

Training on the IEUBK Model, Adult Lead Methodology, and Recent Lead Risk Assessment Updates

TRW Lead Committee



- Introduction (why do we need models?)
- The IEUBK Model -- structure and components
- The U.S. EPA Adult Lead Methodology (ALM)
- Discuss IEUBK & ALM inputs (review data entry windows and input variables, and why some are controversial)
- Risk Assessment Issues and Guidance
- PbB calculation, risk calculation, & PRG

What is a Biokinetic Model?

- Biokinetic models assess the routes of environmental exposure to a substance and determine the distribution of this substance among the various body tissues in humans.
- Biokinetic models work best when there is a known effect that is associated with a specific tissue concentration in humans. -e.g., impaired nerve conduction velocity in children at 10 µg Pb/dL blood.
- Biokinetic models also enable the risk assessor to predict the relative effect of an increase in body tissue that might result from a specific increase in environmental exposure.
 - -e.g., the expected blood lead concentration that would result from an increase in soil lead concentration of 500 mg/kg.

Lead Risk Assessment is Different

- In comparison to most other environmental contaminants, the degree of uncertainty about the health effects of lead is quite low.
- Some of these effects, particularly changes in the levels of certain blood enzymes and in aspects of children's neurobehavioral development, may occur at blood lead levels so low as to be essentially without a threshold.
- EPA decided that it was inappropriate to derive a Reference Dose (RfD) for lead.
- EPA regulates lead exposure by using a biomarker (blood lead concentration).
- Environmental exposures to lead are <u>modeled</u> to predict blood lead levels associated with those exposures.
- CDC established 10 μ g/dL as the Federal level of concern in 1991.

OSWER Lead Risk Assessment Policy

- The IEUBK Model as the primary tool to generate residential riskbased soil cleanup levels.
- OSWER's risk reduction policy is for no child to have greater than a 5% probability of having a blood lead level >10 μ g/dL.
- Modeling is used to associate environmental exposures with risk and inform cleanup decisions (relative to OSWER's risk reduction goal).
- In general, blood lead survey data should not be used as the only basis for cleanup decisions.

Purpose of the Lead Models

IEUBK (Integrated Exposure Uptake Biokinetic Model)

- Predicts the blood lead levels in children (under 7 years old) who are exposed to environmental lead from many sources
- Predicts the risk (probability) that a typical or hypothetical child exposed to specified media lead concentrations will have a blood lead level $\geq 10 \ \mu g/dL$ (the blood lead level of concern)
- Predicts PRG (cleanup levels) for various media for residential land use

ALM (Adult Lead Methodology)

- Predicts the risk of elevated blood lead levels in non-residential settings (adult exposure to soil; ultimate receptor is fetus)
- Predicts PRG (cleanup levels) for soil in non-residential land use

Characteristics of the IEUBK Model

- While IEUBK model risk assessments are more complex than the typical Superfund risk assessment approach, the IEUBK model is not as complex as variance propagation approaches (PRA)
- The IEUBK model employs more site-specific information than other EPA risk assessment models
- The IEUBK model performs well when comparing predicted and observed blood lead levels (*Hogan et al., 1998. Integrated Exposure Uptake Biokinetic Model for Lead in Children: Empirical Comparisons with Epidemiologic Data. Environmental Health Perspectives, Vol. 106 No. S6*)

Intake – Uptake – Biokinetic Relationship

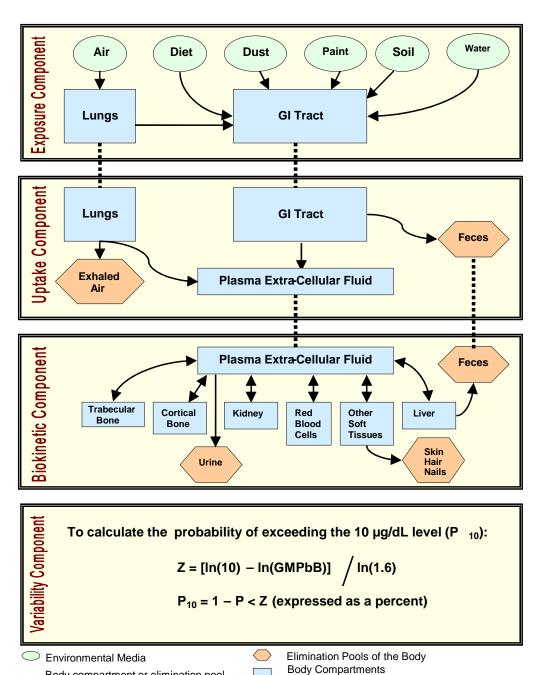
Daily **Intake** of lead is calculated as follows: Intake = Media Concentration x Media Intake Rate For example: µg lead/day = (µg lead / g of media) x (g of media / day)

