

Predicting Blood Lead Levels by Modeling Soil Lead Exposure Frequency and Duration – A Work in Progress –

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Agency for Toxic Substances and Disease Registry

Division of Community Health Investigations

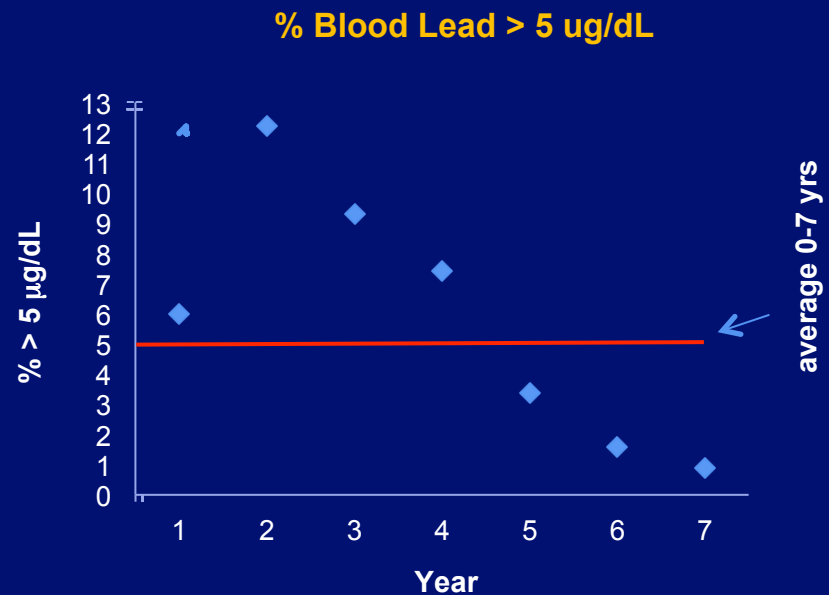


Background

- CDC blood lead reference value lowered to 5 $\mu\text{g}/\text{dL}$ (2012)
- EPA's Integrated Exposure Uptake Biokinetic (IEUBK) model is currently used for predicting blood lead levels (BLLs)
- ATSDR is investigating using other physiologically based pharmacokinetic (PBPK) models for short-term lead exposures
 - Currently testing the All Ages Lead Model (AALM)

Need for more investigation

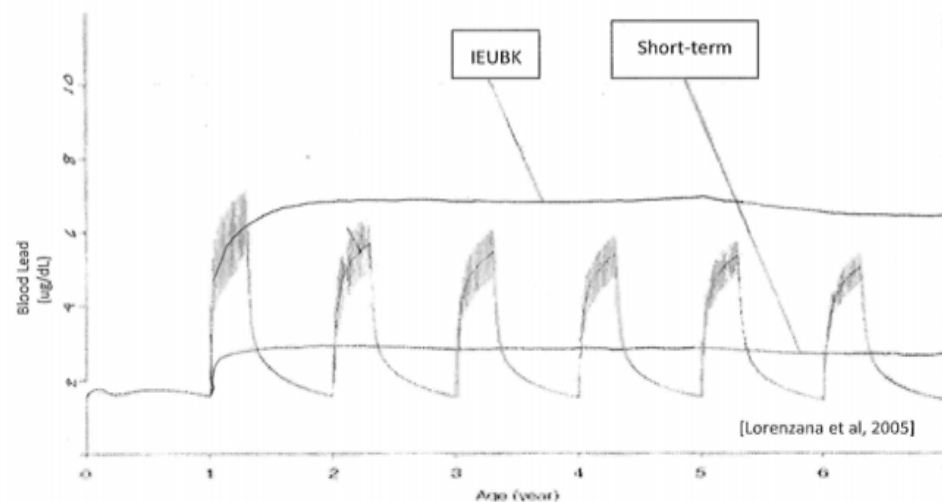
- Need to model shorter-term and less frequent exposures
- **IEUBK model**
 - One year minimum exposure duration
 - Default exposure duration has been averaged 0-7 years



Need for more investigation

- Recent PBPK models allow for more descriptive exposure scenarios and averaging times.

PBPK model of lead exposure with different averaging times



Public Health Implications

Using the AALM provides:

- An estimate of BLLs based on site-specific exposure conditions, including seasonality, frequency, duration, and concentration.
- A means of describing the degree of concern based on the relationship to 5 $\mu\text{g}/\text{dL}$ using probabilities.
- An opportunity for ATSDR to design site-specific interventions, based on site-specific scenarios, to reduce lead exposures.

