

1.1. The Michaelis-Menten enzyme kinetics model (1913 *Biochem Z* 49: 333–369) is named after German biochemist Leonor Michaelis and Canadian physician Maud Menten. Yu and Rappaport (1997 *Environ Health Perspectives* 105 : 496–503) show that the Michaelis-Menten model describes the clearance rate (k) of insoluble dust particles from lungs as a function of the maximum rate (k_{max}), the particulate burden (m), and the particulate burden (m_{half}) at which k is half of k_{max} .

$$k = \frac{k_{max} \cdot m_{half}}{m + m_{half}}$$

$$k = k_{max} \left(\frac{m_{half}}{m + m_{half}} \right)$$

The parameter m_{half} and the variable m have units of milligrams (mg), k has units of %/day

a. Show units for the ratio in parentheses $\left(\frac{\text{mg}}{\text{mg} + \text{mg}} \right)$ and for k_{max} $\frac{\%}{\text{day}}$ [1+1]

b. Explain your answer for units of k_{max} [2]

Same as k because no units for $\left(\frac{m_{half}}{m + m_{half}} \right)$

c. Does the ratio in parentheses increase or decrease as lung burden m decreases?

Write your answer here it increases [no mark]

d. Given $m_{half} = 0.97$ mg for photocopier toner dust (PTT) and $k_{max} = 0.009/\text{day}$ for experimental rats, calculate the expected clearance rate at

$$\begin{aligned} m &= 5 \text{ mg} & E(k) &= \frac{0.001462}{1} \\ m &= 0.5 \text{ mg.} & E(k) &= \frac{0.005939}{1} \end{aligned}$$

d. Show your calculations, with units, for 0.5 mg

$$E(k) = (0.009) \left(\frac{0.97}{0.5 + 0.97} \right) = 0.005939$$

e. Does the expected clearance $E(k)$ change in the direction you expected, with decrease in lung burden m ? [no mark]

2. Using the expected value $E(k)$ at a burden of $m = 0.5$ mg, complete a data equation for an observed value of $k = 0.008$

$$\begin{aligned} k &= E(k) + \text{residual} \\ 0.008 &= 0.005939 + 0.002061 \end{aligned}$$

[3]

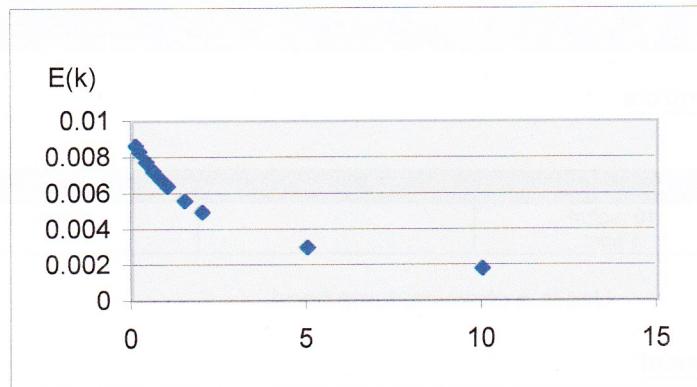
Quiz 3

DEP = Diesel Exhaust particles

kmax	0.009	per day
m	0.5	mg
mhalf	2.49	mg
E(k)	0.007495	per day
k	0.008	per day
residuals	0.000505	per day

A Lung Retention Model Based on Michaelis-Menten
 Rong Chun Yu and Stephen M. Rappaport
 Yu and Rappaport (1997 Environ Health Perspect)

m	E(k)
0.1	0.008653
0.2	0.008331
0.4	0.007754
0.6	0.007252
0.8	0.006812
1	0.006421
1.5	0.005617
2	0.004991
5	0.002992
10	0.001794



PTT = Photocopy Test Toner

kmax	0.009	per day
m	0.5	mg
mhalf	0.97	mg
E(k)	0.005939	per day
k	0.008	per day
residuals	0.002061	per day

m	E(k)
0.1	0.008159
0.2	0.007462
0.4	0.006372
0.6	0.005561
0.8	0.004932
1	0.004431
1.5	0.003534
2	0.002939
5	0.001462
10	0.000796