Uptake is calculated based on media-specific absorption values (defaults are available): Uptake = Intake x Absorption Factor

Biokinetic module estimates transfer rates for Pb moving between compartments and through elimination pathways to derive a predicted long-term steady state geometric mean PbB concentration.

In the final step, the **Probability** module estimates a plausible distribution of PbB concentrations for a given GSD. The distribution is centered on the geometric mean PbB concentration calculated by the Biokinetic Module.





Body compartment or elimination pool required in more than one component

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History of the IEUBK Model Development

IEUBK is the product of many years of development 1985-89: Initially Office of Air Quality Planning Standards

- 1989: Development by Superfund following SAB review 1989-2001: DOS version (0.99d) development.
- 1994-2001: Release of 0.99d version by Superfund with input from EPA, ATSDR, CDC, and SAB.
- 1998: Independent Validation and Verification (IV&V)
 1997-2001: IEUBK (0.99d) was converted to Windows
 2001-present: IEUBKwin 1.0 and IEUBKwin 1.1
 2005: NAS review

Independent Reviews of the IEUBK

The reviewers have generally found that the model was scientifically sound and useful for lead risk assessment

- 1990 SAB review for NAAQS
- 1992 SAB review and External Peer Review of model
- 1998 Independent Validation and Verification
- 1998 SAB review for TSCA Section 403 Regulation
- 2005 National Academies of Science (NAS) review for Coeur d'Alene site report

Evaluation and Validation of the IEUBK

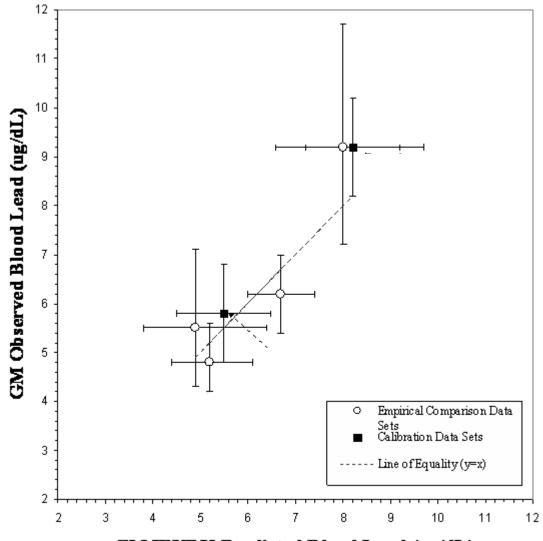
IV&V evaluated the following:

- 1. Scientific underpinnings of the model structure
- 2. Adequacy of parameter estimates
- 3. Mathematical relationships (as computer code)
- 4. Empirical comparisons (predicted vs. observed)

The process and results of the IEUBK validation are available online (TRW web site)

1994 Validation Strategy for the IEUBK1998 Empirical Comparisons Manuscript (Hogan et al., 1998)

Comparison of IEUBK Predictions and Observed PbB



GM IEUBK Predicted Blood Lead (ug/dL)

Correspondence of observed and IEUBK model-predicted blood lead concentrations by site: Kansas/Missouri, Illinois, Pennsylvania, 1991. The solid points are the GMs, whereas the open points provide 95% CIs for the GMs. Adapted from Hogan et al., 1998

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Comparison of IEUBK Predictions and Observed PbB

Comparison of Observed and Predicted Geometric Mean Blood Lead and Risk of Exceeding 10 µg/dL for Three Community Blood Lead Studies

