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## ACCEPTED MANUSCRIPT

#### Toxicology Letter-TOXLET-D-18-00495

#### **Re-Defining Kinetic Lung Overload: Time for New Paradigms**

Running title: Re-Defining Kinetic Lung Overload

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#### Highglights

- Repeated exposure inhalation studies with poorly soluble particles should be structured by kinetic modeling
- The displacement volume of aggrgated particles is the key metric for doseresponse analyses
- The NOAEL and MTD should be expressed relative to the kinetic overload threshold
- The elimination kinetics of poorly soluble particles is linked to the total count of BAL-cells

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