plant tissue (Alloway, 1990).

In plants nickel is necessary for healthy growth and is essential for metabolic processes (Alloway, 1990; NRC, 2005). Nickel is generally not accepted an essential trace element for mammals and birds as there is no clearly defined biochemical function. Under laboratory experimental conditions nickel deprivation can result in adverse effects including growth depression, impaired reproduction, and other biochemical changes (NRC, 2005).

The Eco-SSL values derived to date for nickel are summarized in Table 2.1.

Table 2.1 Nickel Eco-SSLs (mg/kg dry weight in soil)			
Plants	Soil Invertebrates	Wildlife	
		Avian	Mammalian
38	280	210	130

Eco-SSL values were derived for plants, soil invertebrates, avian and mammalian wildlife. The Eco-SSL values for nickel range from 38 mg/kg dry weight (dw) for plants to 280 mg/kg dw for soil invertebrates. All values are higher than the 95th percentile of reported background soil concentrations in eastern and western U.S. soils (Figure 2.1).

3.0 ECO-SSL FOR TERRESTRIAL PLANTS

Of the papers identified from the literature search process, 252 papers were selected for acquisition for further review. Of those papers acquired, 26 met all 11 Study Acceptance Criteria (U.S. EPA, 2003; Attachment 3-1). Each of these papers were reviewed and the studies were scored according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 3-2). Thirty-nine study results received an Evaluation Score greater than ten (U.S. EPA, 2003; Attachment 3-1). These studies are listed in Table 3.1.

The studies in Table 3.1 are sorted by bioavailability score. There are 11 studies eligible for Eco-SSL derivation with a bioavailability score of two. These results are used to derive the plant Eco-SSL for nickel (U.S. EPA, 2003; Attachment 3-2). The Eco-SSL is the geometric mean of the maximum acceptable toxicant concentration (MATC) and 20% effective concentration (EC_{20}) values for 6 species under different test conditions (pH and % organic matter (OM)) and is equal to 38 mg/kg dw.

Table 3.1 Plant Toxicity Data - Nickel

Reference	IP Number	Study ID	Test Organism		Soil pH	OM %	Bio-availability Score	ERE	Tox Parameter	Tox Value-Soil Conc. (mg/kg dw)	Total Evaluation Score	Eligible for Eco-SSL Derivation?	Used for Eco-SSL?
TN&Associates Inc., 2000	56444	G	Alfalfa	<i>Medicago sativa</i>	5.0	5.0	2	GRO	EC ₂₀	17	18	Y	Y
TN&Associates Inc., 2000	56444	H	Barley	<i>Hordeum vulgare</i>	5.0	5.0	2	GRO	EC ₂₀	7	18	Y	Y
TN&Associates Inc., 2000	56444	I	Brassica	<i>Brassica rapa</i>	5.0	5.0	2	GRO	EC ₂₀	10	18	Y	Y
TN&Associates Inc., 2000	56444	J	Alfalfa	<i>Medicago sativa</i>	6.3	0.1	2	GRO	EC ₂₀	25	18	Y	Y
TN&Associates Inc., 2000	56444	K	Barley	<i>Hordeum vulgare</i>	6.3	0.1	2	GRO	EC ₂₀	65	18	Y	Y
TN&Associates Inc., 2000	56444	L	Brassica	<i>Brassica rapa</i>	6.3	0.1	2	GRO	EC ₂₀	32	18	Y	Y
Dixon, 1988	7450	--	Red Oak	<i>Quercus rubra</i>	6.0	1.5	2	GRO	MATC	32	16	Y	Y
Taylor and Allinson, 1981	2790	B	Alfalfa	<i>Medicago sativa</i>	6.9	1.7	2	GRO	MATC	177	15	Y	Y
Khalid and Tinsley, 1980	7481	--	Ryegrass	<i>Lelium perenne</i>	4.7	4.5	2	GRO	MATC	52	15	Y	Y
Taylor, 1974	9645	--	Alfalfa	<i>Medicago sativa</i>	6.4	1.0	2	GRO	MATC	177	11	Y	Y
Halstead et al, 1969	7465	B	Oat	<i>Avena sativa</i>	6.1	1.4	2	REP	MATC	71	13	Y	Y
Geometric Mean										38			
Data Not Used to Derive Plant Eco-SSL													
Narwal et al, 1996	11707	A	Corn	<i>Zea mays</i>	8.0	0.3	1	GRO	MATC	141	15	Y	N
Narwal et al, 1996	11707	B	Corn	<i>Zea mays</i>	7.8	0.2	1	GRO	MATC	71	15	Y	N
Narwal et al, 1996	11707	C	Corn	<i>Zea mays</i>	7.7	1.2	1	GRO	MATC	71	15	Y	N
Taylor and Allinson, 1981	2790	A	Alfalfa	<i>Medicago sativa</i>	6.9	4.8	1	GRO	MATC	177	14	Y	N
Halstead et al, 1969	7465	C	Oat	<i>Avena sativa</i>	5.7	4.1	1	REP	MATC	224	12	Y	N
Halstead et al, 1969	7465	F	Alfalfa	<i>Medicago sativa</i>	5.7	4.1	1	GRO	MATC	32	12	Y	N
Halstead et al, 1969	7465	D	Alfalfa	<i>Medicago sativa</i>	7.8	4.0	0	GRO	MATC	224	11	Y	N
Tikhomirov et al, 1988	4757	B	Oat	<i>Avena sativa</i>	4.6	2.5	2	GRO	LOAEC	100	14	N	N
Singh and Jeng, 1993	12400	--	Ryegrass	<i>Lelium perenne</i>	6.0	0.7	2	GRO	NOAEC	50	14	N	N
Halstead et al, 1969	7465	E	Alfalfa	<i>Medicago sativa</i>	6.1	1.4	2	GRO	LOAEC	20	13	N	N
Wallace et al, 1977	57304	A	Bush Beans	<i>Phaseolus vulgaris</i>	4.0	2.8	2	GRO	NOAEC	250	12	N	N
Rehab and Wallace, 1978	7523	A	Cotton	<i>Gossypium barbadense</i>	6.8	2.2	1	GRO	LOAEC	100	12	N	N
Metwally and Rabie, 1989	1536	D	Corn	<i>Zea mays</i>	8.2	1.2	1	GRO	MATC	57	11	N	N
Wallace et al, 1977	57304	B	Bush Beans	<i>Phaseolus vulgaris</i>	5.8	2.8	1	GRO	LOAEC	100	11	N	N
Elmosly and Abdel-Sabour, 1997	4094	A	Red Clover	<i>Trifolium pratense L.</i>	8.2	1.0	1	GRO	NOAEC	100	14	N	N
Elmosly and Abdel-Sabour, 1997	4094	B	Red Clover	<i>Trifolium pratense L.</i>	7.9	0.6	1	GRO	NOAEC	100	14	N	N
Elmosly and Abdel-Sabour, 1997	4094	C	Red Clover	<i>Trifolium pratense L.</i>	7.6	0.1	1	GRO	NOAEC	100	14	N	N
Tikhomirov et al, 1988	4757	A	Oat	<i>Avena sativa</i>	5.9	2.9	1	GRO	LOAEC	400	13	N	N

Table 3.1 Plant Toxicity Data - Nickel

Reference	IP Number	Study ID	Test Organism		Soil pH	OM %	Bio-availability Score	ERE	Tox Parameter	Tox Value-Soil Conc. (mg/kg dw)	Total Evaluation Score	Eligible for Eco-SSL Derivation?	Used for Eco-SSL?
Dang et al, 1990	12906	A	Onion	<i>Allium cepa</i>	8.3	0.5	1	GRO	LOAEC	50	11	N	N
Dang et al, 1990	12906	B	Fenugreek	<i>Trigonella foenum</i>	8.3	0.5	1	GRO	LOAEC	50	11	N	N
Metwally and Rabie, 1989	1536	C	Fava Bean	<i>Vicia faba</i>	8.2	1.2	1	GRO	NOAEC	50	11	N	N
Wallace et al, 1977	57304	C	Barley	<i>Hordeum vulgare</i>	5.8	2.8	1	GRO	MATC	50	11	N	N
Genovese, 1978	58147	B	Jack Pine	<i>Pinus banksiana</i>	7.7	0.1	1	GRO	NOAEC	150	11	N	N
Genovese, 1978	58147	D	Black Spruce	<i>Picea mariana</i>	7.7	0.1	1	GRO	NOAEC	150	11	N	N
Halstead et al, 1969	7465	A	Oat	<i>Avena sativa</i>	7.8	4.0	0	REP	NOAEC	500	11	N	N
Gupta et al, 1996	13146	A	Chickpea	<i>Cicer arietinum</i>	8.0	2.0	0	GRO	MATC	6	11	N	N
Gupta et al, 1996	13146	B	Lentil	<i>Lens esculentum</i>	8.0	2.0	0	GRO	MATC	4	11	N	N
Gupta et al, 1996	13146	C	Mustard	<i>Brassica juncea</i>	8.0	2.0	0	GRO	MATC	14	11	N	N

EC₂₀ = Effect concentration for 20% of test population

ERE = Ecologically relevant endpoint

GRO = Growth

LOAEC = Lowest observed adverse effect concentration

MATC = Maximum acceptable toxicant concentration. Geometric mean of NOAEC and LOAEC.

N = No

NOAEC = No observed adverse effect concentration

-- Study ID number not assigned

OM = Organic matter content

PHY = Physiology

REP = Reproduction

Y = yes

Bioavailability Score described in *Guidance for Developing Eco-SSLs* (U.S. EPA, 2003)

Total Evaluation Score described in *Guidance for Developing Eco-SSLs* (U.S. EPA, 2003)

4.0 ECO-SSL FOR SOIL INVERTEBRATES

Of the papers identified from the literature search process, 46 papers were selected for acquisition for further review. Of those papers acquired, 9 met all 11 Study Acceptance Criteria (U.S. EPA 2003; Attachment 3-1). Each of these papers were reviewed and the studies were scored according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 3-2). Twelve studies received an Evaluation Score greater than ten. These studies are listed in Table 4.1. The studies in Table 4.1 are sorted by bioavailability score. There are five studies eligible for Eco-SSL derivation. The Eco-SSL is the geometric mean of the maximum acceptable toxicant concentrations (MATCs) for five species under different test conditions (pH and % organic matter (OM)) and is equal to 280 mg/kg dw.

5.0 ECO-SSL FOR AVIAN WILDLIFE

The derivation of the Eco-SSL for avian wildlife was completed as two parts. First, the toxicity reference value (TRV) was derived according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-5). Second, the Eco-SSL (soil concentration) was back-calculated for each surrogate species representing different trophic levels based on the wildlife exposure model and the TRV (U.S. EPA, 2003).

5.1 Avian TRV

The literature search completed according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-1) identified 1,169 papers with possible toxicity data for either avian or mammalian species. Of these studies, 1,101 were rejected for use as described in Section 7.5. Of the remaining studies, 11 contained data for avian test species. These papers were reviewed and the data were extracted and scored according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-3 and 4-4). The results of the data extraction and review are provided as Table 5.1. The complete results are included as Appendix 5-1.

Within the reviewed papers, there are 28 results for biochemical (BIO), behavior (BEH), physiology (PHY), pathology (PTH), reproduction (REP), growth (GRO), and survival (MOR) effects that meet the Data Evaluation Score of >65 for use to derive the TRV (U.S. EPA, 2003; Attachment 4-4). These data are plotted in Figure 5.1 and correspond directly with the data presented in Table 5.1. The no-observed adverse effect level (NOAEL) results for growth and reproduction are used to calculate a geometric mean. This result is examined in relationship to the lowest bounded lowest-observed adverse effect level (LOAEL) for reproduction, growth, and survival to derive the TRV according to procedures in the Eco-SSL guidance (U.S. EPA, 2003); Attachment 4-5).

A geometric mean of the NOAEL values for reproduction and growth was calculated at 6.71 mg nickel/kg bw/day. This value is lower than the lowest bounded LOAEL for reproduction, growth, or survival. Therefore, the TRV is equal to the geometric mean of the NOAEL values for reproduction and growth and is equal to 6.71 mg nickel/kg bw/day.

Table 4.1 Invertebrate Toxicity Data - Nickel

Reference	IP Number	Study ID	Test Organism		Soil pH	OM %	Bio-availability Score	ERE	Tox Parameter	Tox Value-Soil Conc. (mg/kg dw)	Total Evaluation Score	Eligible for Eco-SSL Derivation?	Used for Eco-SSL?
Scott-Fordsmand et al, 1999	15163	--	Springtail	<i>Folsomia fimetaria L.</i>	5.8	4.0	1	REP	MATC	387	16	Y	Y
Scott-Fordsmand et al, 1998	18892	a	Earthworm	<i>Eisenia veneta</i>	5.8	4.0	1	REP	MATC	173	13	Y	Y
Lock and Janssen, 2002	66195	a	Earthworm	<i>Eisenia fetida</i>	6.0	10.0	1	REP	MATC	240	14	Y	Y
Lock and Janssen, 2002	66195	b	Earthworm	<i>Enchytraeus albidus</i>	6.0	10.0	1	REP	MATC	240	14	Y	Y
Lock and Janssen, 2002	66195	c	Springtail	<i>Folsomia candida</i>	6.0	10.0	1	REP	MATC	423	14	Y	Y
Geometric Mean										280			

Data Not Used to Derive Invertebrate Eco-SSL

Peredney and Williams, 2000b	56449	m	Roundworm	<i>Caenorhabditis elegans</i>	4	1.14	2	MOR	LC ₅₀	144	13	N	N
Peredney and Williams, 2000b	56449	n	Roundworm	<i>Caenorhabditis elegans</i>	4	1.14	2	MOR	LC ₅₀	44	13	N	N
Peredney and Williams, 2000b	56449	o	Roundworm	<i>Caenorhabditis elegans</i>	4	4.2	2	MOR	LC ₅₀	387	13	N	N
Peredney and Williams, 2000b	56449	p	Roundworm	<i>Caenorhabditis elegans</i>	4	4.2	2	MOR	LC ₅₀	165	13	N	N
Neuhauser et al, 1985	6812	--	Earthworm	<i>E. fetida</i>	6.0	10.0	1	MOR	LC ₅₀	242	11	N	N
Korthals et al, 1996	4402	--	Nematodes	nematoda (various sp.)	4.1	1.9	1	POP	LOAEC	100	14	N	N
Peredney and Williams, 2000a	53082	--	Roundworm	<i>Caenorhabditis elegans</i>	4	10	1	MOR	LC ₅₀	797	12	N	N
Peredney and Williams, 2000b	56449	q	Roundworm	<i>Caenorhabditis elegans</i>	4	10	1	MOR	LC ₅₀	800	12	N	N
Peredney and Williams, 2000b	56449	r	Roundworm	<i>Caenorhabditis elegans</i>	4	10	1	MOR	LC ₅₀	348	12	N	N
Neuhauser et al, 1986	17707	--	Earthworm	<i>E. fetida</i>	6.0	10.0	0	MOR	LC ₅₀	242	14	N	N