Dataset	N	Observed Blo	ood Lead (µg/dL)	Model Predictions (µg/dL)		
		GM (95% CI)	Percent >10 (95% CI)	GM (95% CI)	Percent >10 (95% CI)	
Galena, KA Jasper Co, MI ^a	111	5.2 (4.5-5.9)	20 (13-27)	4.6 (4.0-5.3)	18 (11-25)	
Madison Co, IL ^a	333	5.9 (5.5-6.4)	19 (15-23)	5.9 (5.4-6.3)	23 (19-28)	
Palmerton, PA ^b	34	6.8 (5.6-8.2)	29 (14-44)	7.5 (6.6-8.6)	31 (16-47)	

Excerpts from Air Criteria Document for Lead (October 2006). Original data from Hogan et al. (1998)

CI, confidence interval; GM, geometric means ^aChildren away from home ≤10 hours/week ^bChildren away from home ≤20 hours/week

IEUBK Exposure Module Components

Media Concentrations for Input							
Soil	Soil must be sampled. Site- specific data required.	Refer to the IEUBK User's Guide and 1994 Guidance Manual for additional information on this input parameter.					
Dust	Site-specific data or a value can be derived from soil concentration using multiple source analysis.	Refer to the IEUBK User's Guide and 1994 Guidance Manual for additional information on this input parameter.					
Air (default)	0.1 µg/m ³	Ratio of indoor to outdoor air lead concentration is 30%. Site-specific data may be substituted.					
Drinking Water (default)	4 µg/L	Site-specific data may be substituted.					

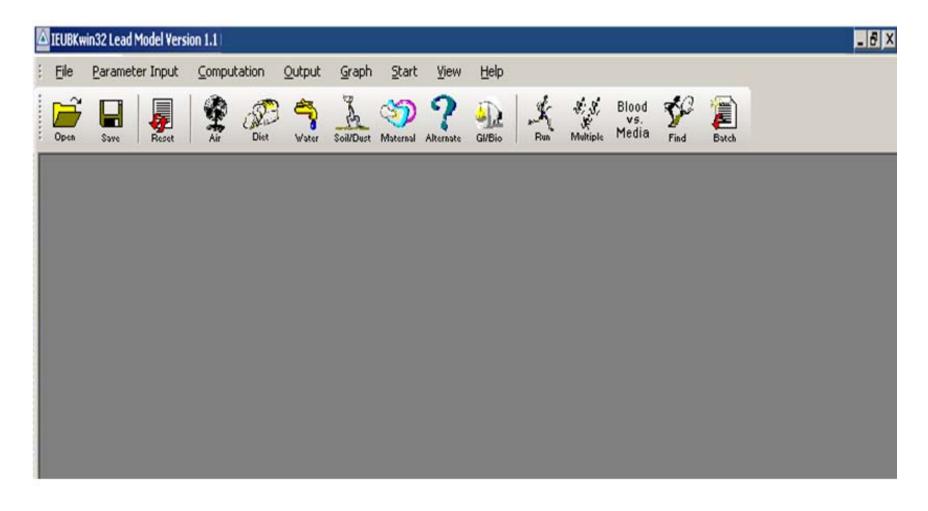


IEUBK Components (continued)

Media		Age-specific Intake Rates						Comments
	0-1	1-2	2-3	3-4	4-5	5-6	6-7	
	year	yrs	yrs	yrs	yrs	yrs	yrs	
Soil/dust (mg/day)	85	135	135	135	100	90	85	Default values recommended. Intake is apportioned 55% dust & 45% soil
Air (m³/day)	2	3	5	5	5	7	7	Default values recommended
Drinking Water	0.2	0.5	0.52	0.53	0.55	0.58	0.59	Default values recommended
(L/day)								
Diet (µg Pb/day)	2.26	1.96	2.13	2.04	1.95	2.05	2.22	Site-specific data may be used to assess exposure to fish, game, or home-grown produce.
Alt. Source	Site-specific data may be used to account for intake of lead in other sources						or	Refer to the IEUBK User's Guide and 1994 Guidance Manual for more information