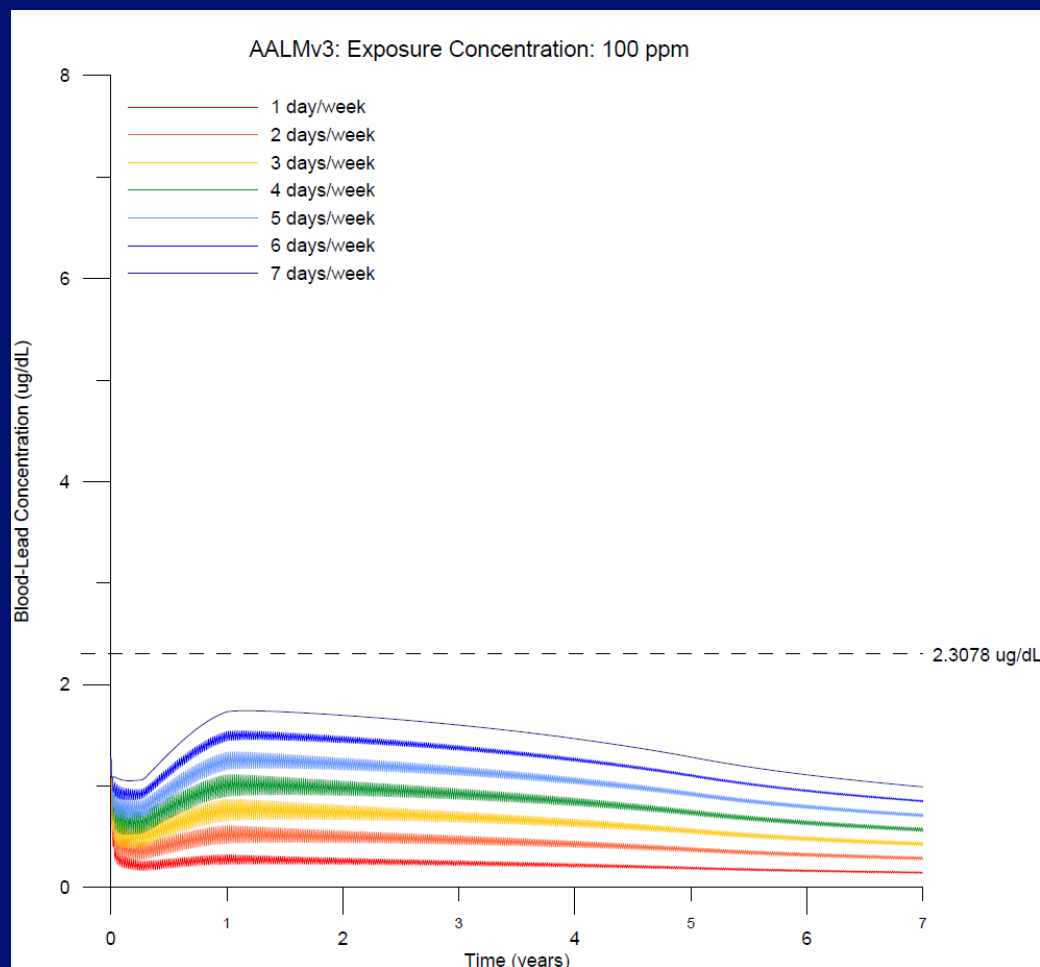
ATSDR activities (completed)

- **Investigated PBPK models for short-term, intermittent, and seasonal exposures.**
- **During *beta* testing of the EPA All Ages Lead Model (AALM) worked with Syracuse Research to develop model execution files to predict ATSDR exposure scenarios.**
 - Modeled blood lead levels (BLLs) for different frequencies, durations, and soil lead levels.
 - Modeled seasonal and day-care exposure scenarios.
 - Derived probabilities of simulated BLLs (% probability of exceeding 5 $\mu\text{g}/\text{dL}$ BLL).

PBPK Results – AALMv3

predicted BLL by exposure frequency and duration

- Soil Concentration: 100 ppm
- Exposure from birth to age 7.
- Predicted geometric mean BLL does not exceed a 5% probability of exceeding $5 \mu\text{g/dL}$ for any exposure frequency or duration.

