

Energy intake, physical activity and body weight: a simulation model

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In adults, body mass (BM) and its components fat-free mass (FFM) and fat mass (FM) are normally regulated at a constant level. Changes in FM and FFM are dependent on energy intake (EI) and energy expenditure (EE). The body defends itself against an imbalance between EI and EE by adjusting, within limits, the one to the other. When, at a given EI or EE, energy balance cannot be reached, FM and FFM will change, eventually resulting in an energy balance at a new value. A model is described which simulates changes in FM and FFM using EI and physical activity (PA) as input variables. EI can be set at a chosen value or calculated from dietary intake with a database on the net energy of foods. PA can be set at a chosen multiple of basal metabolic rate (BMR) or calculated from the activity budget with a database on the energy cost of activities in multiples of BMR. BMR is calculated from FFM and FM and, if necessary, FFM is calculated from BM, height, sex and age, using empirical equations. The model uses existing knowledge on the adaptation of energy expenditure (EE) to an imbalance between EI and EE, and to resulting changes in FM and FFM. Mobilization and storage of energy as FM and FFM are functions of the relative size of the deficit (EI/EE) and of the body composition. The model was validated with three recent studies measuring EE at a fixed EI during an interval with energy restriction, overfeeding and exercise training respectively. Discrepancies between observed and simulated changes in energy stores were within the measurement precision of EI, EE and body composition. Thus the consequences of a change in dietary intake or a change in physical activity on body weight and body composition can be simulated.

Energy intake: Physical activity: Simulation model

Body mass (BM) in adult man is regulated at a constant level, an everyday experience backed by surprisingly little data in the literature. One of the few studies providing information on the constancy of BM is the Framingham Study, a long-term sampling of 5209 adults, 30–59 years of age, living in the town of Framingham at the start of the study from 1948 to 1949. They underwent, every 2 years, a standard medical examination, including the measurement of BM, for at least 20 years if not prevented by illness or death. Most subjects lost or gained no more than 5–10 kg over some part of the 20-year period as calculated by James (1985). This demonstrates a nearly perfect system for preserving energy balance as the total energy turnover of an average adult subject over 20 years is 73000 MJ. A discrepancy of 1% between energy intake (EI) and energy expenditure (EE) would add up to 730 MJ over 20 years, equivalent to about 24 kg BM as fat tissue with an energy density of 30 MJ/kg. Thus, in the long term EI matches EE within 1%.

There are situations where subjects are brought into a positive or negative energy balance by an intervention and where one wants to know the resulting consequences for BM. Examples are overeating or undereating and a restriction of physical activity or exercise

training. The consequences of the intervention for energy balance are not a linear function of the size of the intervention and are not linear over time. Reducing EI by 1 MJ/d or increasing EE by 1 MJ/d by eating less at the same level of physical activity or exercising more at the same level of EI respectively does not result in a BM loss of 1 kg over 30 d, using an energy equivalent of 30 MJ/kg as mentioned above. The BM loss is smaller as the body defends itself against a negative energy balance. A negative energy balance results in a reduction of basal metabolic rate (BMR) through a loss of BM and a reduction of EE per unit BM. The opposite holds as well: increasing EI by 1 MJ/d or reducing EE by 1 MJ/d by eating more at the same level of physical activity or exercising less at the same level of EI respectively does not result in a BM gain of 1 kg over 30 d. A positive energy balance results in an increase of BMR through an increase of BM. An increase of EE per unit BM, i.e. as a consequence of so-called futile cycles, has not been shown.

We will describe a model which uses existing knowledge on the relationship between EI, fat-free mass (FFM) and fat mass (FM), and EE in energy balance, and on the adaptation of EE to a changing EI, to a changing activity budget, and to resulting changes in FM and FFM. Thus the consequences of a change in dietary intake or a change in physical activity on FM and FFM can be simulated.

METHODS

The model is a continuous-time dynamic mathematical model. Input variables are energy intake (EI) and energy expenditure (EE). When EI does not match EE, energy is stored in or mobilized from the body assuming two compartments: fat mass (FM) and fat-free mass (FFM). The two input variables and the resulting consequences when EI does not match EE will be described separately.

Energy intake

EI, as an input variable, is estimated or can be calculated from information on dietary intake as collected with e.g. a dietary recall or a dietary record. The energy content of the diet is calculated with a nutrient table in a separate database (Stichting Nederlands Voedingsstoffenbestand, 1989). The resulting figure is the net energy intake, i.e. gross energy intake corrected for energy losses in faeces and urine, available for energy production.