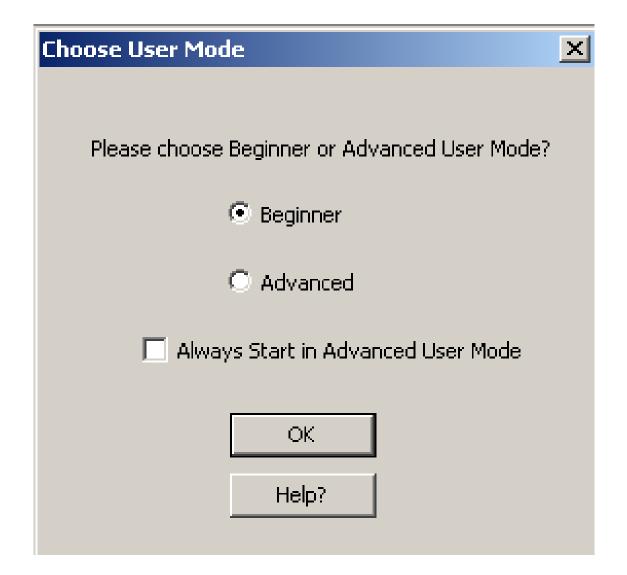


IEUBK Model screenshot showing input icons for pathways





Choosing the User Mode





ata								
Indoor air lead concentration (percentag	e of outdo	oor): 30	_				OK	
Outdoor Air Pb Concentration (ug/m²):							Cancel	
Constant Value: 0.1							Reset	
C Variable Values							Help?	
nput for different age groups	AGE (Years)							
	0-1	1-2	2-3	3-4	4-5	5-6	6-7	
Outdoor Air Pb Concentration (µg/m²):	0.1	0.1	0.1	0.1	0.1	0.1	0.1	
Time Spent Outdoors (hr/day):	1	2	3	4	4	4	4	
Ventilation Rate (m³/day):	2	3	5	5	5	7	7	
	32	32	32	32	32	32	32	

TRW Homepage: http://www.epa.gov/superfund/health/contaminants/lead/index.htm

Dietary Exposure Input

Dietary Data				? ×
		AGE (Years)		
Dietary Lead Intake (ug/day)	0-1 1-2 26 1.96	2-3 3-4 2.13 2.04	4-5 5-6 6-7 1.95 2.05 2.22	<u>0</u> K
DIETARY VALUES				<u>C</u> ancel
Use alternate dietary values?	No	C Yes		Help?
Cone	centration (ug P	b/g) Percent of Food (Class	
Home Grown Fruits	0	0	(% of all fruits)	
Home Grown Vegetables	0	0	(% of all vegetables)	
Fish from Fishing	0	0	(% of all meat)	
Game Animals from Hunting	0	0	(% of all meat)	
Ethnic Preferences				
Regional Preferences				
GI Values / Bioavailability GI / Bio Change Values		TRW Ho http://wv	mepage: vw.epa.gov/superfund/progra	ams/lead



Drinking Water Exposure Input

inking Water Data	?
Water Consumption (L/day) AGE (Years) 0-1 1-2 2-3 3-4 4-5 5-6 6-7 0.2 0.5 0.52 0.53 0.55 0.58 0.59	OK Cancel
Use alternate water values? No If No, please enter the lead concentration in drinking water (µg/L): 4 Yes If Yes, please fill in the information below.	Help?
LEAD CONCENTRATION IN DRINKING WATER Percent of Total Consumed as First Draw: 50 Concentration of Lead in First Draw (µg/L): 4 Concentration of Lead in Flushed (µg/L): 1 Percentage of Total Consumed from Fountains: 15 Concentration of Lead in Fountain Water (µg/L): 10	
GI / Bio Change Values TRW Homepage: http://www.epa.gov/superfund/health/contaminant	ts/lead/index.htm



Soil and Dust Exposure Input

pecific Soil Dust Data							
oil/Dust Ingestion Weighting	Factor (per	cent soil):	45				ОК
Outdoor Soil Lead Concentr	ation (µg/g)	_ Ind	loor Dust Le	ad Concer	tration (µg/	g)	Cancel
			C Constar	nt Value	200		Reset
Constant Value 20	U		C Variable	Values			Help?
C Variable Values			Multiple	Source An	alysis	Set	
			Multiple	Source Av	g: 150	_	
Soil/Indoor Dust Concentral	tion (μg/g) —		<u>م</u>	iE (Years)			
	0-1	1.2	2-3	3-4	4-5	5-6	6-7
Outdoor Soil Lead Levels:	200	200	200	200	200	200	200
Indoor Dust Lead Levels:	150	150	150	150	150	150	150
Amount of Soil/Dust Ingeste	ed Dailv (g/d	avì					
			A	GE (Years)			
	0-1	1-2	2-3	3-4	4-5	5-6	6-7
Total Dust + Soil Intake:	0.085	0.135	0.135	0.135	0.100	0.090	0.085
GI Values/Bioavailability		TRWH	omepage:				
GI / Bio Change	202012		www.epa.g				