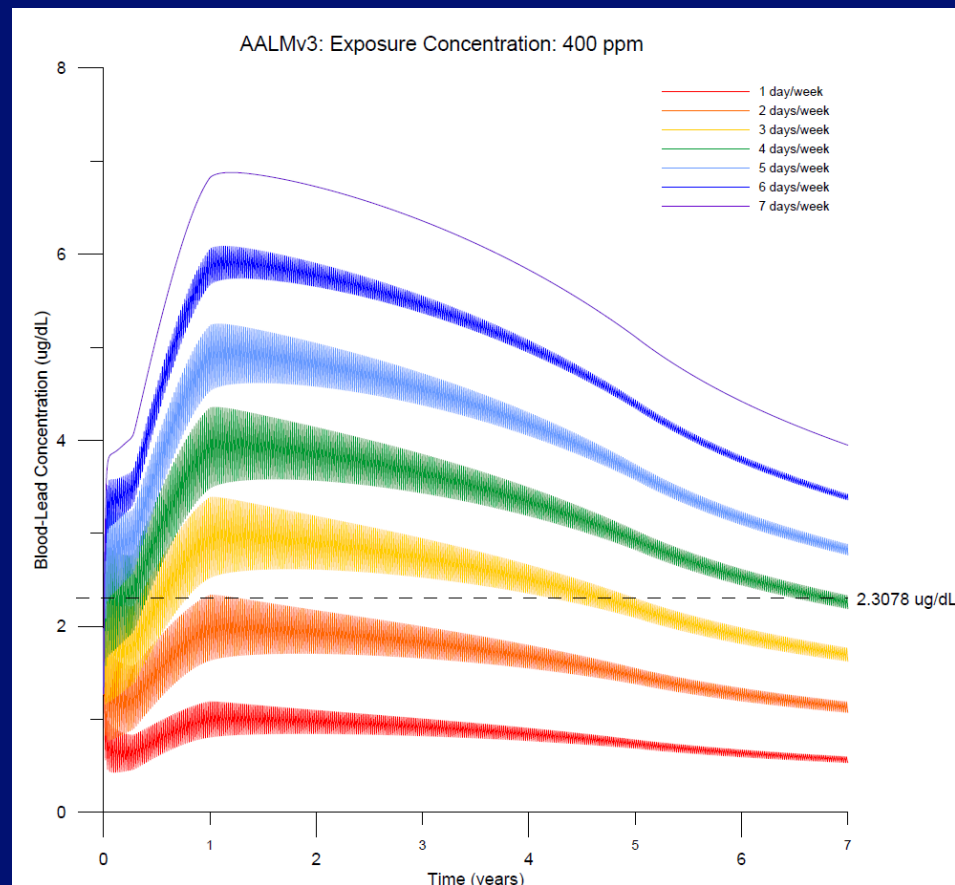


* The 2.3078 $\mu\text{g/dL}$ line is the geometric mean resulting in a probability of 5% of BLLs equaling $5 \mu\text{g/dL}$

PBPK Results – AALMv3

predicted BLL by exposure frequency and duration

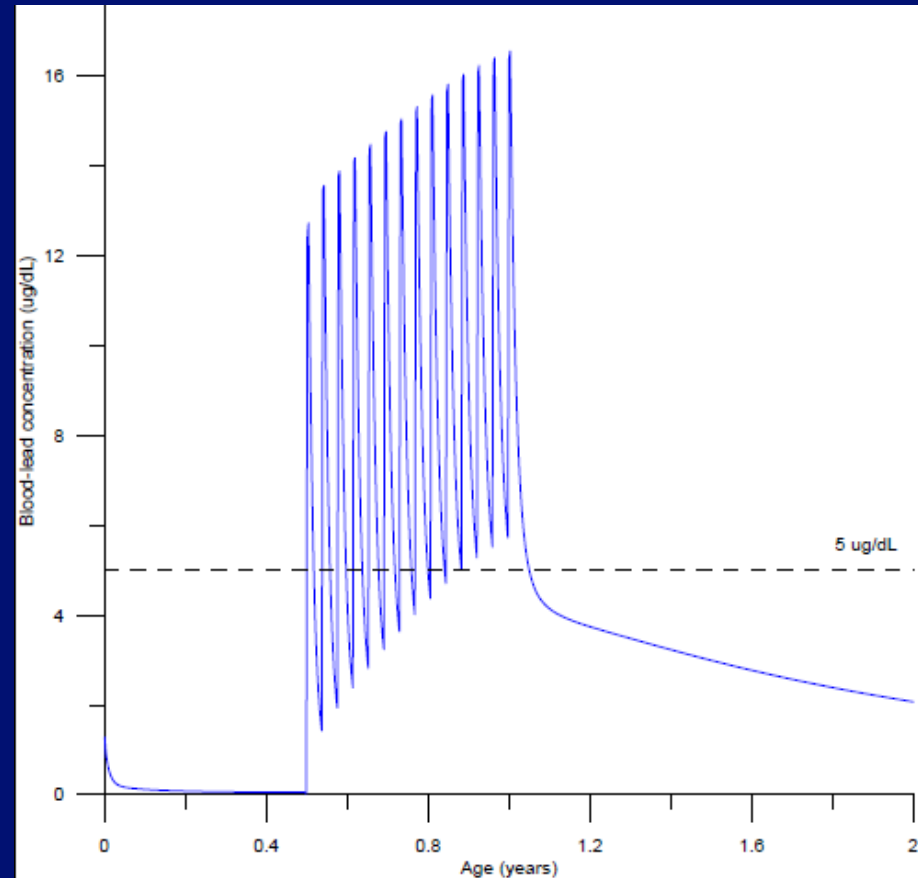
- Soil Concentration: 400 ppm
- Exposure duration: birth to age 7
- Predicted geometric mean BLL does not exceed $5 \mu\text{g/dL}$ for exposures occurring 2 days/week or less for all ages.
- Exposures occurring 3 days/week may result in a greater than 5% probability of a BLL exceeding $5 \mu\text{g/dL}$ for ages 6 months to about 5 years.
- Exposures occurring 4 days/week or more may result in a greater than 5% probability of a BLL exceeding $5 \mu\text{g/dL}$ for ages 7 and under.