ERE = Ecologically relevant endpoint

LC₅₀ = Concentration lethal to 50% of test population

LOAEC = Lowest observed adverse effect concentration

MATC = Maximum acceptable toxicant concentration

MOR = Mortality

N = No

-- Study ID not assigned

NOAEC = No observed adverse effect concentration

OM = Organic matter content

POP = Population

REP = Reproduction

Y = Yes

Bioavailability Score described in *Guidance for Developing Eco-SSLs* (U.S. EPA, 2003)

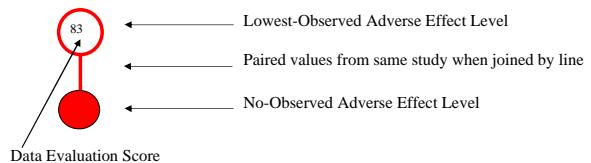
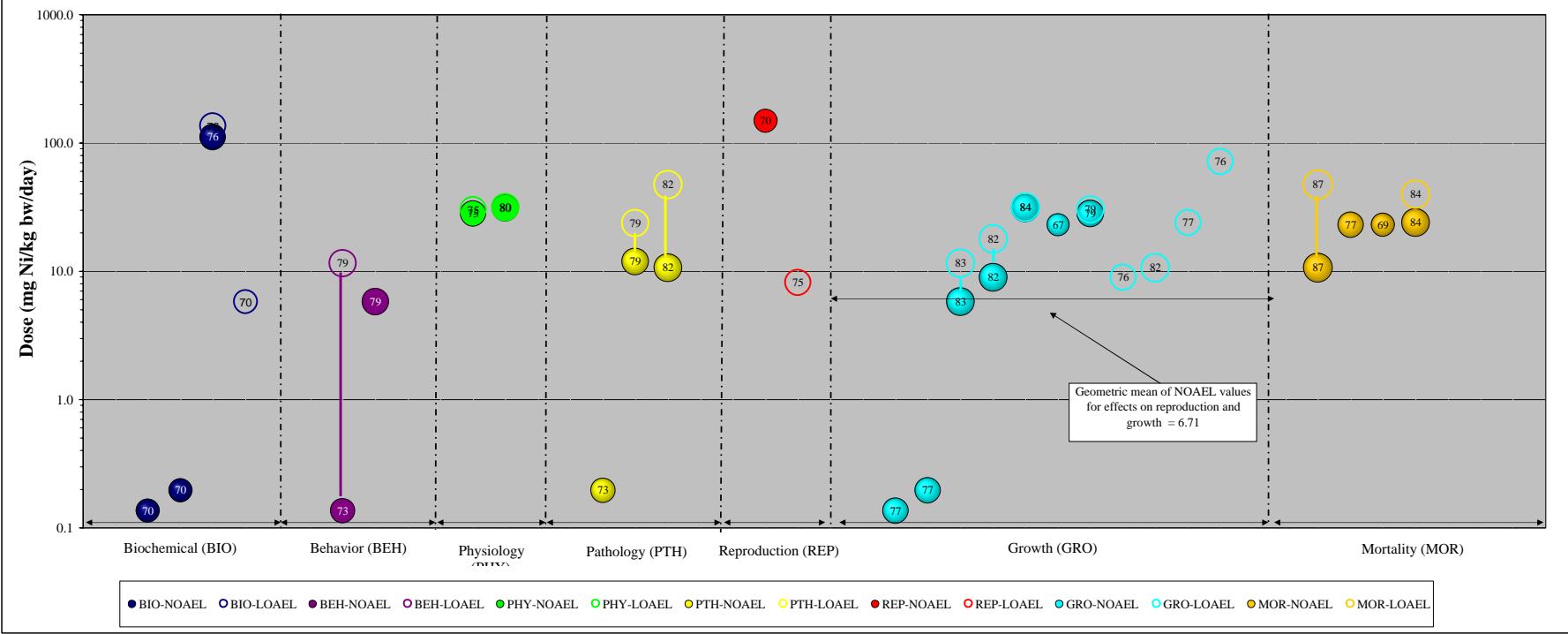
Total Evaluation Score described in *Guidance for Developing Eco-SSLs* (U.S. EPA, 2003)

Table 5.1
Avian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)
Nickel

Result #	Reference	Ref No.	Test Organism	# of Conc/ Doses	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Effect Type	Effect Measure	Response Site	NOAEL Dose (mg/kg bw/day)	LOAEL Dose (mg/kg bw/day)	Total
Biochemical (BIO)																		
1	Oscar and Mitchell, 1995	5407	Chicken (<i>Gallus domesticus</i>)	3	U	FD	28	d	21	d	JV	F	CHM	LIPD	WO	0.136		70
2	Nielsen et al, 1975	6885	Chicken (<i>Gallus domesticus</i>)	4	U	FD	3	w	1	d	JV	M	CHM	HMCT	BL	0.195		70
3	Ling and Leach, 1979	6666	Chicken (<i>Gallus domesticus</i>)	6	U	FD	3	w	1	d	JV	M	CHM	HMGL	BL	110	135	76
4	Martinez and Diaz, 1996	5345	Chicken (<i>Gallus domesticus</i>)	4	U	FD	42	d	1	d	JV	M	CHM	HMGL	BL		5.76	70
Behavior (BEH)																		
5	Oscar and Mitchell, 1995	5407	Chicken (<i>Gallus domesticus</i>)	3	U	FD	28	d	21	d	JV	F	FDB	FCNS	WO	0.136		73
6	Martinez and Diaz, 1996	5345	Chicken (<i>Gallus domesticus</i>)	4	U	FD	42	d	1	d	JV	M	FDB	FCNS	WO	5.76	11.5	79
Physiology (PHY)																		
7	Weber and Reid, 1968	60	Chicken (<i>Gallus domesticus</i>)	8	U	FD	4	w	1	d	JV	B	PHY	FDCV	WO	28.3	30.2	75
8	Weber and Reid, 1968	60	Chicken (<i>Gallus domesticus</i>)	8	U	FD	4	w	1	d	JV	B	PHY	META	WO	31.0	31.5	80
Pathology (PTH)																		
9	Nielsen et al, 1975	6885	Chicken (<i>Gallus domesticus</i>)	4	U	FD	3	w	1	d	JV	M	ORW	SMIX	SK	0.195		73
10	Cain and Pafford, 1981	6461	Duck (<i>Anas platyrhynchos</i>)	4	M	FD	28	d	1	d	JV	B	ITX	ATAX	WO	10.7	47.0	82
11	Martinez and Diaz, 1996	5345	Chicken (<i>Gallus domesticus</i>)	4	U	FD	42	d	1	d	JV	M	HIS	GHIS	HE	11.9	23.8	79
Reproduction (REP)																		
12	Eastin and O'Shea, 1981	6492	Duck (<i>Anas platyrhynchos</i>)	5	U	FD	90	d	20	mo	LB	F	REP	HTCH	EG	149		70
13	Meluzzi et al, 1996	2771	Chicken (<i>Gallus domesticus</i>)	4	U	FD	60	d	22	w	LB	F	EGG	ESWT	EG		8.16	75
Growth (GRO)																		
14	Oscar et al, 1995	5407	Chicken (<i>Gallus domesticus</i>)	3	U	FD	28	d	21	d	JV	F	GRO	BDWT	WO	0.136		77
15	Nielsen et al, 1975	6885	Chicken (<i>Gallus domesticus</i>)	4	U	FD	3	w	1	d	JV	M	GRO	BDWT	WO	0.195		77
16	Martinez and Diaz, 1996	5345	Chicken (<i>Gallus domesticus</i>)	4	U	FD	42	d	1	d	JV	M	GRO	BDWT	WO	5.76	11.5	83
17	Hill, 1979	397	Chicken (<i>Gallus domesticus</i>)	3	U	FD	5	w	1	d	JV	F	GRO	BDWT	WO	8.95	17.9	82
18	Blalock and Hill, 1985	36697	Chicken (<i>Gallus domesticus</i>)	2	U	FD	25	d	1	d	JV	B	GRO	BDWT	WO	22.9		67
19	Weber and Reid, 1968	60	Chicken (<i>Gallus domesticus</i>)	8	U	FD	4	w	1	d	JV	B	GRO	BDWT	WO	28.3	30.2	79
20	Weber and Reid, 1968	60	Chicken (<i>Gallus domesticus</i>)	8	U	FD	4	w	1	d	JV	B	GRO	BDWT	WO	31.0	31.5	84
21	Hill, 1979	397	Chicken (<i>Gallus domesticus</i>)	3	U	FD	5	w	1	d	JV	F	GRO	BDWT	WO		8.95	76
22	Cain and Pafford, 1981	6461	Duck (<i>Anas platyrhynchos</i>)	4	M	FD	60	d	1	d	JV	F	MPH	HULT	HM		10.7	82
23	Ling and Leach, 1979	6666	Chicken (<i>Gallus domesticus</i>)	6	U	FD	3	w	1	d	JV	M	GRO	BDWT	WO		23.9	77
24	Hill, 1985	36708	Chicken (<i>Gallus domesticus</i>)	2	U	FD	19	d	NR	NR	JV	B	GRO	BDWT	WO		71.8	76
Survival (MOR)																		
25	Cain and Pafford, 1981	6461	Duck (<i>Anas platyrhynchos</i>)	4	M	FD	30	d	1	d	JV	B	MOR	SURV	WO	10.7	47.0	87
26	Blalock and Hill, 1985	36697	Chicken (<i>Gallus domesticus</i>)	2	U	FD	25	d	1	d	JV	B	MOR	MORT	WO	22.9		77
27	Martinez and Diaz, 1996	5345	Chicken (<i>Gallus domesticus</i>)	4	U	FD	42	d	1	d	JV	M	MOR	MORT	WO	23.0		69
28	Ling and Leach, 1979	6666	Chicken (<i>Gallus domesticus</i>)	6	U	FD	3	w	1	d	JV	M	MOR	SURV	WO	23.9	39.9	84

AD = adult; ATAX = ataxia; B = both; BDWT = body weight changes; BEH = behavior; BIO = biochemical effects; BL = blood; bw = body weight; CHM = chemical changes; d = day; EGG = effects on eggs; EG = egg; ESWT = eggshell weight; F = female; FCNS = food consumption; FD = food; FDB = feeding behavior; FDCV = food conversion efficiency; GHIS = general histology; GRO = growth; HE = heart; HIS = histological changes; HM = humerus; HMCT = hematocrit; HMGL = hemoglobin; HTCH = hatch; HULT = humerus length; ITX = intoxication; JV = juvenile; kg = kilograms; LB = egg laying bird; LIPD = lipid; LOAEL = lowest observed adverse effect level; M = male; M = measured; META = metabolic rate; mo = months; MOR = effects on mortality and survival; MORT = mortality; MPH = morphology effects; NOAEL = No Observed Adverse Effect Level; NR = Not reported; ORW = organ weight changes; ORWT = organ weight changes; PHY = physiology; PTH = pathology; REP = reproduction; SK = skin, epidermis; SMIX = weight relative to body weight; SURV = survival; U = unmeasured; w = weeks; WO = whole organism.

Figure 5.1 Avian TRV Derivation for Nickel



Wildlife TRV Derivation Process

- 1) There are at least three results available for two test species within the growth, reproduction, and mortality effect groups.
There are enough data to derive a TRV.
- 2) There are at least three NOAEL results available within the growth and reproduction effect groups for calculation of a geometric mean.
- 3) The geometric mean is equal to 6.71 mg nickel/kg bw/d and is lower than the lowest bounded LOAEL for results within the reproduction, growth, and survival (MOR) effect groups.
- 3) The avian wildlife TRV for nickel is equal to 6.71 mg nickel/kg bw/day which is the geometric mean of NOAEL values for effects on reproduction and growth.

5.2 Estimation of Dose and Calculation of the Eco-SSL

Two separate Eco-SSL values were calculated for avian wildlife, one for each of two surrogate receptor species representing different trophic levels. The avian Eco-SSLs were calculated according to the Eco-SSL guidance (U.S. EPA, 2003) and are summarized in Table 5.2. An Eco-SSL value could not be calculated for the avian ground insectivore (woodcock) as a reliable method to estimate uptake of nickel from soil into soil invertebrates could not be identified (see Attachment 4-1 of the Eco-SSL guidance; U.S. EPA, 2003).

Table 5.2 Calculation of the Avian Eco-SSLs for Nickel					
Surrogate Receptor Group	TRV for Nickel (mg dw/kg bw/d) ¹	Food Ingestion Rate (FIR) ² (kg dw/kg bw/d)	Soil Ingestion as Proportion of Diet (P_s) ²	Concentration of Nickel in Biota Type (i) ^{2,3} (B_i) (mg/kg dw)	Eco-SSL (mg/kg dw) ⁴
Avian herbivore (dove)	6.71	0.190	0.139	$\ln(B_i) = 0.748 * \ln(Soil_j) - 2.223$ where i = plants	210
Avian ground insectivore (woodcock)	6.71	0.214	0.164	NA	NA
Avian carnivore (hawk)	6.71	0.0353	0.057	$\ln(B_i) = 0.4658 * \ln(Soil_j) - 0.2462$ where i = mammals	2800

¹ The process for derivation of wildlife TRVs is described in Attachment 4-5 of U.S. EPA (2003).
² Parameters (FIR, P_s , B_i values, regressions) are provided in U.S. EPA (2003) Attachment 4-1 (revised February 2005).
³ B_i = Concentration in biota type (i) which represents 100% of the diet for the respective receptor.
⁴ HQ = [FIR * (Soil_j * P_s + B_i)] / TRV solved for HQ=1 where Soil_j = Eco-SSL (Equation 4-2; U.S. EPA, 2003).
NA = Not Available.

6.0 ECO-SSL FOR MAMMALIAN WILDLIFE

The derivation of the Eco-SSL for mammalian wildlife was completed as two parts. First, the TRV was derived according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-5). Second, the Eco-SSL (soil concentration) was back-calculated for each surrogate receptor species based on the wildlife exposure model and the TRV (U.S. EPA, 2003).