Multiple Source Analysis Detail

Multiple Source Analysis				<u>? ×</u>
Contribution of soil lead to indoor house (conversion factor): Contribution of outdoor airborne lead to household dust lead (conversion factor	indoor	0.7		<u>D</u> K <u>C</u> ancel Help?
Indoor Dust Lead Sources				
Use Alternate Indoor Dust Lead	Sources?	No	O Yes	
Co	ncentration (ug	Pb/g)	Percent	
Household Dust (average)	150		100.000	
Secondary Occupational Dust	1200		0.000	
Dust at School	200		0.000	
Dust at Daycare	200		0.000	
Second Home Dust	200		0.000	
Lead-based Paint in Home	1200		0.000	
	TRW Ho http://ww	mepage: ww.epa.gov/su	uperfund/prog	grams/lead



Bioavailability Information Input

	GI Values/Bio	availability Inform	mation		? ×
Drinking Water Data Water Consumption (L/day)= 0-1 1-2 0.2 0.5 0 Use alternate water values? No If No, please enter 1 Yes If Yes, please fill in LEAD CONCENTRATION IN Percent of To	MEDIA Soil Dust Water Diet Alternate	ABSORPTION FRACTION PERCENT 30 30 50 50 50 0 TRW Homepage:	Acess alternate bioavaila paramenters? FRACTION PASSIVE/ TOTAL ACCESSIBLE 0.2	HALF SATURATION Level (µg/day)	OK Cancel Reset Help?
Percentage of 1	Lead in Flushed (µg/l Total Consumed from Fo of Lead in Fountain Wa TRW Hom	L): 1 ountains: 15 ter (μg/L): 10 epage:	Ith/contaminants/lead/index.htm		



Alternate Exposure Input

X

Alternate Source Intake

If you change the alternate source intake, remember to change the Alternate Source Absorption Percent on the GI/Bioavailability data entry screen.

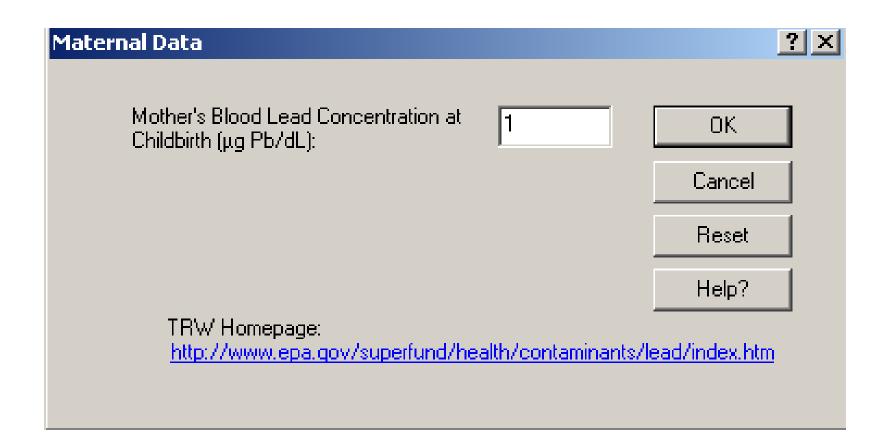
Its default value is 0.0% which must be changed if intakes are not 0.0.