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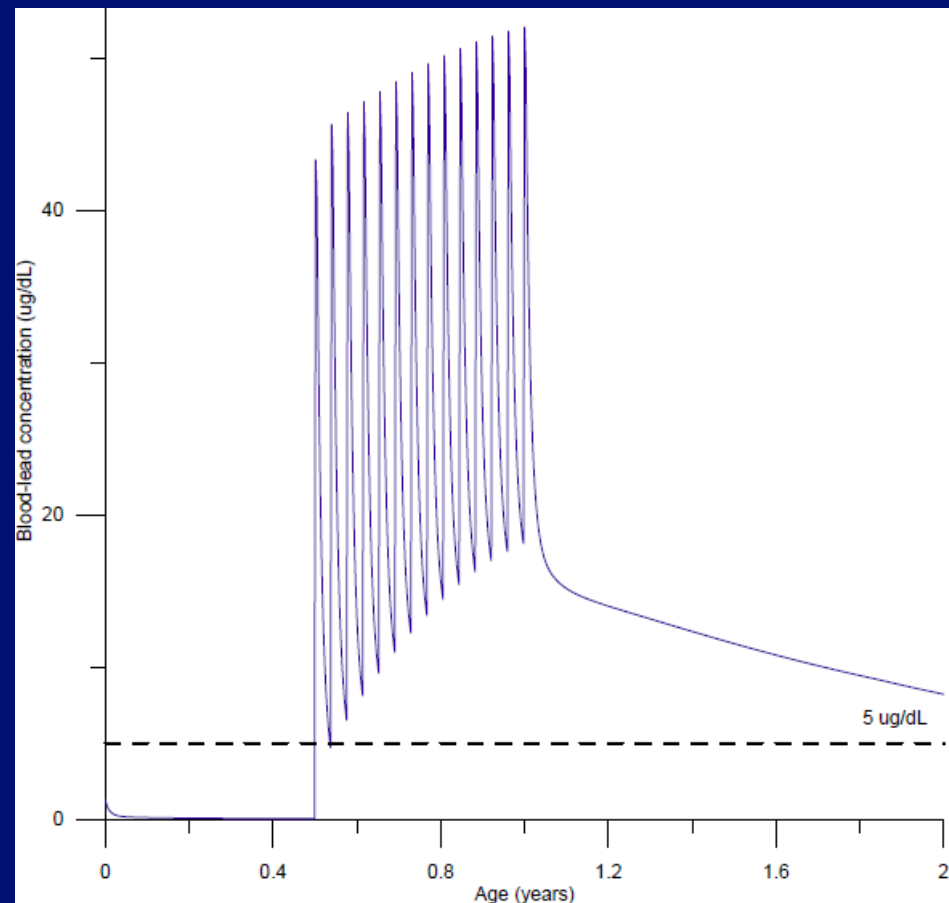
Simulation of Pica Exposure Scenario – AALMv3

- Soil Concentration: 100 ppm
- Pica Ingestion Rate: 5000 mg soil/day
- Exposure Duration: 6 months – 1 year
- Exposure Frequency: once every two weeks
- Peak BLLs: 12 – 16 $\mu\text{g}/\text{dL}$
- **Between pica episodes, BLLs return to below 5 $\mu\text{g}/\text{dL}$ for ages 6 – 10 months.**
- BLLs fall below 5 $\mu\text{g}/\text{dL}$ quickly after end of pica episodes.