6.1 Mammalian TRV

The literature search was completed according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-2) and identified 1,169 papers with possible toxicity data for nickel for either avian or mammalian species. Of these studies, 1,101 were rejected for use as described in Section 7.5. Of the remaining papers, 52 contained data for mammalian test species. These papers were reviewed and the data were extracted and scored according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-3 and 4-4). The results of the data extraction and review are summarized in Table 6.1. The complete results are provided as Appendix 6-1.

Table 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)

Nickel

Page 1 of 3

Result #	Reference	Ref No.	Test Organism	# of Conc/ Doses	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Effect Type	Effect Measure	Response Site	NOAEL Dose*	(mg/kg bw/day)	LOAEL Dose*	(mg/kg bw/day)	Total
Biochemical (BIO)																				
1	Spears et al, 1984	19671	Rat (<i>Rattus norvegicus</i>)	3	U	FD	21	d	1	d	JV	NR	ENZ	ALPH	SR	0.265	1.32	74		
2	Chatterjee et al, 1979	19098	Rat (<i>Rattus norvegicus</i>)	2	U	FD	21	d	NR	NR	JV	M	CHM	ASCA	LI	0.335		74		
3	Nielsen, 1980	19478	Rat (<i>Rattus norvegicus</i>)	3	U	FD	10	w	NR	NR	JV	F	CHM	HMCT	PL	0.456	4.56	73		
4	Pandey et al, 1999	19521	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	ENZ	LADH	TE	1.35	2.71	83		
5	Spears and Hatfield, 1985	19666	Rat (<i>Rattus norvegicus</i>)	3	U	FD	21	d	NR	NR	JV	M	CHM	LEUK	BL	1.47	22.0	72		
6	O'Dell et al, 1970	14477	Cattle (<i>Bos taurus</i>)	4	U	FD	8	w	14	w	JV	M	CHM	CALC	NR	1.63	6.53	75		
7	Kadiiska et al, 1985	19290	Rat (<i>Rattus norvegicus</i>)	2	U	DR	30	d	NR	NR	JV	M	ENZ	P450	LI	4.47		67		
8	Obone et al, 1999	19507	Rat (<i>Rattus norvegicus</i>)	4	U	DR	13	w	NR	NR	JV	M	CHM	PRTL	PL	4.70	11.7	71		
9	ODell et al, 1970	14476	Cattle (<i>Bos taurus</i>)	3	U	FD	6	w	NR	NR	LC	F	CHM	PRTL	MK	6.75		70		
10	Smith et al, 1993	62	Rat (<i>Rattus norvegicus</i>)	4	M	DR	26	w	62	d	LC	F	CHM	GBCM	PL	6.80	31.6	78		
11	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	3	M	FD	28	d	31	d	JV	M	CHM	HMGL	BL	7.78		75		
12	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	2	M	FD	28	d	31	d	JV	M	CHM	HMGL	BL	8.20		75		
13	Whanger, 1973	63	Rat (<i>Rattus norvegicus</i>)	4	U	FD	6	w	35	d	JV	NR	CHM	HMGL	BL	9.49	47.4	69		
14	Nation et al, 1985	19460	Rat (<i>Rattus norvegicus</i>)	3	U	FD	14	d	80	d	JV	M	CHM	GBCM	LI	20.0		74		
15	Uthus and Poellot, 1997	19764	Rat (<i>Rattus norvegicus</i>)	2	U	FD	57	d	NR	NR	JV	M	ENZ	GENZ	LI		0.0844	70		
16	Spears et al, 1986	19664	Cattle (<i>Bos taurus</i>)	2	U	FD	28	d	50	d	JV	M	CHM	AMMO	GT		0.101	71		
17	Milne et al, 1990	19437	Sheep (<i>Ovis aries</i>)	2	U	GV	4	w	5	mo	JV	F	ENZ	GENZ	SH	0.147	73			
18	Nielsen et al, 1979	19461	Rat (<i>Rattus norvegicus</i>)	3	U	FD	9	w	21	d	JV	F	CHM	HMCT	BL	0.456	69			
19	Pandey and Singh, 1999	19522	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	ENZ	LADH	TE	0.797	72			
20	Spears and Hatfield, 1985	19666	Rat (<i>Rattus norvegicus</i>)	2	U	FD	42	d	NR	NR	JV	M	CHM	HMGL	BL	2.63	70			
21	Tandon and Mathur, 1986	19746	Rat (<i>Rattus norvegicus</i>)	2	U	GV	4	w	NR	NR	JV	M	ENZ	MADH	KI	3.00	77			
22	Vyskocil et al, 1994	19787	Rat (<i>Rattus norvegicus</i>)	2	U	DR	6	mo	10	w	JV	M	CHM	ALBM	UR	6.25	66			
23	Novelli et al, 1998	19496	Rat (<i>Rattus norvegicus</i>)	2	U	DR	2	mo	15	w	JV	M	ENZ	AATT	SR	12.5	66			
24	Waltschewa et al, 1972	19792	Rat (<i>Rattus norvegicus</i>)	2	U	GV	120	d	5	mo	JV	M	ENZ	GENZ	TE	25.0	77			
25	Dieter et al, 1988	61	Mouse (<i>Mus musculus</i>)	4	UX	DR	180	d	6-8	w	JV	F	ENZ	G6PD	BM	43.3	71			
26	Mathur, 1987	19416	Rat (<i>Rattus norvegicus</i>)	2	U	FD	3	w	3	w	JV	M	ENZ	GOTR	LI	44.5	69			
27	Weber and Reid, 1969	14485	Mouse (<i>Mus musculus</i>)	3	U	FD	4	w	NR	NR	JV	B	ENZ	CCOX	LI		179	66		
Behavior (BEH)																				
28	Spears et al, 1986	19664	Cattle (<i>Bos taurus</i>)	2	U	FD	140	d	50	d	JV	M	FDB	FCNS	WO	0.101		72		
29	O'Dell et al, 1970	14477	Cattle (<i>Bos taurus</i>)	4	U	FD	8	w	14	w	JV	M	FDB	FCNS	WO	1.63	6.53	78		
30	Gershbein et al 1983	136	Rat (<i>Rattus norvegicus</i>)	2	U	FD	80	d	44	d	JV	M	BEH	NMVM	WO	5.89		66		
31	Vyskocil et al, 1994	19787	Rat (<i>Rattus norvegicus</i>)	2	U	DR	6	mo	10	w	JV	M	FDB	WCON	WO	6.25		69		
32	Smith et al, 1993	62	Rat (<i>Rattus norvegicus</i>)	4	M	DR	11	w	62	d	LC	F	FDB	WCON	WO	6.80	31.63	81		
33	O'Dell et al, 1971	14479	Cattle (<i>Bos taurus</i>)	4	UX	FD	8	w	13-21	w	JV	M	FDB	FCNS	WO	7.00	14.6	88		
34	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	3	M	FD	28	d	31	d	JV	M	FDB	FCNS	WO	7.78		78		
35	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	2	M	FD	28	d	31	d	JV	M	FDB	FCNS	WO	8.20		78		
36	Nation et al, 1985	19460	Rat (<i>Rattus norvegicus</i>)	3	U	FD	14	d	80	d	JV	M	BEH	ACTP	WO	10.0	20.0	83		
37	Alexander et al, 1978	36694	Meadow vole (<i>Microtus pennsylvanicus</i>)	4	M	FD	49	d	14	d	JV	NR	FDB	FCNS	WO	29.4	309	81		
38	Dieter et al, 1988	61	Mouse (<i>Mus musculus</i>)	4	UX	DR	180	d	6-8	w	JV	F	FDB	WCON	WO	43.3	107	80		
39	Weber and Reid, 1969	14485	Mouse (<i>Mus musculus</i>)	3	U	FD	4	w	NR	NR	JV	M	FDB	FCNS	WO	179	265	75		
40	Cempel and Janicka, 2002	36331	Rat (<i>Rattus norvegicus</i>)	3	U	DR	90	d	NR	NR	JV	M	FDB	WCON	WO		37.6	69		
Physiology (PHY)																				
41	Szakmary et al, 1995	19729	Rat (<i>Rattus norvegicus</i>)	2	U	GV	8	d	NR	NR	GE	F	PHY	BLPR	HE	1.36		80		
42	O'Dell et al, 1971	14479	Cattle (<i>Bos taurus</i>)	4	UX	FD	8	w	13	w	JV	M	PHY	EXCR	UR	1.80	7.00	86		
43	Obone et al, 1999	19507	Rat (<i>Rattus norvegicus</i>)	4	U	DR	13	w	NR	NR	JV	M	PHY	EXCR	UR	4.70	11.7	74		
44	O'Dell et al, 1970	14477	Cattle (<i>Bos taurus</i>)	4	U	FD	8	w	14	w	JV	M	PHY	DIFD	WO	6.55		74		
45	ODell et al, 1970	14476	Cattle (<i>Bos taurus</i>)	3	U	FD	6	w	NR	NR	LC	F	PHY	GPHY	MK	6.75		73		
Pathology (PTH)																				
46	Chatterjee et al, 1979	19098	Rat (<i>Rattus norvegicus</i>)	2	U	FD	21	d	NR	NR	JV	M	ORW	SMIX	LI	0.335		77		
47	Nielsen, 1980	19478	Rat (<i>Rattus norvegicus</i>)	3	U	FD	10	w	NR	NR	JV	F	ORW	SMIX	LI	0.456	4.56	76		
48	O'Dell et al, 1970	14477	Cattle (<i>Bos taurus</i>)	4	U	FD	8	w	14	w	JV	M	ORW	ORWT	LU	1.64	6.55	78		
49	Ambrose et al, 1976	14474	Rat (<i>Rattus norvegicus</i>)	4	U	FD	24	mo	28	d	JV	B	ORW	SMIX	HE	8.06	80.6	77		
50	Ambrose et al, 1976	14474	Dog (<i>Canis familiaris</i>)	4	U	FD	24	mo	6	mo	JV	B	ORW	SMIX	KI	45.0	112	78		
51	Schroeder and Mitchener, 1975	1858	Mouse (<i>Mus musculus</i>)	2	U	DR	520	d	19-20	d	JV	M	HIS	EDMA	WO	0.705	68			
52	Pandey et al, 1999	19521	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	ORW	ORWT	PG	1.35	80			
53	Kakela et al, 1999	19293	Rat (<i>Rattus norvegicus</i>)	3	U	DR	62	d	3.5-9	mo	GE	F	ORW	SMIX	LI	3.40	68			
54	Obone et al, 1999	19507	Rat (<i>Rattus norvegicus</i>)	4	U	DR	13	w	NR	NR	JV	M	ORW	ORWT	HE	4.70	68			
55	Schroeder et al, 1974	14484	Rat (<i>Rattus norvegicus</i>)	2	M	DR	1217	d	30	d	JV	B	HIS	GHIS	HE	5.44	71			
56	Vyskocil et al, 1994	19787	Rat (<i>Rattus norvegicus</i>)	2	U	DR	6	mo	10	w	JV	M	ORW	SMIX	KI	6.25	69			
57	Dieter et al, 1988	61	Mouse (<i>Mus musculus</i>)	4	UX	DR	180	d	6-8	w	JV	F	ORW	ORWT	LI		43.3	74		
58	Seidenberg et al 1986	113	Mouse (<i>Mus musculus</i>)	2	U	GV	4	d	NR	NR	GE	F	GRS	BDWT	WO	90.6	80			

Table 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)