OK

Alte	ernate So	ource Dat	a					? ×
	Alternate	e Lead Intai	ke (μg/day)	AGE (Years)			ОК
	0-1	1-2	2-3	3-4	4-5	5-6	6-7	Cancel
	0	0	0	0	0	0	0	Reset
								Help?
	GI Value	es / Bioavai	ilability ——					
	GI 7	Bio C	ihange Valu	ies				
	TRW H	lomepage:	http://ww	w.epa.go	v/superfu	nd/health/	contaminant	ts/lead/index.htm

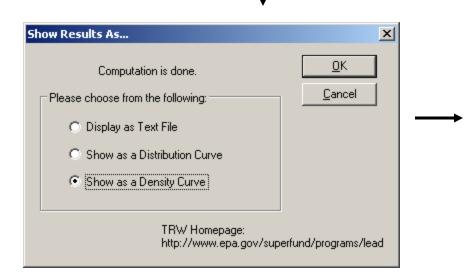


Maternal Exposure Input



Run Risk Calculation (forward equation)

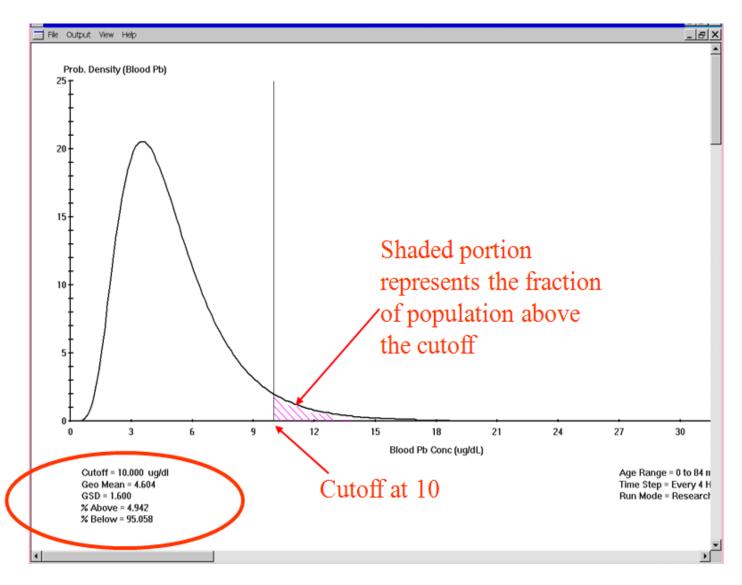
Run the Model	? >
Enter the Result File Name RunModel	Run
Export data into a Spreadsheet Format 🛛 🔲	Cancel
TRW Homepage: http://www.epa.gov/superfund/health/contaminants/lead/index.htm	Help?



Cutoff, GSD, and Comment Information	<u>? ×</u>
Select Age Group for Graph 0 to 84 months	<u>S</u> how Graph
	<u>C</u> ancel
Parameter Change	
Change Cutoff 10	
Change GSD(Geometric Standard Deviation) 1.6	
Comment	
TRW Homepage:http://www.epa.gov/superfund	d/programs/lead



Output from the IEUBK Model





Find Soil Pb Concentration	×
Select Age Group for Graph 0 to 84 months .	Find Cancel Help?
Change Cutoff 10 µg/dl 1 Change GSD (Geometric Standard Deviation) 1.6	Depending on the values enter,
Probability of Exceeding the Cutoff (PC) 5 % "	Find Soil Pb Concentration
	Find
Soil and/or Dust Concentration 0 PPM	Select Age Group for Graph 0 to 84 months Cancel
TRW Homepage: <u>http://www.epa.gov/superfund/health/contaminants/l</u>	Help?
	Parameter Change Please note
	Change Cutoff 10 µg/dl Depending on the values
	Change GSD (Geometric Standard Deviation) 1.6 enter, calculating PRG
	Probability of Exceeding the Cutoff (PC) 5 % may take a few moments.
	Soil and/or Dust Concentration 418 PPM
	TRW Homepage: <u>http://www.epa.gov/superfund/health/contaminants/lead/index.htm</u>



Sensitivity Analysis

- Predicted PbB and total lead uptake were most sensitive to the amount of soil/dust ingested per day.
- Predicted PbB and total lead uptake were moderately sensitive to the following (listed in decreasing relative sensitivity):
 - absorption fraction for soil dust and diet,
 - soil lead concentration,
 - indoor dust lead concentration,
 - dietary lead concentration,
 - contribution of soil lead to indoor dust lead, and
 - half-saturation absorbable intake (based on output-input ratio).
- The predicted probability of exceeding a specified level of concerns is very sensitive to changes in the GSD.