Simulation of Pica Exposure Scenario – AALMv3

- Soil Concentration: 400 ppm
- Pica Ingestion Rate: 5,000 mg soil/day
- Exposure Duration: 6 months – 1 year
- Exposure Frequency: once every two weeks
- Peak BLLs: 43 – 52 ug/dL
- **Between pica episodes, BLLs generally stay above 5 ug/dL from the start of the pica episodes until over one year after they end.**



Current ATSDR Discussions

▪ Scientific

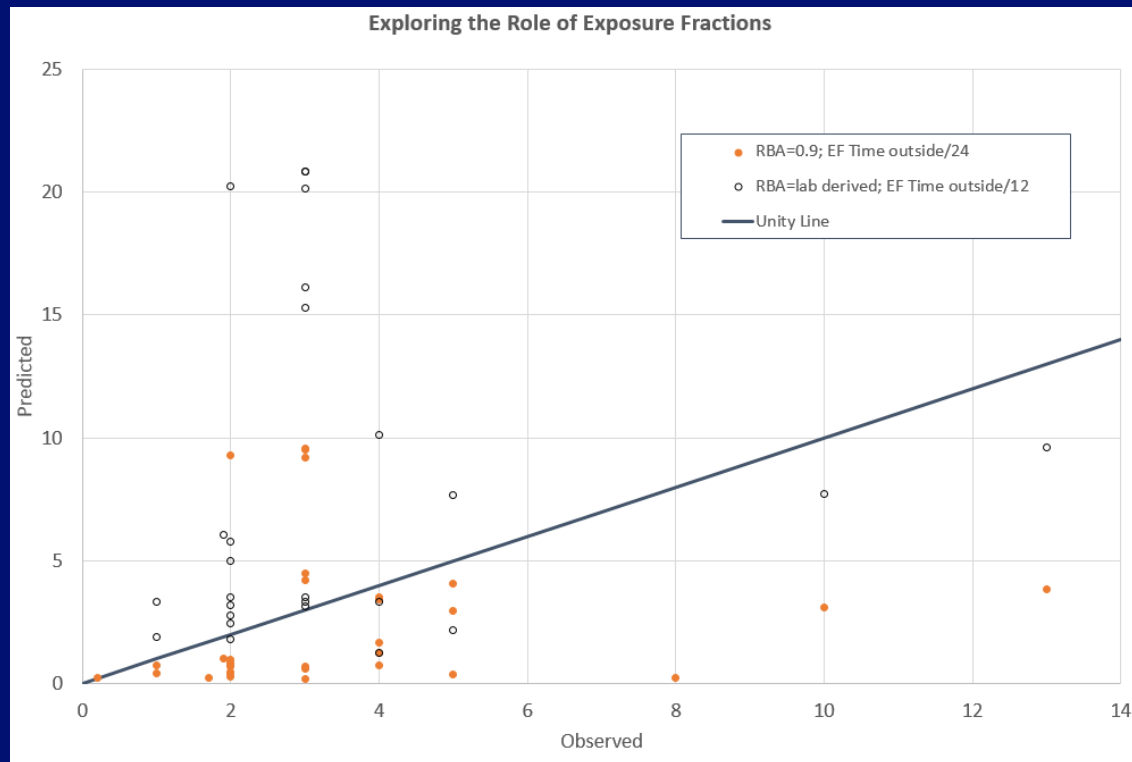
- Can the AALM PBPK model be used to inform short-term, acute, and *pica* lead exposures?
- How should exposure fractions be assigned in the AALM?
- How should ATSDR average lead exposures?
 - Yearly? 3 months? 6 months?
- How should ATSDR consider peak lead exposures?

▪ Logistic

- Vetting/approval of ATSDR's approach for lead evaluations

Current Activities AALM V4.2

- Currently testing the model using JT Lewis data.
- Refining exposure fractions



- May require data from other sites (new and/or existing)

Acknowledgements

- **Modeling team**
 - Rachel Worley
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- **Lead SMEs**
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- **JT Lewis site team**