Nickel

Page 2 of 3

Result #	Reference	Ref No.	Test Organism	# of Conc/ Doses	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Effect Type	Effect Measure	Response Site	NOAEL Dose*	(mg/kg bw/day)	LOAEL Dose*	(mg/kg bw/day)	Total
Reproduction (REP)																				
59	Kakela et al, 1999	19293	Rat (<i>Rattus norvegicus</i>)	4	U	DR	28	d	3.5-9	mo	SM	M	REP	GIDX	WO	1.10	3.31	80		
60	Pandey and Srivastava, 2000	36722	Mouse (<i>Mus musculus</i>)	4	U	OR	35	d	NR	mo	JV	M	REP	SPCL	SM	1.35	2.71	92		
61	Pandey and Srivastava, 2000	36722	Mouse (<i>Mus musculus</i>)	4	U	OR	35	d	NR	NR	JV	M	REP	SPCL	SM	1.70	3.40	92		
62	Phatak and Patwardhan, 1950	14480	Rat (<i>Rattus norvegicus</i>)	4	U	FD	4	mo	4	w	GE	F	REP	PRWT	WO	9.30		71		
63	Chernoff and Kavlock, 1982	1260	Mouse (<i>Mus musculus</i>)	2	U	GV	5	d	60	d	GE	F	REP	PROG	WO	45.3		86		
64	Berman and Rehnberg, 1983	19064	Mouse (<i>Mus musculus</i>)	3	U	DR	15	d	NR	NR	GE	F	REP	PRFM	WO	85.3	171	81		
65	Seidenberg et al 1986	113	Mouse (<i>Mus musculus</i>)	2	U	GV	4	d	NR	NR	GE	F	REP	PRWT	WO	90.6		82		
66	Ambrose et al, 1976	14474	Dog (<i>Canis familiaris</i>)	4	U	FD	24	mo	6	mo	JV	B	REP	TEWT	TE	112		74		
67	Ambrose et al, 1976	14474	Rat (<i>Rattus norvegicus</i>)	4	U	FD	118	d	28	d	GE	F	REP	PRWT	WO	164	327	84		
68	Ambrose et al, 1976	14474	Rat (<i>Rattus norvegicus</i>)	4	U	FD	2	yr	28	d	JV	M	REP	TEWT	TE	205		70		
69	Schroeder and Mitchener, 1971	66	Rat (<i>Rattus norvegicus</i>)	2	U	DR	9	mo	21	d	GE	F	REP	DEYO	WO		0.551	67		
70	Pandey and Singh, 1999	19522	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	REP	SPCV	SM		0.797	81		
71	Smith et al, 1993	62	Rat (<i>Rattus norvegicus</i>)	4	M	DR	23	w	62	d	LC	F	REP	DEYO	WO		1.33	83		
72	Pandey et al, 1999	19521	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	REP	TEWT	TE		1.35	86		
73	Pandey et al, 1999	19520	Mouse (<i>Mus musculus</i>)	2	U	GV	35	d	NR	NR	AD	M	REP	RSUC	WO		1.59	86		
74	Obone et al, 1999	19507	Rat (<i>Rattus norvegicus</i>)	4	U	DR	13	w	NR	NR	JV	M	REP	TEWT	TE		4.70	74		
75	Waltschewa et al, 1972	19792	Rat (<i>Rattus norvegicus</i>)	2	U	GV	120	d	5	mo	JV	M	REP	SPCL	SM		25.0	86		
Growth (GRO)																				
76	Uthus and Poellot, 1997	19764	Rat (<i>Rattus norvegicus</i>)	2	U	FD	57	d	NR	NR	JV	M	GRO	BDWT	WO	0.0844		77		
77	Spears et al, 1986	19664	Cattle (<i>Bos taurus</i>)	2	U	FD	140	d	50	d	JV	M	GRO	BDWT	WO	0.101		78		
78	Chatterjee et al, 1979	19098	Rat (<i>Rattus norvegicus</i>)	2	U	FD	21	d	NR	NR	JV	M	GRO	BDWT	WO	0.335		81		
79	Spears et al, 1984	19671	Rat (<i>Rattus norvegicus</i>)	3	U	FD	49	d	1	d	JV	NR	GRO	BDWT	WO	1.17		77		
80	Smith et al, 1993	62	Rat (<i>Rattus norvegicus</i>)	4	M	DR	15	w	62	d	LC	F	GRO	BDWT	WO	1.33	6.80	85		
81	Szakmary et al, 1995	19729	Rat (<i>Rattus norvegicus</i>)	2	U	GV	8	d	NR	NR	GE	F	GRO	BDWT	WO	1.36		84		
82	Spears and Hatfield, 1985	19666	Rat (<i>Rattus norvegicus</i>)	3	U	FD	21	d	NR	NR	JV	M	GRO	BDWT	WO	1.47	22.0	79		
83	O'Dell et al, 1970	14477	Cattle (<i>Bos taurus</i>)	4	U	FD	8	w	14	w	JV	M	GRO	BDWT	WO	1.64	6.55	82		
84	Cikrt et al, 1992	19109	Rat (<i>Rattus norvegicus</i>)	2	U	DR	90	d	NR	NR	JV	F	GRO	BDWT	WO	2.97		73		
85	Nielsen et al, 1979	19461	Rat (<i>Rattus norvegicus</i>)	3	U	FD	9	w	21	d	JV	F	GRO	BDWT	WO	4.56		67		
86	Nielsen, 1980	19479	Rat (<i>Rattus norvegicus</i>)	3	U	FD	10	w	NR	NR	JV	F	GRO	BDWT	WO	4.56		67		
87	Schroeder et al, 1974	14484	Rat (<i>Rattus norvegicus</i>)	2	M	DR	1217	d	30	d	JV	B	GRO	BDWT	WO	5.44		75		
88	Gershbein et al 1983	136	Rat (<i>Rattus norvegicus</i>)	2	U	FD	80	d	44	d	JV	M	GRO	BDWT	WO	5.89		68		
89	ODell et al, 1970	14476	Cattle (<i>Bos taurus</i>)	3	U	FD	6	w	NR	NR	LC	F	GRO	BDWT	WO	6.75		68		
90	O'Dell et al, 1971	14479	Cattle (<i>Bos taurus</i>)	4	UX	FD	8	w	13-21	w	JV	M	GRO	BDWT	WO	7.00	14.6	92		
91	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	3	M	FD	28	d	31	d	JV	M	GRO	BDWT	WO	7.78		82		
92	Ambrose et al, 1976	14474	Rat (<i>Rattus norvegicus</i>)	4	U	FD	6	w	28	d	JV	B	GRO	BDWT	WO	9.11	91.1	81		
93	Phatak and Patwardhan, 1950	14480	Rat (<i>Rattus norvegicus</i>)	4	U	FD	8	w	4	w	JV	B	GRO	BDWT	WO	9.30		69		
94	Whanger, 1973	63	Rat (<i>Rattus norvegicus</i>)	4	U	FD	6	w	35	d	JV	NR	GRO	BDWT	WO	9.49	47.4	76		
95	Kakela et al, 1999	19293	Rat (<i>Rattus norvegicus</i>)	3	U	DR	62	d	3.5-9	mo	GE	F	GRO	BDWT	WO	11.4		70		
96	Obone et al, 1999	19507	Rat (<i>Rattus norvegicus</i>)	4	U	DR	13	w	NR	NR	JV	M	GRO	BDWT	WO	11.7	23.4	78		
97	Alexander et al, 1978	36694	Meadow vole (<i>Microtus pennsylvanicus</i>)	3	U	FD	45	d	14	d	JV	NR	GRO	BDWT	WO	12.5		68		
98	Nation et al, 1985	19460	Rat (<i>Rattus norvegicus</i>)	3	U	FD	14	d	80	d	JV	M	GRO	BDWT	WO	20.0		72		
99	Alexander et al, 1978	36694	Meadow vole (<i>Microtus pennsylvanicus</i>)	4	M	FD	49	d	14	d	JV	NR	GRO	BDWT	WO	29.4	309	85		
101	Ambrose et al, 1976	14474	Dog (<i>Canis familiaris</i>)	4	U	FD	65	w	6	mo	JV	B	GRO	BDWT	WO	45.0	112	82		
100	Chernoff and Kavlock, 1982	1260	Mouse (<i>Mus musculus</i>)	2	U	GV	5	d	60	d	GE	F	GRO	BDWT	WO	45.3		82		
102	Berman and Rehnberg, 1983	19064	Mouse (<i>Mus musculus</i>)	3	U	DR	15	d	NR	NR	GE	F	GRO	BDWT	WO	85.3	171	79		
103	Dieter,et al, 1988	61	Mouse (<i>Mus musculus</i>)	4	UX	DR	180	d	6-8	w	JV	F	GRO	BDWT	WO	107	148	84		
104	Spears and Hatfield, 1985	19666	Rat (<i>Rattus norvegicus</i>)	2	U	FD	21	d	NR	NR	JV	M	GRO	BDWT	WO		2.81	77		
105	Oosting et al, 1991	19514	Rat (<i>Rattus norvegicus</i>)	2	M	FD	28	d	31	d	JV	M	GRO	BDWT	WO		8.20	82		
106	Clary, 1975	19111	Rat (<i>Rattus norvegicus</i>)	2	U	DR	4	mo	NR	NR	JV	M	GRO	BDWT	WO		24.7	72		
107	Weber and Reid, 1969	14485	Mouse (<i>Mus musculus</i>)	3	U	FD	4	w	NR	NR	JV	F	GRO	BDWT	WO		208	73		

Table 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)
Nickel
Page 3 of 3

Result #	Reference	Ref No.	Test Organism	# of Conc/ Doses	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Effect Type	Effect Measure	Response Site	NOAEL Dose*	LOAEL Dose*	Total
Survival (MOR)																		
108	Pandey et al, 1999	19521	Mouse (<i>Mus musculus</i>)	3	U	GV	35	d	NR	NR	JV	M	MOR	MORT	WO	2.71		85
109	Nielsen et al 1982	19483	Rat (<i>Rattus norvegicus</i>)	3	U	FD	11	w	NR	NR	JV	F	MOR	MORT	WO	4.56		77
110	Pandey and Srivastava, 2000	36722	Mouse (<i>Mus musculus</i>)	4	U	OR	35	d	NR	NR	JV	M	MOR	MORT	WO	5.42		85
111	Pandey and Srivastava, 2000	36722	Mouse (<i>Mus musculus</i>)	4	U	OR	35	d	NR	NR	JV	M	MOR	MORT	WO	6.47		85
112	Alexander et al, 1978	36694	Meadow vole (<i>Microtus pennsylvanicus</i>)	4	M	FD	14	d	14	d	JV	NR	MOR	MORT	WO	29.4	309	86
113	Chernoff and Kavlock, 1982	1260	Mouse (<i>Mus musculus</i>)	2	U	GV	5	d	60	d	GE	F	MOR	MORT	WO	45.3		76
114	Seidenberg et al 1986	113	Mouse (<i>Mus musculus</i>)	2	U	GV	4	d	NR	NR	GE	F	MOR	MORT	WO	90.6		85
115	Ambrose et al, 1976	14474	Dog (<i>Canis familiaris</i>)	4	U	FD	24	mo	6	mo	JV	B	MOR	MORT	WO	112		77
116	Cempel and Janicka, 2002	36331	Rat (<i>Rattus norvegicus</i>)	3	U	DR	90	d	NR	NR	JV	M	MOR	MORT	WO	138		74
117	Ambrose et al, 1976	14474	Rat (<i>Rattus norvegicus</i>)	4	U	FD	24	mo	28	d	JV	B	MOR	MORT	WO	205		78
118	Schroeder et al, 1964	14447	Mouse (<i>Mus musculus</i>)	2	U	DR	16	mo	21	d	JV	B	MOR	SURV	WO	0.620		73
119	Schroeder and Mitchener, 1975	1858	Mouse (<i>Mus musculus</i>)	2	U	DR	520	d	19-20	d	AD	F	MOR	LFSP	WO	0.716		68

AATT = alanine aminotransferase; ACTP = accuracy of learned behavior; AD = adult; ALBM = albumins; ALPH = alkaline phosphatase; AMMO = ammonia; ASCA = ascorbic acid; B = both; BDWT = body weight changes; BEH = behavior; BL = blood; BLPR = blood pressure; BM = bone marrow; bw = body weight; CALC = calcium; CCOX = cytochrome C-oxidase; CHM = chemical changes; d - day; DEYO = development of young; DIFD = digestibility of food; DR = drinking water; EDMA = edema; ENZ = enzyme level changes; EXCR = excretion; F = female; FCNS = food consumption; FD = food; FDB = feeding behavior; G6PD = glucose-6-phosphate dehydrogenase; GBCM = general biochemical changes; GE = gestation; GENZ = general enzyme changes; GHIS = general histology; GIDX = gestation index; GOTR = glutamic-oxaloacetic transaminase; GPHY = general physiology changes; GRO = growth; GRS = gross body weight changes; GT = gastrointestinal tract; GV = gavage; HE = heart; HIS = histological changes; HMCT = hematocrit; HMGL = hemoglobin; JV = juvenile; kg = kilograms; KI = kidney; L = liter; LADH = lactate dehydrogenase; LEUK = leukocytes; LFSP = lifespan; LI = liver; LOAEL = lowest observed adverse effect level; LU = lungs; M = male; M = measured; MADH = malic dehydrogenase; MK = milk; mo = months; MOR = effects on mortality and survival; M^t NMVM = number of movements; NOAEL = No Observed Adverse Effect Level; NR = Not reported; OR = other oral; ORW = organ weight changes; ORWT = organ weight changes; P450 = cytochrome P450 proteins; PG = prostate gland; PHY = physiology; PL = plasma; PRFM = pregnant females in a population; PROG = progeny numbers/counts; PRTL = protein, total; PRWT = progeny weight; PTH = pathology; REP = reproduction; RSUC = reproductive success (general); SH = stomach; SM = sperm; SM = sexually mature SMIX = weight relative to body weight; SPCL = sperm cell counts; SPCV = sperm cell volume; SR = serum; SURV = survival; TE = testes; TEWT = testes weight; U = unmeasured; UR = urine; UX = measured but values not reported; w = weeks; WCON = water consumption; WO = whole organism; yr = year.

*NOAEL and LOAEL values that are equal and from the same reference represent different experimental designs.

Within the reviewed papers there are 119 results for biochemical (BIO), behavior (BEH), physiology (PHY), pathology (PTH), reproduction (REP), growth (GRO), and survival (MOR) endpoints with a total Data Evaluation Score >65 that were used to derive the TRV (U.S. EPA 2003; Attachment 4-4). These data are plotted in Figure 6.1 and correspond directly with the data presented in Table 6.1. The NOAEL results for growth and reproduction are used to calculate a geometric mean NOAEL. This geometric mean is examined in relationship to the lowest bounded LOAEL for reproduction, growth, and survival to derive the TRV according to the Eco-SSL guidance (U.S. EPA 2003; Attachment 4-5).

A geometric mean of the NOAEL values for reproduction and growth was calculated at 7.70 mg nickel/kg bw/day. However this value is higher than the lowest bounded LOAEL for reproduction, growth, or mortality results. Therefore, the TRV is equal to the highest bounded NOAEL below the lowest bounded LOAEL for reproduction, growth, or survival, and is equal to 1.70 mg nickel/kg bw/day.