IEUBK Strengths and Limitations

- Strengths:
 - Integrates multimedia exposure and relates it to a well characterized biomarker of effect
 - Risk predictions and PRG over a range of exposure scenarios
 - > Inputs tailored to support Superfund site risk assessment
 - Risk information complementary to a public health (PbB) study or when no public health (PbB) study is available
- Limitations:
 - Cannot assess short-term, periodic or acute exposures (exposures must be for at least 1 day per week for 90 consecutive days)
 - Cannot assess pica exposures
 - > Cannot assess dust exposures using loading data
 - Cannot assess age groups >7 years

윶 EPA Adult Methodology (ALM)

- Adopted and modified from Bowers et al. (1994)
- Uses a simplified biokinetic slope factor (BKSF)
- Slope relates change in PbB ($\mu g/dL$) per $\mu g/day$ Pb absorbed
- Exposure and other variables differ from IEUBK (IR, bioavailability, etc.)



ALM Spreadsheet (Risk Calculation)

Calculations of Blood Lead Concentrations (PbBs)

U.S. EPAT echnical Review Workgroup for Lead, Adult Lead Committee

Version date 6/21/09

Variable	Description of Variable	Units	GSDi and PbBo from Analysis of NHANES 1999-2004	GSDi and PbBo from Analysis of NHANES III (Phases 1&2)
PbS	Soil lead concentration	ugłg or ppm	2240	1235
R _{fetal/maternal}	Fetal/maternal PbB ratio		0.9	0.9
BKSF	Biokinetic Slope Factor	ug/dL per ug/day	0.4	0.4
GSDi	Geometric standard deviation PbB		1.8	2.1
PbBo	Baseline PbB	ug/dL	1.0	1.5
IR_s	Soil ingestion rate (including soil-derived indoor dust)	g/day	0.050	0.050
IR_{S+D}	Total ingestion rate of outdoor soil and indoor dust	g/day		
Ws	Weighting factor; fraction of IR5.0 ingested as outdoor soil			
Ksd	Mass fraction of soil in dust			
AF _{S, D}	Absorption fraction (same for soil and dust)		0.12	0.12
EF _{S,D}	Exposure frequency (same for soil and dust)	daysłyr	219	219
AT _{S, D}	Averaging time (same for soil and dust)	days/yr	365	365
PbB _{adult}	PbB of adult worker, geometric mean	ug/dL	4.2	3.3
PbB _{fetal, 0.95}	95th percentile PbB among fetuses of adult workers	ug/dL	10.0	10.0
PbB _t	Target PbB level of concern (e.g., 10 ug/dL)	ug/dL	10.0	10.0
P(PbB _{fetal} > PbB _t)	Probability that fetal PbB > PbB1, assuming lognormal distribution	×	5.0%	5.0%

Recommended PbB₀ and GDS_i Input

 $Calculations \ of \ Preliminary Remediation \ Goals \ (PRGs)$

U.S. EPA Technical Review Workgroup for Lead, Adult Lead Committee Version date 6/21/09 EDIT RED CELLS

GSDi and PbBo from GSDi and PbBo from Analysis of NHANES Analysis of NHANES Variable Description of Variable 1999-2004 III (Phases 1&2) Units PbB_{feul, 0.95} 95th percentile PbB in fetus 10 10 uałdL 09 09 Retainaternal Fetal/maternal PbB ratio **Biokinetic Slope Factor** ug/dL per BKSF 04 0.4 uq/day. GSD_i 1.8 Geometric standard deviation PbB PbB_o 10 15 **Baseline PbB** ualdL $\mathbb{IR}_{\mathbb{R}}$ 0.050 0.050 Soil ingestion rate (including soil-derived indoor dust) o/dau AF_{s. d} 0.12Absorption fraction (same for soil and dust) 012EF_{S.D} 219 219Exposure frequency (same for soil and dust) daystyr AT_{s n} Averaging time (same for soil and dust) 365 365 daysiyr PRG. 2.2401.235 ppm