6.2 Estimation of Dose and Calculation of the Eco-SSL

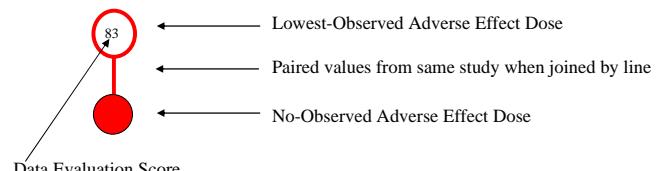
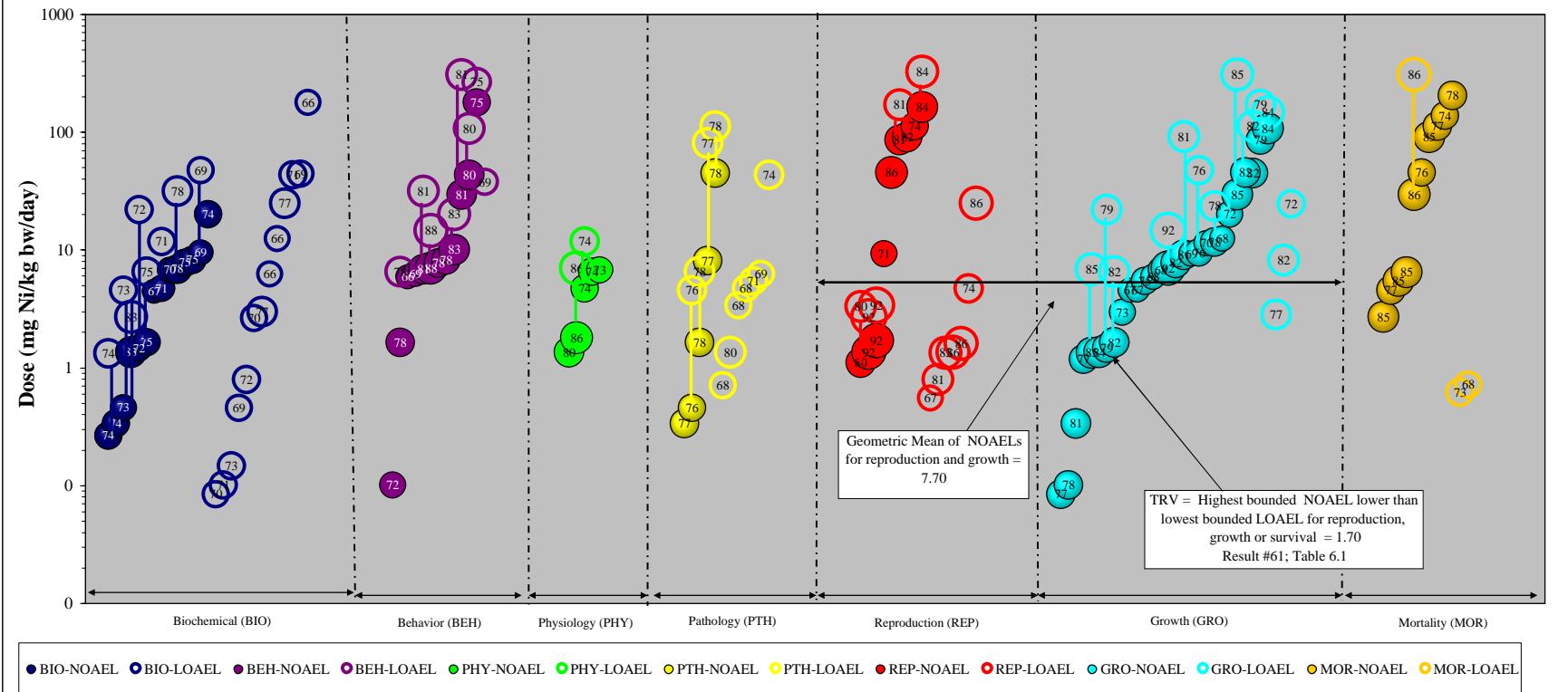
Two separate Eco-SSL values were calculated for mammalian wildlife, one for each of two surrogate receptor groups representing different trophic levels. The mammalian Eco-SSLs derived for nickel were calculated according to the Eco-SSL guidance (U.S. EPA, 2003; Attachment 4-5) and are summarized in Table 6.2. An Eco-SSL value could not be calculated for the mammalian ground insectivore (shrew) as a reliable method to estimate uptake of nickel from soil into soil invertebrates could not be identified (see Attachment 4-1 of the Eco-SSL guidance (U.S. EPA, 2003)).

Table 6.2 Calculation of the Mammalian Eco-SSLs for Nickel

Surrogate Receptor Group	TRV for Nickel (mg dw/kg bw/d) ¹	Food Ingestion Rate (FIR) ² (kg dw/kg bw/d)	Soil Ingestion as Proportion of Diet (P_s) ²	Concentration of Nickel in Biota Type (i) ^{2,3} (B_i) (mg/kg dw)	Eco-SSL (mg/kg dw) ⁴
Mammalian herbivore (vole)	1.70	0.0875	0.032	$\ln(B_i) = 0.748 * \ln(\text{Soil}_i) - 2.223$ where i = plants	340
Mammalian ground insectivore (shrew)	1.70	0.209	0.030	NA	NA
Mammalian carnivore (weasel)	1.70	0.130	0.043	$\ln(B_i) = 0.4658 * \ln(\text{Soil}_i) - 0.2462$ where i = mammals	130

¹ The process for derivation of wildlife TRVs is described in Attachment 4-5 of U.S. EPA (2003).
² Parameters (FIR, P_s , B_i values, regressions) are provided in U.S. EPA (2003) Attachment 4-1 (revised February 2005).
³ B_i = Concentration in biota type (i) which represents 100% of the diet for the respective receptor.
⁴ HQ = [FIR * (Soil_j * P_s + B_i)] / TRV solved for HQ=1 where Soil_j = Eco-SSL (Equation 4-2; U.S. EPA, 2003).
NA = Not Available

Figure 6.1 Mammalian TRV Derivation for Nickel



Wildlife TRV Derivation Process

- 1) There are at least three results available for two test species within the growth, reproduction, and mortality effect groups.
There are enough data to derive a TRV.
- 2) There are three NOAEL results available within the growth and reproduction effect groups for calculation of a geometric mean.
- 3) The geometric mean is equal to 7.70 mg nickel/kg bw/d. However this value is higher than the lowest bounded LOAEL for results within the reproduction, growth, and survival (MOR) effect groups.
- 4) The mammalian wildlife TRV for nickel is equal to 1.70 mg nickel/kg bw/day which is the highest bounded NOAEL lower than the lowest bounded LOAEL within the reproduction, growth or survival effect groups.

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7.5 References Rejected for Use in Derivation of Wildlife TRV

These references were reviewed and rejected for use in derivation of the Eco-SSL. The definition of the codes describing the basis for rejection is provided at the end of the reference sections.

- Unrel** addition of calcium ions for enhancing the safety of metal-ligand chelates as magnetic resonance imaging agents and x-ray contrast agents. *PCT Int. Appl.* 10 pp.
- Diss** arsenic: an analytical procedure to determine its total content in biological samples and signs of its deprivation in rats and chicks. 788794 *ORDER NO: AAD82-20750*
- Diss** aspects of crack growth in structural materials in light water reactors (stainless steel, nickel alloys). 01642159 *ORDER NO: NOT AVAILABLE FROM UNIVERSITY MICROFILMS INT'L.*
- Diss** attempts to isolate with gas liquid chromatography chemical differences between reward and frustrative nonreward odor emissions in the rat. 699745 *ORDER NO: AAD80-25899*
- CP** 1971. *Bulletin of the Chemical Society - Belgrade. Volume 34, Number 8-9-10, 1969. SFCSI-COMM(TT-69-51006/8-9-10)*
- Diss** content and evolution of cadmium, cobalt, chromium, copper, nickel, lead, and zinc in soils of l'horta and ribera baixa regions (valencia) (spain) original title: contenido y evolucion de cadmio, cobalto, cromo, cobre, niquel, plomo, cinc en suelos de las comarcas de l'horta y la baixa (valencia). 01269400 *ORDER NO: NOT AVAILABLE FROM UNIVERSITY MICROFILMS INT'L.*
- Diss** ecophysiology of the common cockle (*cerastoderma edule* l.) in southampton water, with particular reference to pollution (england). 1092481 *ORDER NO: AADDX-87466*

- Diss** effects of ethanol on muscarinic receptor-induced responses in astroglia (carbachol, calcium).
01698206 ORDER NO: AAD99-24077
- CP** 1971-1977. fifty-sixth annual conference of the north central entomologists.| au-yuill, t. m.; scholl, p. j.; meyer, j. a.; knapp, f. w.; christensen,c. m.; peterson, r. d.; williams, r. e.; hair, j. a.; johnson, o. a.;nickel, c. a.; mcneal, c. d., jr.; campbell, j. b.; moneyham, g. e.;shugart, j. i.; boxler, d.; coffey, m. d.; boxler, d. j.; berry, r.l.; parsons, m. a.; lalonde-weigert, b. j.; restifo, r. a.; keiper,d.; lebio, j.; sipos, c.; bear, f. t.; halpin, t. j.; jennings, m. r.;zaim, m.; newson, h. d. *Proceedings, North Central Branch, Entomological Society of America*
- Diss** heavy metal toxicity and nutrient depletion in upper bear creek reservoir. *01229302 ORDER NO: AAD13-46169*
- Diss** immune response to orthopaedic biomaterials. *01440048 ORDER NO: AADAA-I9534554*
- Diss** the murine dna methyltransferase: purification, heterologous expression, steady-state kinetics and post-translational processing. *01572339 ORDER NO: AAD97-27712*
- No COC** news about chemicals. *IRPTC Bull. V8, N1, P17(16)*
- No Oral** nickel permeation pathways in the small intestine and the olfactory system (iron deficiency).
01626234 ORDER NO: NOT AVAILABLE FROM UNIVERSITY MICROFILMS INT'L.
- Diss** nickelous nitrate hexahydrate and its effect on reproduction in the mouse. *702149 ORDER NO: AAD13-01167*
- Mix** 1993. ntp technical report on toxicity studies of a chemical mixture of 25 groundwater contaminants administered in ***drinking*** ***water*** to f344/n ***rats*** and b6c3fl ***mice***. *Natl. Toxicol. Program Toxic. Rep. Ser.* 35: 184 pp.
- FL** 1978. (on the use of radioisotopic methods in an hygienic experiment).| au- moscow, ussr.
GIGIENA I SANITARIYA No. 9: 72-75.
- Unrel** 1999. *Report on the Proposed Voisey's Bay Mine and Mill Project. SSC-EN106-45/1999E; ISBN-0-662-27511-X*
- Diss** the role of the nmda receptor and cyclic guanosine 3',5'-monophosphate (cgmp) in the pathophysiological mechanisms of experimental brain injury: a time-course study (trauma, intracellular signaling, memory). *01603489 ORDER NO: AAD98-04188*
- No Oral** soft actuators and artificial muscles. *PCT Int. Appl.* 53 pp..
- Diss** studies on the muscarinic receptors in neuroblastoma cells. *750140 ORDER NO: AAD81-14988*
- Diss** a study of the chemistry and mutagenicity of welding fume. *910380 ORDER NO: NOT AVAILABLE FROM UNIVERSITY MICROFILMS INT'L.*
- Diss** the susceptibility of pulmonary immunologic phenomena to the toxic effects of nickel subsulfide. *933904 ORDER NO: AAD86-25095*
- Diss** the synthesis and characterization of some polymer-supported macrocyclic complexes and their interaction with oxygen. *01134166 ORDER NO: AADDX-90556*
- Diss** theoretical studies of the electronic structure and bonding in some transition metal complexes of

- vanadium, chromium, iron, nickel and palladium. 765403 ORDER NO: AAD81-24610
- Drug** treatment of septic shock with transition metal complexes. *PCT Int. Appl.* 13 pp.
- Diss** ultraviolet radiation and allergic contact dermatitis: an experimental and clinical study. 0997098 ORDER NO: NOT AVAILABLE FROM UNIVERSITY MICROFILMS INT'L.
- No Oral** 1979. whole-body autoradiography of 63ni in mice throughout gestation.| au- blindern, oslo 3, Norway. *Toxicology* 12(2): 165-172.
- Diss** zinc and calcium effects on nickel dermatitis in the guinea pig. 0961875 ORDER NO: AAD87-17748
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- Bio Acc** Adam, J. and Pinta, M. 1986. tibial dyschondroplasia of broilers. syndrome of s-shaped tibias. *Bulletin Mensuel De La Societe Veterinaire Pratique De France* 70(7): 425-428.
- Unrel** Adam, J., Pinta, M., and Goyon, M. 1986. hyena disease of cattle. *Bulletin Mensuel De La Societe Veterinaire Pratique De France* 70(7): 405...416.
- No Dose** Adam, J., Pinta, M., and Viel, M. 1981. hyaena disease in cattle and nickel deficiency. *Bulletin De L'Academie Veterinaire De France* 54(3): 329-335.
- No Oral** Adamis, Z., Tatrai, E., Honma, K., Karpati, J., and Ungvary, G. 1997. a study on lung toxicity of respirable hard metal dusts in ***rats***. *Ann. Occup. Hyg.* (1997) 41(5): 515-526 .
- CP** <Additional> Rapp, W. F., Amin, O. M., Amrine, J. A., Lewis, R. E., McGivern, J. J., Rai, K. S., Treece, R. E., Wagoner, D. E., Johnson, O. A., Childress, D., Nickel, C. A., Birkenmeyer, D. R., Dame, D. A., Pinger, R. R. Jr., Degruillier, M. E., Leopold, R. A., and Heller, P. R. 1974. medical and veterinary insects. *Proceedings, North Central Branch, Entomological Society of America* 29: 184-188.
- Abstract** Adkins, B. Jr and Gardner, D. E. the effect of nickel on the enhancement of induced streptococcal infections. *Abstracts of the Annual Meeting of the American Society for Microbiology*. 76. 1976 B45
- Acu** Adler, M. W. and Adler, C. H. 1977. toxicity to heavy metals and relationship to seizure thresholds. *Clin Pharmacol Ther.* 22(5 Pt 2): 774-9.
- FL** Ado, V. A. and Horyachkina, L. A. modeling of chemical allergoses and their selective and nonspecific inhibition. *Fiziol Zh (Kiev)*; 21 (4). 1975 481-485
- Phys** Ahmed, Abu, Kobayashi, Sei, Shikasho, Tomomi, Nishimura, Junji, and Kanaide, Hideo. differential effects of ca²⁺ channel blockers on ca²⁺ transients and cell cycle progression in vascular smooth muscle cells. *Eur. J. Pharmacol.* (1998) 344(2/3): 323-331
- Diss** Ahpasri Chaigool. 1985. genotoxicity of some heavy metals on chick embryos. <original> phit khong loha nak bangchanit to san phanthukam nai embryo kai. 83 Leaves
- Aquatic** Albers, Peter H. and Camardese, Michael B. effects of acidification on metal accumulation by aquatic plants and invertebrates. 2. wetlands, ponds, and small lakes. *Environ. Toxicol. Chem.*

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In Vit	Alcain, F. J., Low, H., and Crane, F. L. iron at the cell surface controls both dna synthesis and plasma membrane redox system. <i>Protoplasma</i> (1995) 184(1-4): 233-7
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CP	Anke, M., Kronemann, H., Groppel, B., Hennig, A., Meissner, D., and Schneider, H. J. 1980. the influence of nickel-deficiency on ***growth*** , ***reproduction*** , longevity and different biochemical parameters of goats. <i>Spurenelem.-Symp.: Nickel 3rd : 3-10. Editor(s): Anke, Manfred; Schneider, Hans-Joachim; Brueckner, Chr. Publisher: Friedrich-Schiller- Univ. Jena Abt. Wiss. Publ., Jena, Ger. Dem. Rep.</i>
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- No Oral** Zissu, D., Cavelier, C., and De Ceaurriz, J. 1987. experimental sensitization of guinea-pigs to nickel and patch testing with metal samples. *Food and Chemical Toxicology* 25(1): 83-5.
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Literature Rejection Categories		
Rejection Criteria	Description	Receptor
ABSTRACT (Abstract)	Abstracts of journal publications or conference presentations.	Wildlife Plants and Soil Invertebrates
ACUTE STUDIES (Acu)	Single oral dose or exposure duration of three days or less.	Wildlife
AIR POLLUTION (Air P)	Studies describing the results for air pollution studies.	Wildlife Plants and Soil Invertebrates
ALTERED RECEPTOR (Alt)	Studies that describe the effects of the contaminant on surgically-altered or chemically-modified receptors (e.g., right nephrectomy, left renal artery ligation, hormone implant, etc.).	Wildlife
AQUATIC STUDIES (Aquatic)	Studies that investigate toxicity in aquatic organisms.	Wildlife Plants and Soil Invertebrates
ANATOMICAL STUDIES (Anat)	Studies of anatomy. Instance where the contaminant is used in physical studies (e.g., silver nitrate staining for histology).	Wildlife
BACTERIA (Bact)	Studies on bacteria or susceptibility to bacterial infection.	Wildlife Plants and Soil Invertebrates
BIOACCUMULATION SURVEY (Bio Acc)	Studies reporting the measurement of the concentration of the contaminant in tissues.	Wildlife Plants and Soil Invertebrates
BIOLOGICAL PRODUCT (BioP)	Studies of biological toxicants, including venoms, fungal toxins, <i>Bacillus thuringiensis</i> , other plant, animal, or microbial extracts or toxins.	Wildlife Plants and Soil Invertebrates
BIOMARKER (Biom)	Studies reporting results for a biomarker having no reported association with an adverse effect and an exposure dose (or concentration).	Wildlife
CARCINOGENICITY STUDIES (Carcin)	Studies that report data only for carcinogenic endpoints such as tumor induction. Papers that report systemic toxicity data are retained for coding of appropriate endpoints.	Wildlife Plants and Soil Invertebrates
CHEMICAL METHODS (Chem Meth)	Studies reporting methods for determination of contaminants, purification of chemicals, etc. Studies describing the preparation and analysis of the contaminant in the tissues of the receptor.	Wildlife Plants and Soil Invertebrates
CONFERENCE PROCEEDINGS (CP)	Studies reported in conference and symposium proceedings.	Wildlife Plants and Soil Invertebrates
DEAD (Dead)	Studies reporting results for dead organisms. Studies reporting field mortalities with necropsy data where it is not possible to establish the dose to the organism.	Wildlife Plants and Soil Invertebrates
DISSERTATIONS (Diss)	Dissertations are excluded. However, dissertations are flagged for possible future use.	Wildlife
DRUG (Drug)	Studies reporting results for testing of drug and therapeutic effects and side-effects. Therapeutic drugs include vitamins and minerals. Studies of some minerals may be included if there is potential for adverse effects.	Wildlife Plants and Soil Invertebrates
DUPLICATE DATA (Dup)	Studies reporting results that are duplicated in a separate publication. The publication with the earlier year is used.	Wildlife Plants and Soil Invertebrates

Literature Rejection Categories		
Rejection Criteria	Description	Receptor
ECOLOGICAL INTERACTIONS (Ecol)	Studies of ecological processes that do not investigate effects of contaminant exposure (e.g., studies of “silver” fox natural history; studies on ferrets identified in iron search).	Wildlife Plants and Soil Invertebrates
EFFLUENT (Effl)	Studies reporting effects of effluent, sewage, or polluted runoff.	Wildlife Plants and Soil Invertebrates
ECOLOGICALLY RELEVANT ENDPOINT (ERE)	Studies reporting a result for endpoints considered as ecologically relevant but is not used for deriving Eco-SSLs (e.g., behavior, mortality).	Plants and Soil Invertebrates
CONTAMINANT FATE/METABOLISM (Fate)	Studies reporting what happens to the contaminant, rather than what happens to the organism. Studies describing the intermediary metabolism of the contaminant (e.g., radioactive tracer studies) without description of adverse effects.	Wildlife Plants and Soil Invertebrates
FOREIGN LANGUAGE (FL)	Studies in languages other than English.	Wildlife Plants and Soil Invertebrates
FOOD STUDIES (Food)	Food science studies conducted to improve production of food for human consumption.	Wildlife
FUNGUS (Fungus)	Studies on fungus.	Wildlife Plants and Soil Invertebrates
GENE (Gene)	Studies of genotoxicity (chromosomal aberrations and mutagenicity).	Wildlife Plants and Soil Invertebrates
HUMAN HEALTH (HHE)	Studies with human subjects.	Wildlife Plants and Soil Invertebrates
IMMUNOLOGY (IMM)	Studies on the effects of contaminants on immunological endpoints.	Wildlife Plants and Soil Invertebrates
INVERTEBRATE (Invert)	Studies that investigate the effects of contaminants on terrestrial invertebrates are excluded.	Wildlife
IN VITRO (In Vit)	<i>In vitro</i> studies, including exposure of cell cultures, excised tissues and/or excised organs.	Wildlife Plants and Soil Invertebrates
LEAD SHOT (Lead shot)	Studies administering lead shot as the exposure form. These studies are labeled separately for possible later retrieval and review.	Wildlife
MEDIA (Media)	Authors must report that the study was conducted using natural or artificial soil. Studies conducted in pore water or any other aqueous phase (e.g., hydroponic solution), filter paper, petri dishes, manure, organic or histosols (e.g., peat muck, humus), are not considered suitable for use in defining soil screening levels.	Plants and Soil Invertebrates
METHODS (Meth)	Studies reporting methods or methods development without usable toxicity test results for specific endpoints.	Wildlife Plants and Soil Invertebrates
MINERAL REQUIREMENTS (Mineral)	Studies examining the minerals required for better production of animals for human consumption, unless there is potential for adverse effects.	Wildlife
MIXTURE (Mix)	Studies that report data for combinations of single toxicants (e.g. cadmium and copper) are excluded. Exposure in a field setting from contaminated natural soils or waste application to soil may be coded as Field Survey.	Wildlife Plants and Soil Invertebrates

Literature Rejection Categories		
Rejection Criteria	Description	Receptor
MODELING (Model)	Studies reporting the use of existing data for modeling, i.e., no new organism toxicity data are reported. Studies which extrapolate effects based on known relationships between parameters and adverse effects.	Wildlife Plants and Soil Invertebrates
NO CONTAMINANT OF CONCERN (No COC)	Studies that do not examine the toxicity of Eco-SSL contaminants of concern.	Wildlife Plants and Soil Invertebrates
NO CONTROL (No Control)	Studies which lack a control or which have a control that is classified as invalid for derivation of TRVs.	Wildlife Plants and Soil Invertebrates
NO DATA (No Data)	Studies for which results are stated in text but no data is provided. Also refers to studies with insufficient data where results are reported for only one organism per exposure concentration or dose (wildlife).	Wildlife Plants and Soil Invertebrates
NO DOSE or CONC (No Dose)	Studies with no usable dose or concentration reported, or an insufficient number of doses/concentrations are used based on Eco-SSL SOPs. These are usually identified after examination of full paper. This includes studies which examine effects after exposure to contaminant ceases. This also includes studies where offspring are exposed in utero and/or lactation by doses to parents and then after weaning to similar concentrations as their parents. Dose cannot be determined.	Wildlife Plants and Soil Invertebrates
NO DURATION (No Dur)	Studies with no exposure duration. These are usually identified after examination of full paper.	Wildlife Plants and Soil Invertebrates
NO EFFECT (No Efect)	Studies with no relevant effect evaluated in a biological test species or data not reported for effect discussed.	Wildlife Plants and Soil Invertebrates
NO ORAL (No Oral)	Studies using non-oral routes of contaminant administration including intraperitoneal injection, other injection, inhalation, and dermal exposures.	Wildlife
NO ORGANISM (No Org) or NO SPECIES	Studies that do not examine or test a viable organism (also see in vitro rejection category).	Wildlife Plants and Soil Invertebrates
NOT AVAILABLE (Not Avail)	Papers that could not be located. Citation from electronic searches may be incorrect or the source is not readily available.	Wildlife Plants and Soil Invertebrates
NOT PRIMARY (Not Prim)	Papers that are not the original compilation and/or publication of the experimental data.	Wildlife Plants and Soil Invertebrates
NO TOXICANT (No Tox)	No toxicant used. Publications often report responses to changes in water or soil chemistry variables, e.g., pH or temperature. Such publications are not included.	Wildlife Plants and Soil Invertebrates
NO TOX DATA (No Tox Data)	Studies where toxicant used but no results reported that had a negative impact (plants and soil invertebrates).	Plants and Soil Invertebrates
NUTRIENT (Nutrient)	Nutrition studies reporting no concentration related negative impact.	Plants and Soil Invertebrates
NUTRIENT DEFICIENCY (Nut def)	Studies of the effects of nutrient deficiencies. Nutritional deficient diet is identified by the author. If reviewer is uncertain then the administrator should be consulted. Effects associated with added nutrients are coded.	Wildlife
NUTRITION (Nut)	Studies examining the best or minimum level of a chemical in the diet for improvement of health or maintenance of animals in captivity.	Wildlife
OTHER AMBIENT CONDITIONS (OAC)	Studies which examine other ambient conditions: pH, salinity, DO, UV, radiation, etc.	Wildlife Plants and Soil Invertebrates

Literature Rejection Categories		
Rejection Criteria	Description	Receptor
OIL (Oil)	Studies which examine the effects of oil and petroleum products.	Wildlife Plants and Soil Invertebrates
OM, pH (OM, pH)	<p>Organic matter content of the test soil must be reported by the authors, but may be presented in one of the following ways; total organic carbon (TOC), particulate organic carbon (POC), organic carbon (OC), coarse particulate organic matter (CPOM), particulate organic matter (POM), ash free dry weight of soil, ash free dry mass of soil, percent organic matter, percent peat, loss on ignition (LOI), organic matter content (OMC).</p> <p>With the exception of studies on non-ionizing substances, the study must report the pH of the soil, and the soil pH should be within the range of 4 and 8.5. Studies that do not report pH or report pH outside this range are rejected.</p>	Plants and Soil Invertebrates
ORGANIC METAL (Org Met)	Studies which examine the effects of organic metals. This includes tetraethyl lead, triethyl lead, chromium picolinate, phenylarsonic acid, roxarsone, 3-nitro-4-phenylarsonic acid., zinc phosphide, monomethylarsonic acid (MMA), dimethylarsinic acid (DMA), trimethylarsine oxide (TMAO), or arsenobetaine (AsBe) and other organo metallic fungicides. Metal acetates and methionines are not rejected and are evaluated.	Wildlife
LEAD BEHAVIOR OR HIGH DOSE MODELS (Pb Behav)	<p>There are a high number of studies in the literature that expose rats or mice to high concentrations of lead in drinking water (0.1, 1 to 2% solutions) and then observe behavior in offspring, and/or pathology changes in the brain of the exposed dam and/or the progeny. Only a representative subset of these studies were coded.</p> <p>Behavior studies examining complex behavior (learned tasks) were also not coded.</p>	Wildlife
PHYSIOLOGY STUDIES (Phys)	Physiology studies where adverse effects are not associated with exposure to contaminants of concern.	Wildlife
PLANT (Plant)	Studies of terrestrial plants are excluded.	Wildlife
PRIMATE (Prim)	Primate studies are excluded.	Wildlife
PUBL AS (Publ as)	The author states that the information in this report has been published in another source. Data are recorded from only one source. The secondary citation is noted as Publ As.	Wildlife Plants and Soil Invertebrates
QSAR (QSAR)	Derivation of Quantitative Structure-Activity Relationships (QSAR) is a form of modeling. QSAR publications are rejected if raw toxicity data are not reported or if the toxicity data are published elsewhere as original data.	Wildlife Plants and Soil Invertebrates
REGULATIONS (Reg)	Regulations and related publications that are not a primary source of data.	Wildlife Plants and Soil Invertebrates
REVIEW (Rev)	Studies in which the data reported in the article are not primary data from research conducted by the author. The publication is a compilation of data published elsewhere. These publications are reviewed manually to identify other relevant literature.	Wildlife Plants and Soil Invertebrates

Literature Rejection Categories		
Rejection Criteria	Description	Receptor
SEDIMENT CONC (Sed)	Studies in which the only exposure concentration/dose reported is for the level of a toxicant in sediment.	Wildlife Plants and Soil Invertebrates
SCORE (Score)	Papers in which all studies had data evaluation scores at or lower than the acceptable cut-off (#10 of 18) for plants and soil invertebrates).	Plants and Soil Invertebrates
SEDIMENT CONC (Sed)	Studies in which the only exposure concentration/dose reported is for the level of a toxicant in sediment.	Wildlife Plants and Soil Invertebrates
SLUDGE	Studies on the effects of ingestion of soils amended with sewage sludge.	Wildlife Plants and Soil Invertebrates
SOIL CONC (Soil)	Studies in which the only exposure concentration/dose reported is for the level of a toxicant in soil.	Wildlife
SPECIES	Studies in which the species of concern was not a terrestrial invertebrate or plant or mammal or bird.	Plants and Soil Invertebrates Wildlife
STRESSOR (QAC)	Studies examining the interaction of a stressor (e.g., radiation, heat, etc.) and the contaminant, where the effect of the contaminant alone cannot be isolated.	Wildlife Plants and Soil Invertebrates
SURVEY (Surv)	Studies reporting the toxicity of a contaminant in the field over a period of time. Often neither a duration nor an exposure concentration is reported.	Wildlife Plants and Soil Invertebrates
REPTILE OR AMPHIBIAN (Herp)	Studies on reptiles and amphibians. These papers flagged for possible later review.	Wildlife Plants and Soil Invertebrates
UNRELATED (Unrel)	Studies that are unrelated to contaminant exposure and response and/or the receptor groups of interest.	Wildlife
WATER QUALITY STUDY (Wqual)	Studies of water quality.	Wildlife Plants and Soil Invertebrates
YEAST (Yeast)	Studies of yeast.	Wildlife Plants and Soil Invertebrates

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Appendix 5-1

Avian Toxicity Data Extracted and Reviewed for Wildlife Toxicity Reference Value (TRV) - Nickel

March 2007

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Appendix 5.1 Avian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)
Nickel
Page 1 of 1