Guidance for the IEUBK and ALM

- Model documentation (user's guides and validation information)
- Short Sheets
 - > MSD
 - Ingestion Rate
 - > Sieving
 - Sampling Guidance
 - Small Arms Firing Range
 - Exposure Point Concentration
 - Residential Dust Guidance
- Residential Sites Handbook
- TRW Bioavailability Guidance
- Frequently Asked Questions (FAQs)

TRW Lead Committee Web Site: www.epa.gov/superfund/lead

Soil Lead Bioavailability Guidance

Guidance for Evaluating the Oral Bioavailability of Metals in Soils for Use in Human Health Risk Assessment

OSWER 9285.7-80

United States Environmental Protection Agency



Guidance for Evaluating the Oral Bioavailability of Metals in Soils for Use in Human Health Risk Assessment



OSWER 9285.7-77 May 2007

ESTIMATION OF RELATIVE BIOAVAILABILITY OF LEAD IN SOIL AND SOIL-LIKE MATERIALS USING *IN VIVO* AND *IN VITRO* METHODS

Assessing Risk Example Exposure Scenarios

- 1. Site exposure to lead for a residential community where single (arithmetic mean) media concentrations is available
 - ➢ Risk: IEUBK single run mode
 - > PRG: IEUBK find media concentration
- 2. Residential exposure to lead in soil with individual residence media concentrations
 - ➢ Risk: IEUBK batch mode run
 - PRG: IEUBK find media concentration
- 3. Exposure to lead contaminated soil on an industrial property
 - ➢ Risk: ALM
 - > PRG: ALM
- 4. Adolescent recreational exposure to contaminated soil
 - Risk: ALM
 - > PRG: ALM



1. IEUBK: Single run with 500 ppm soil and default (old) dietary data. Risk calculation.

2. IEUBK: Single run with 500 ppm soil and new dietary data (attached). Risk calculation.

3. IEUBK: PRG calculation using new dietary data and modified drinking water value. Also saving and reloading a data file and reset all parameters.

4. IEUBK: Multiple runs for soil range using new dietary data and modified drinking water value. Plot of risk calculations.

5. IEUBK: Find media concentration for soil without and with new dietary data and interpretation of GM PbB output.

6. IEUBK: Creating a batch mode input file from a spreadsheet file (attached).

7. IEUBK: Running a batch mode to calculate risk.

8. ALM: Example data entry for non-residential scenario. Calculation of risk and PRG.



ALM example scenario: non residential exposure scenario for a soil-intensive contact scenario. What is PRG?

Use $PbB_0 \& GSD_i$ from recent NHANES and IRs of 100 mg/day

PRG = Approx 620 ppm



Is the following health protective? Residential soil PRG is 600 ppm for future use. PbA = $0.05 \ \mu g/m^3$, PbW = 1 ppb, Soil bioavailability = 21%.

Enter media data and use new dietary values

Use find media concentration

Yes, PRG is <5% NTE 10 μ g/dL



Is the following health protective?

PRG of 2500 ppm for a fire monitoring station occupied by adults 7 days per week for 4 months (16 weeks) of the year.

Use ALM with EF = 112 days/year

IRs=50 mg/day

No, PRG is >5% NTE 10 μ g/dL



Assuming that 2500 ppm is the selected soil lead concentration for those fire monitoring stations (occupied by adults), what is the maximum duration that they can be occupied (assuming continuous exposure)?

IRs = 50 mg/day

Reduce EF to achieve is <5% NTE 10 μ g/dL

Approximately 15 weeks



EPA provides risk tools and guidance to assess lead exposure at hazardous waste sites. The TRW Lead Committee is available to support users when questions or when novel applications arise

- Evaluate & develop models and other risk tools
- Provide technical support for the development and implementation of EPA guidance on lead
- Review application of risk assessment tools
- Provide technical assistance to end users for use of non-standard (site-specific) values

TRW Lead Committee Members are EPA staff from Regions, Headquarters, and Labs



Co-Chairs

- Mary Ballew (Region 1)
- •Mike Beringer (Region 7)
- Jim Konz (OSRTI HQ)

TRW Web page: www.epa.gov/superfund/lead

Contact the TRW hotline

- Send an e-mail to pbhelp@epa.gov
- Call the toll-free TRW hotline at 1-866-282-8622