Ref		Reference	Chemical Form	MW%	Test Species	Phase #	# of Conc/ Doses	Exposure												Conc/Dose Units	Wet Weight Reported?	Percent Moisture	Application Frequency	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Control Type	Test Location	General Effect Group	Effects			Conversion to mg/kg bw/day			Result			Data Evaluation Score										
Result #	Ref N.							DLY	U	FD	28	d	21	d	JV	F	C	Lab	BIO	CHM	LIPD	WO	12.0		Y	1.98	N	0.09079	0.136		10	10	5	10	6	1	4	10	10	4	Total													
Biochemical																																																						
1	5407	Oscar et al, 1995	Nickel chloride hexahydrate	24.69	Chicken (<i>Gallus domesticus</i>)	1	3	0/6/12										mg/kg diet	N	na	DLY	U	FD	28	d	21	d	JV	F	C	Lab	BIO	CHM	LIPD	WO	12.0		Y	1.98	N	0.09079	0.136		10	10	5	10	6	1	4	10	10	4	70
2	6885	Nielsen et al, 1975	Nickel chloride hexahydrate	100	Chicken (<i>Gallus domesticus</i>)	1	4	0/50/250/2500									ng/g diet	N	na	NR	U	FD	3	w	1	d	JV	M	C	Lab	BIO	CHM	HMCT	BL	2500		Y	0.433	N	0.03375	0.195		10	10	5	10	6	1	4	10	10	4	70	
3	6666	Ling and Leach, 1979	Nickel chloride	100	Chicken (<i>Gallus domesticus</i>)	1	6	0/300/500/700/900/1100									mg/kg diet	N	na	ADL	U	FD	3	w	1	d	JV	M	C	Lab	BIO	CHM	HMGL	BL	900	1100	Y	0.1179	N	0.01447	110	135	10	10	5	10	6	1	10	10	4	76		
4	5345	Martinez and Diaz, 1996	Nickel chloride hexahydrate	100	Chicken (<i>Gallus domesticus</i>)	1	4	0/123.5/247/494									mg/kg diet	N	na	ADL	U	FD	42	d	1	d	JV	M	C	Lab	BIO	CHM	HMGL	BL	124	Y	1.8901	N	0.08809	5.76	10	10	5	10	6	1	4	10	10	4	70			
Behavior																																																						
5	5407	Oscar et al, 1995	Nickel chloride hexahydrate	24.69	Chicken (<i>Gallus domesticus</i>)	1	3	0/6/12									mg/kg diet	N	na	DLY	U	FD	28	d	21	d	JV	F	C	Lab	BEH	FDB	FCNS	WO	12.0		Y	1.98	N	0.09079	0.136		10	10	5	10	6	4	4	10	10	4	73	
6	5345	Martinez and Diaz, 1996	Nickel chloride hexahydrate	100	Chicken (<i>Gallus domesticus</i>)	1	4	0/123.5/247/494								mg/kg diet	N	na	ADL	U	FD	42	d	1	d	JV	M	C	Lab	BEH	FDB	FCNS	WO	123.5	247	Y	1.8901	N	0.08809	5.76	11.5	10	10	5	10	6	4	10	10	4	79			
Physiology																																																						
7	60	Weber and Reid, 1968	Nickel acetate	100	Chicken (<i>Gallus domesticus</i>)	2	8	0/3.04/9.25/13.68/14.61/15.86/16.21/17.25								mg/org/d	N	na	ADL	U	FD	4	w	1	d	JV	B	C	Lab	PHY	PHY	FDCV	WO	13.7	14.6	Y	0.484	Y	0.001	28.3	30.2	10	10	5	5	7	4	10	10	4	75			
8	60	Weber and Reid, 1968	Nickel sulfate	100	Chicken (<i>Gallus domesticus</i>)	1	8	0/3.11/9.61/13.32/14.14/14.50/14.71/17.07							mg/org/d	N	na	ADL	U	FD	4	w	1	d	JV	B	C	Lab	PHY	PHY	META	WO	14.5	14.71	Y	0.467	Y	0.001	31.0	31.5	10	10	5	10	7	4	10	10	4	80				
Pathology																																																						
9	6885	Nielsen et al, 1975	Nickel chloride hexahydrate	100	Chicken (<i>Gallus domesticus</i>)	1	4	0/50/250/2500								ng/g diet	N	na	NR	U	FD	3	w	1	d	JV	M	C	Lab	PTH	ORW	SMIX	SK	2500		Y	0.433	N	0.03375	0.195		10	10	5	10	6	4	4	10	10	4	73		
10	6461	Cain and Pafford, 1981	Nickel sulfate	100	Duck (<i>Anas platyrhynchos</i>)	1	4	0/176/774/1069								mg/kg diet	N	na	ADL	M	FD	28	d	1	d	JV	B	C	Lab	PTH	ITX	ATAX	WO	176	774	Y	1.023	N	0.05907	10.7	47.0	10	10	10	10	6	4	8	10	10	4	82		
11	5345	Martinez and Diaz, 1996	Nickel chloride hexahydrate	100	Chicken (<i>Gallus domesticus</i>)	1	4	0/123.5/247/494								mg/kg diet	N	na	ADL	U	FD	42	d	1	d	JV	M	C	Lab	PTH	HIS	GHIS	HE	247	494	Y	1.7285	N	0.08311	11.9	23.8	10	10	5	10	6	4	10	10	4	79			
Reproduction																																																						
12	6492	Eastin and O'Shea, 1981	Nickel sulfate	100	Duck (<i>Anas platyrhynchos</i>)	1	5	0/12.5/50/200/800								mg/kg diet	N	na	ADL	U	FD	90	d	20	mo	LB	F																											



Appendix 6-1

*Mammalian Toxicity Data Extracted and Reviewed for Wildlife
Toxicity Reference Value (TRV) - Nickel*

March 2007

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Appendix 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)
Nickel
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Ref		Chemical Form	Reference	Exposure												Effects				Conversion to mg/kg bw/day				Result		Data Evaluation Score																		
Result #	Ref N.			MW%	Test Species	# of Conc/ Doses	Conc/ Doses	Conc/Dose Units	Wet Weight Reported?	Percent Moisture	Application Frequency	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Control Type	Test Location	General Effect Group	Effect Type	Effect Measure	Response Site	Study NOAEL	Study LOAEL	Body Weight Reported?	Body Weight in kg	Ingestion Rate Reported?	Ingestion Rate in kg or L/day	NOAEL Dose (mg/kg/day)	Data Source	Dose Route	Test Concentrations	Chemical form	Dose Quantification	Endpoint	Dose Range	Statistical Power	Exposure Duration	Test Conditions	Total	
Biochemical																																												
1	19671	Spears et al, 1984	Nickel chloride hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/25	mg/kg diet	N	na	ADL	U	FD	21	d	1	d	JV	NR	C	Lab	BIO	ENZ	ALPH	SR	5	25	Y	4.329	N	0.22912	0.265	1.32	10	10	5	10	6	1	8	10	10	4	74
2	19098	Chatterjee et al, 1979	Nickel sulfate hexahydrate	22.33	Rat (<i>Rattus norvegicus</i>)	2	0/1.5	mg/kg bw/d	N	na	NR	U	FD	21	d	NR	NR	JV	M	C	Lab	BIO	CHM	ASCA	LI	1.50		Y	0.0755	N	0.008215	0.335		10	10	5	10	10	1	4	10	10	4	74
3	19478	Nielsen, 1980	Nickel chloride trihydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	DLY	U	FD	10	w	NR	NR	JV	F	C	Lab	BIO	CHM	HMCT	PL	5.00	50.0	N	0.204	N	0.0186	0.456	4.56	10	10	5	10	5	1	8	10	10	4	73
4	19521	Pandey et al, 1999	Nickel sulfate	37.94	Mouse (<i>Mus musculus</i>)	3	0/3.57/7.14	mg/kg bw/d	N	na	5 per w	U	GV	35	d	NR	NR	JV	M	C	Lab	BIO	ENZ	LADH	TE	3.57	7.14	Y	0.025	N	0.00331	1.35	2.71	10	8	10	10	1	10	10	10	4	83	
5	19666	Spears and Hatfield, 1985	Nickel chloride hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/15/225	mg/kg diet	N	na	ADL	U	FD	21	d	NR	NR	JV	M	C	Lab	BIO	CHM	LEUK	BL	15	225	Y	0.137	N	0.01341	1.47	22.0	10	10	5	10	6	1	6	10	10	4	72
6	14477	O'Dell et al, 1970	Nickelous carbonate	100	Cattle (<i>Bos taurus</i>)	4	0/62.5/250/1000	mg/kg diet	N	na	DLY	U	FD	8	w	14	w	JV	M	C	Lab	BIO	CHM	CALC	NR	250	1000	Y	180.8	Y	1.180	1.63	6.53	10	10	5	10	7	1	8	10	10	4	75
7	19290	Kadiiska et al, 1985	Nickel sulfate hexahydrate	22.33	Rat (<i>Rattus norvegicus</i>)	2	0/20	mg/kg bw/d	N	na	DLY	U	DR	30	d	NR	NR	JV	M	C	Lab	BIO	ENZ	P450	LI	20.0		Y	0.175	N	0.02062	4.47		10	5	5	10	10	1	4	8	10	4	67
8	19507	Obone et al, 1999	Nickel sulfate	100	Rat (<i>Rattus norvegicus</i>)	4	0/44.7/111.75/223.5	mg/L	N	na	ADL	U	DR	13	w	NR	NR	JV	M	C	Lab	BIO	CHM	PRTL	PL	44.7	112	Y	0.55	N	0.0578	4.70	11.7	10	5	5	10	6	1	10	10	4	71	
9	14476	ODell et al, 1970	Nickelous carbonate	100	Cattle (<i>Bos taurus</i>)	3	0/365/1835	mg/org/d	N	na	1 per d	U	FD	6	w	NR	NR	LC	F	C	NR	BIO	CHM	PRTL	MK	1835		N	272	Y	7.350	6.75		10	10	5	10	6	1	4	10	10	4	70
10	62	Smith et al, 1993	Nickel chloride	100	Rat (<i>Rattus norvegicus</i>)	4	0/1.33/6.8/31.63	mg/kg bw/d	N	na	ADL	M	DR	26	w	62	d	LC	F	C	Lab	BIO	CHM	GBCM	PL	6.80	31.6	Y	0.3325	Y	0.06389	6.80	31.6	10	5	10	10	1	8	10	10	4	78	
11	19514	Oosting et al, 1991	Nickel chloride	100	Rat (<i>Rattus norvegicus</i>)	3	0/49.5/104.9	mg/kg diet	N	na	ADL	M	FD	28	d	31	d	JV	M	C	Lab	BIO	CHM	HMGL	BL	105		N	0.217	Y	0.01610	7.78		10	10	10	10	6	1	4	10	10	4	75
12	19514	Oosting et al, 1991	Nickel chloride	100	Rat (<i>Rattus norvegicus</i>)	2	0/100	mg/kg diet	N	na	ADL	M	FD	28	d	31	d	JV	M	C	Lab	BIO	CHM	HMGL	BL	100		N	0.217	Y	0.0178	8.20		10	10	5	10	6	1	4	10	10	4	75
13	63	Whanger, 1973	Nickel acetate	100	Rat (<i>Rattus norvegicus</i>)	4	0/100/500/1000	mg/kg diet	N	na	DLY	U	FD	6	w	35	d	JV	NR	C	Lab	BIO	CHM	HMGL	BL	100	500	Y	0.163	N	0.01547	9.49	47.4	10	5	5	10	6	1	8	10	10	4	69
14	19460	Nation et al, 1985	Nickel Chloride	100	Rat (<i>Rattus norvegicus</i>)	3	0/10/20	mg/kg bw/d	N	na	DLY	U	FD	14	d	80	d	JV	M	C	Lab	BIO	CHM	GBCM	LI	20.0		Y	0.1867	N	0.0173	20.0		10	10	5	10	10	1	4	10	10	4	74
15	19764	Uthus and Poellot, 1997	Nickel chloride hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	2	0/1	ug/g diet	N	na	DLY	U	FD	57	d	NR	NR	JV	M	C	Lab	BIO	ENZ	GENZ	LI	1	Y	0.315	N	0.0265808		0.0844	10	10	5	10	6	1	4	10	10	4	70	
16	19664	Spears et al, 1986	Nickel chloride	100	Cattle (<i>Bos taurus</i>)	2	0/5	mg/kg diet	N	na	2 per d	U	FD	28	d	50	d	JV	M	C	FieldA	BIO	CHM	AMMO	GT	5	Y	247.6	Y	5.00		0.101	10	10	5	10	7	1	4	10	10	4	71	
17	19437	Mine et al, 1990	Nickel Chloride hexahydrate	100	Sheep (<i>Ovis aries</i>)	2	0/5	mg/org/d	N	na	DLY	U	GV	4	w	5	mo	JV	F	C	Lab	BIO	ENZ	GENZ	SH	5.00	Y	34	N	1.247		0.147	10	8	10	10	6	1	4	10	10	4	73	
18	19461	Nielsen et al, 1979	Nickel chloride trihydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	DLY	U	FD	9	w	21	d	JV	F	C	Lab	BIO	CHM	HMCT	BL	5.00	N	0.204	N	0.0186		0.456	10	5	10	5	1	4	10	10	4	69		
19	19522	Pandey and Singh, 1999	Nickel sulfate hexahydrate	22.33	Mouse (<i>Mus musculus</i>)</td																																							

Appendix 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)
Nickel
Page 2 of 3

Ref		Reference	Chemical Form	MW%	Test Species	Exposure												Effects				Conversion to mg/kg bw/day				Result		Data Evaluation Score															
Result #	Ref N.					# of Conc/ Doses	Conc/ Doses	Conc/Dose Units	Wet Weight Reported?	Percent Moisture	Application Frequency	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Control Type	General Effect Group	Effect Type	Effect Measure	Response Site	Study NOAEL	Body Weight Reported?	Body Weight in kg	Ingestion Rate Reported?	Ingestion Rate in kg or L/day	NOAEL Dose (mg/kg/day)	Dose Source	Dose Route	Test Concentrations	Chemical form	Dose Quantification	Endpoint	Dose Range	Statistical Power	Exposure Duration	Test Conditions	Total		
55	14484	Schroeder et al, 1974	Nickel	100	Rat (<i>Rattus norvegicus</i>)	2	0/5.44	mg/kg bw/d	N	na	DLY	M	DR	1217	d	30	d	JV	B	C Lab	PTH	HIS	GHIS	HE		5.44	Y	0.397	Y	0.027290		5.44	10	5	10	4	4	10	10	4	71		
56	19787	Vyskocil et al, 1994	Nickel sulfate	100	Rat (<i>Rattus norvegicus</i>)	2	0/6.25	mg/kg bw/d	N	na	ADL	U	DR	6	mo	10	w	JV	M	C Lab	PTH	ORW	SMIX	KI		6.25	Y	0.5	Y	0.03120		6.25	10	5	10	7	4	4	10	10	4	69	
57	61	Dieter et al, 1988	Nickel sulfate	37.4	Mouse (<i>Mus musculus</i>)	4	0/115.7/285.7/395.7	mg/kg bw/d	N	na	ADL	UX	DR	180	d	6-8	w	JV	F	C Lab	PTH	ORW	ORWT	LI		116	Y	0.03	Y	0.00347		43.3	10	5	10	10	7	4	4	10	10	4	74
58	113	Seidenberg et al, 1986	Nickelous chloride	45.29	Mouse (<i>Mus musculus</i>)	2	0/200	mg/kg/bw/d	N	na	DLY	U	GV	4	d	NR	NR	GE	F	C Lab	PTH	GRS	BDWT	WO		200	Y	0.0392	N	0.00479		90.6	10	8	10	10	4	4	10	10	4	80	
Reproduction																																											
59	19293	Kakela et al, 1999	Nickel chloride hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	4	0/10/30/100	mg/L	N	na	ADL	U	DR	28	d	3.5-9	mo	SM	M	C Lab	REP	REP	GIDX	WO	10.0	30.0	Y	0.3379	N	0.03729	1.10	3.31	10	5	5	10	6	10	10	4	80		
60	36722	Pandey and Srivastava, 2000	Nickel sulfate	37.93	Mouse (<i>Mus musculus</i>)	4	0/3.57/7.14/14.29	mg/kg bw/d	N	na	5 per w	U	OR	35	d	NR	mo	JV	M	C Lab	REP	REP	SPCL	SM	7.14	14.29	Y	0.025	N	0.00331	1.35	2.71	10	8	10	10	10	10	10	4	92		
61	36722	Pandey and Srivastava, 2000	Nickelous chloride	45.29	Mouse (<i>Mus musculus</i>)	4	0/3.57/7.14/14.29	mg/kg bw/d	N	na	5 per w	U	OR	35	d	NR	NR	JV	M	C Lab	REP	REP	SPCL	SM	3.57	7.14	Y	0.025	N	0.00331	1.70	3.40	10	8	10	10	10	10	10	4	92		
62	14480	Phatak and Patwardhan, 1950	Nickel carbonate	100	Rat (<i>Rattus norvegicus</i>)	4	0/25.5/1	mg/g diet	N	na	ADL	U	FD	4	mo	4	w	GE	F	C Lab	REP	REP	PRWT	WO	1.00		Y	0.115	Y	0.001070	9.30	10	5	10	7	10	4	1	4	71			
63	1260	Chernoff and Kavlock, 1982	Nickel chloride	45.29	Mouse (<i>Mus musculus</i>)	2	0/100	mg/kg bw/d	N	na	DLY	U	GV	5	d	60	d	GE	F	C Lab	REP	REP	PROG	WO	100		N	0.0225	N	0.00304	45.3	10	8	10	10	4	10	10	4	86			
64	19064	Berman and Rehnberg, 1983	Nickel chloride	100	Mouse (<i>Mus musculus</i>)	3	0/85.3/170.7	mg/kg bw/d	N	na	ADL	U	DR	15	d	NR	NR	GE	F	C Lab	REP	REP	PRFM	WO	85.3	170.7	Y	0.0297	Y	5.387	85.3	171	10	5	5	10	7	10	10	4	81		
65	113	Seidenberg et al, 1986	Nickelous chloride	45.29	Mouse (<i>Mus musculus</i>)	2	0/200	mg/kg bw/d	N	na	DLY	U	GV	4	d	NR	NR	GE	F	C Lab	REP	REP	PRWT	WO	200		Y	0.0392	N	0.00479	90.6	10	8	10	10	4	6	10	4	82			
66	14474	Ambrose et al, 1976	Nickel sulfate hexahydrate	100	Dog (<i>Canis familiaris</i>)	4	0/100/1000/2500	mg/kg diet	N	na	1 per d	U	FD	24	mo	6	mo	JV	B	C Lab	REP	REP	TEWT	TE	2500		N	10.8	N	0.4858	112	10	10	5	10	5	10	4	74				
67	14474	Ambrose et al, 1976	Nickel sulfate hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	4	0/250/500/1000	mg/kg diet	N	na	DLY	U	FD	118	d	28	d	GE	F	C Lab	REP	REP	PRWT	WO	500	1000	N	0.000156	N	0.0001	164	327	10	10	5	10	5	10	10	4	84		
68	14474	Ambrose et al, 1976	Nickel sulfate hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	4	0/100/1000/2500	mg/kg diet	N	na	ADL	U	FD	2	yr	28	d	JV	M	C Lab	REP	REP	TEWT	TE	2500		Y	0.367	N	0.0301	205	10	10	5	10	6	10	4	70				
69	66	Schroeder and Mitchener, 1971	Nickel	100	Rat (<i>Rattus norvegicus</i>)	2	0/5	mg/L	N	na	ADL	U	DR	9	mo	21	d	GE	F	C Lab	REP	REP	DEYO	WO	5.00	N	0.344	N	0.03789		0.551	10	5	5	4	5	10	4	10	4	67		
70	19522	Pandey and Singh, 1999	Nickel sulfate hexahydrate	22.33	Mouse (<i>Mus musculus</i>)	3	0/3.57/7.14	mg/kg bw/d	N	na	5 per w	U	GV	35	d	NR	NR	JV	M	C Lab	REP	REP	SPCV	SM	3.57	Y	0.02	N	0.00276		0.797	10	8	5	10	10	4	10	4	81			
71	62	Smith et al, 1993	Nickel chloride	100	Rat (<i>Rattus norvegicus</i>)	4	0/1.33/6.8/31.63	mg/kg bw/d	N	na	ADL	M	DR	23	w	62	d	LC	F	C Lab	REP	REP	DEYO	WO	1.33	Y	0.2689	Y	0.02479		1.33	10	5	10	10	4	10	4	83				
72	19521	Pandey et al, 1999	Nickel sulfate	37.94	Mouse (<i>Mus musculus</i>)	3	0/3.57/7.14	mg/kg bw/d	N	na	5 per w	U	GV	35	d	NR	NR	JV	M	C Lab	REP	REP	TEWT	TE	3.57	Y	0.025	N	0.00331		1.35	10	8	10	10	4	10	4	86				
73	19520	Pandey et al, 1999	Nickel sulfate hexahydrate	22.33	Mouse (<i>Mus musculus</i>)	2	0/7.14	mg/kg bw/d	N	na	5 per w	U	GV	35	d	NR	NR	AD	M	V Lab	REP	REP	RSUC	WO	7.14	Y	0.02	N	0.00276		1.59	10	8	10	10	4	10	4	86				
74	19507	Obone et al, 1999	Nickel sulfate	100	Rat (<i>Rattus norvegicus</i>)	4	0/44.7/111.75/223.5	mg/L	N	na	ADL																																

Appendix 6.1 Mammalian Toxicity Data Extracted for Wildlife Toxicity Reference Value (TRV)

Nickel

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Ref		Reference	Chemical Form	MW%	Test Species	Exposure										Effects				Conversion to mg/kg bw/day				Result		Data Evaluation Score																	
Result #	Ref N.					# of Conc/ Doses	Conc/ Doses	Conc/Dose Units	Wet Weight Reported?	Percent Moisture	Application Frequency	Method of Analyses	Route of Exposure	Exposure Duration	Duration Units	Age	Age Units	Lifestage	Sex	Control Type	General Effect Group	Effect Type	Effect Measure	Response Site	Study NOAEL	Study LOAEL	Body Weight Reported?	Body Weight in kg	Ingestion Rate Reported?	Ingestion Rate in kg or L/day	NOAEL Dose (mg/kg/day)	Data Source	Dose Route	Test Concentrations	Chemical form	Dose Quantification	Endpoint	Dose Range	Statistical Power	Exposure Duration	Test Conditions	Total	
110	36722	Pandey and Srivastava, 2000	Nickel sulfate	37.93	Mouse (<i>Mus musculus</i>)	4	0/3.57/7.14/14.29	mg/kg bw/d	N	na	5 per w	U	OR	35	d	NR	NR	JV	M	C Lab	MOR	MOR	MORT	WO	14.29		Y	0.025	N	0.00331	5.42	10	8	10	10	9	4	10	10	4	85		
111	36722	Pandey and Srivastava, 2000	Nickelous chloride	45.29	Mouse (<i>Mus musculus</i>)	4	0/3.57/7.14/14.29	mg/kg bw/d	N	na	5 per w	U	OR	35	d	NR	NR	JV	M	C Lab	MOR	MOR	MORT	WO	14.29		Y	0.025	N	0.00331	6.47	10	8	10	10	9	4	10	10	4	85		
112	36694	Alexander et al, 1978	Nickel chloride hexahydrate	100	Meadow vole (<i>Microtus pennsylvanicus</i>)	4	0/21/187/1968	mg/kg diet	N	na	ADL	M	FD	14	d	14	d	JV	NR	C Lab	MOR	MOR	MORT	WO	187	1968	Y	0.035	Y	0.0055	29.4	309	10	10	10	10	7	9	6	10	10	4	86
113	1260	Chernoff and Kavlock, 1982	Nickel chloride	45.29	Mouse (<i>Mus musculus</i>)	2	0/100	mg/kg bw/d	N	na	DLY	U	GV	5	d	60	d	GE	F	C Lab	MOR	MOR	MORT	WO	100		N	0.0225	N	0.00304	45.3		10	8	10	10	10	9	4	1	10	4	76
114	113	Seidenberg et al, 1986	Nickelous chloride	45.29	Mouse (<i>Mus musculus</i>)	2	0/200	mg/kg bw/d	N	na	DLY	U	GV	4	d	NR	NR	GE	F	C Lab	MOR	MOR	MORT	WO	200		Y	0.0392	N	0.00479	90.6		10	8	10	10	10	9	4	10	10	4	85
115	14474	Ambros et al, 1976	Nickel sulfate hexahydrate	100	Dog (<i>Canis familiaris</i>)	4	0/100/1000/2500	mg/kg diet	N	na	1 per d	U	FD	24	mo	6	mo	JV	B	C Lab	MOR	MOR	MORT	WO	2500		N	10.8	N	0.4858	112		10	10	5	10	5	9	4	10	10	4	77
116	36331	Cempel and Janicka, 2002	Nickel chloride hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/7.9/29.0	mg/org/d	N	na	ADL	U	DR	90	d	NR	NR	JV	M	C Lab	MOR	MOR	MORT	WO	29		Y	0.21	Y	0.000242	138		10	5	5	10	7	9	4	10	10	4	74
117	14474	Ambrose et al, 1976	Nickel sulfate hexahydrate	100	Rat (<i>Rattus norvegicus</i>)	4	0/100/1000/2500	mg/kg diet	N	na	ADL	U	FD	24	mo	28	d	JV	B	C Lab	MOR	MOR	MORT	WO	2500		Y	0.367	N	0.0301	205		10	10	5	10	6	9	4	10	10	4	78
118	14447	Schroeder et al, 1964	Divalent Nickel	100	Mouse (<i>Mus musculus</i>)	2	0/5	mg/L	N	na	DLY	U	DR	16	mo	21	d	JV	B	C Lab	MOR	MOR	SURV	WO	5.00		N	0.0375	Y	0.0047		0.62	10	5	5	10	6	9	4	10	10	4	73
119	1858	Schroeder and Mitchener, 1975	Nickelous acetate	100	Mouse (<i>Mus musculus</i>)	2	0/5	mg/L	N	na	NR	U	DR	520	d	19-20	d	AD	F	C Lab	MOR	MOR	LFSP	WO	5.00		Y	0.025	N	0.003579		0.716	10	5	5	5	6	9	4	10	10	4	68
Data Not Used to Derive TRV																																											
120	19787	Vyskocil et al, 1994	Nickel sulfate	100	Rat (<i>Rattus norvegicus</i>)	2	0/6.25	mg/kg bw/d	N	na	ADL	U	DR	6	mo	10	w	JV	M	C Lab	GRO	GRO	BDWT	WO	6.25		Y	0.5	Y	0.03120	6.25		10	5	5	10	7	8	4	1	10	4	64
121	1858	Schroeder and Mitchener, 1975	Nickelous acetate	100	Mouse (<i>Mus musculus</i>)	2	0/5	mg/L	N	na	NR	U	DR	520	d	19-20	d	JV	F	C Lab	GRO	GRO	BDWT	WO	5.00		Y	0.0428	N	0.005807	0.678		10	5	5	5	6	8	4	6	10	4	63
122	19461	Nielsen et al, 1979	Nickel chloride trihydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	DLY	U	FD	9	w	21	d	JV	F	C Lab	PTH	ORW	SMIX	LI	50.0		N	0.204	N	0.0186	4.56		10	10	5	10	5	4	4	1	10	4	63
123	19483	Nielsen et al 1982	Nickel chloride trihydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	ADL	U	FD	11	w	NR	NR	JV	F	C Lab	BIO	CHM	LIPD	PL	50.0		N	0.204	N	0.0186	4.56		10	10	5	10	5	1	4	1	10	4	60
124	19479	Nielsen, 1980	Nickel chloride trihydrate	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	DLY	U	FD	10	w	NR	NR	JV	F	C Lab	BIO	CHM	HMCT	BL	50		N	0.204	N	0.0186	4.56		10	10	5	10	5	1	4	1	10	4	60
125	15690	Nielsen et al, 1980	Nickel chloride	100	Rat (<i>Rattus norvegicus</i>)	3	0/5/50	ug/g diet	N	na	DLY	U	FD	10	w	NR	NR	JV	F	C Lab	BIO	CHM	HMGL	BL	50.00		N	0.2024	N	0.018478	4.56		10	10	5	10	5	1	4	1	10	4	60
126	136	Gershbein et al, 1983	Nickelous chloride	100	Rat (<i>Rattus norvegicus</i>)	2	0/75	mg/kg diet	N	na	ADL	U	FD	80	d	44	d	JV	M	C Lab	PTH	HIS	GHIS	MT	75		Y	0.4700	N	0.03693	5.89		10	10	5	10	6	4	4	1	10	4	64
127	14476	ODell et al, 1970	Nickelous carbonate	100	Cattle (<i>Bos taurus</i>)	3	0/365/1835	mg/org/d	N	na	1 per d	U	FD	6	w	NR	NR	LC	F	C NR	BEH	FDB	FCNS	WO	1835		N	272	Y	7.300	6.75		10	10	5	10	6	4	4	1	10	4	64
128	36732	Tomokuni and Ichiba, 1990	Nickel nitrate hexahydrate	100	Mouse (<i>Mus musculus</i>)	2	0/57	mg/L	N	na																																	