Energy expenditure

EE is the sum of diet-induced energy expenditure (DEE) and physical activity (PA), i.e.

$$EE(t) = DEE(t) + PA(t), \quad (1)$$

where t is time in days. DEE is estimated at 10% EI (Schutz *et al.* 1984), i.e.

$$DEE(t) = 0.1 EI(t). \quad (2)$$

PA, as an input variable, is estimated or can be calculated from information on the activity budget as collected with e.g. an activity recall or an activity record. The activity budget is converted to PA with the energy cost of the activities expressed as a multiple of BMR, i.e.

$$PA(t) = \alpha(t) BMR(t), \quad (3)$$

where α is the physical activity factor. The value of the factor $\alpha (\geq 1)$, which is also an input of the system, for a range of physical activities is stored in a separate database (Ainsworth *et al.* 1993). For BMR the following empirical expression is used (see Appendix 1):

$$BMR(t) = 0.102FFM(t) + 0.024FM(t) + 0.85. \quad (4)$$

In a negative energy balance there is a lowering of BMR per unit BM depending on the size of the energy deficit. EE is lowered by an energy expenditure reduction (EER):

$$EER(t) = e(t)PA(t). \quad (5)$$

The value of e , which value is a function of EI/EE , ranges from 0.0 to 0.1 (James *et al.* 1990):

$$e(t) = \begin{cases} 0 & \text{for } (EI(t)/EE(t)) \geq 1 \\ 0.1 & \text{for } (EI(t)/EE(t)) \leq 0.5 \\ 0.2(1 - (EI(t)/EE(t))) & \text{for } 0.5 < (EI(t)/EE(t)) < 1. \end{cases} \quad (6)$$

Energy storage and mobilization

In a positive energy balance situation the energy surplus is stored and EE goes up because when an energy surplus is stored it has to be converted to a suitable compound, i.e. carbohydrate is deposited as glycogen or fat, fat is deposited as fat, and protein is deposited as protein, glycogen or fat. The conversion cost (EC) is estimated at a mean value of 10% of the energy surplus (Stock & Rothwell, 1982), close to the cost of depositing fat surplus as body fat, by far the most important process to store an energy surplus.

$$EC(t) = h(t)(EI - EE(t)), \quad (7)$$

where the conversion cost factor h is defined as:

$$h(t) = \begin{cases} 0 & \text{for } EI(t) \leq EE(t) \\ 0.1 & \text{for } EI(t) > EE(t). \end{cases} \quad (8)$$

Thus, the energy balance (EB) is:

$$EB(t) = EI - EE(t) + EER(t) - EC(t). \quad (9)$$

The change per day of FM and FFM depends on EB. The energy densities of FM and FFM are assumed to be 38 MJ/kg and 6 MJ/kg respectively, based on the energy density of fat and protein and FM being pure fat and FFM being 73% water and 27% protein. The ratio of mobilization or storage of energy between FM and FFM is not a constant (Forbes, 1987; Prentice *et al.* 1991). When EI exceeds EE excess energy is stored as FM and FFM in a ratio which is a linear function of the body composition. The surplus is stored as FM and FFM in a mass ratio of 75:25 or in an energy ratio of 95:5. When EI is lower than EE the energy deficit is covered from FM and FFM in a ratio which is a linear function of the relative size of the deficit ($EI:EE$) and of the body composition. In a situation where $EI:EE$ is between 1 and 0.5 and where FM is higher than 7.5% and 15% of BM in males and females respectively, the deficit is mobilized from FM and FFM in a mass ratio of 75:25 or in an energy ratio of 95:5. When $EI:EE$ is lower than 0.5 and/or FM is lower than 7.5% and 15% of BM in males and females respectively, relatively more energy is mobilized from FFM linearly increasing up to 100% FFM. There is a minimum FM of 3% BM for both males and females representing structural fat. Thus, the model is a system of two ordinary differential equations:

$$\begin{cases} \frac{dFM(t)}{dt} = \frac{fef(t)EB(t)}{fm_{ed}}, & (10a) \\ \frac{dFFM(t)}{dt} = \frac{(1 - fef(t))EB(t)}{ffm_{ed}}, & (10b) \end{cases}$$

where the change per day of FM and FFM depends on EB, the fat energy factor (fef), which

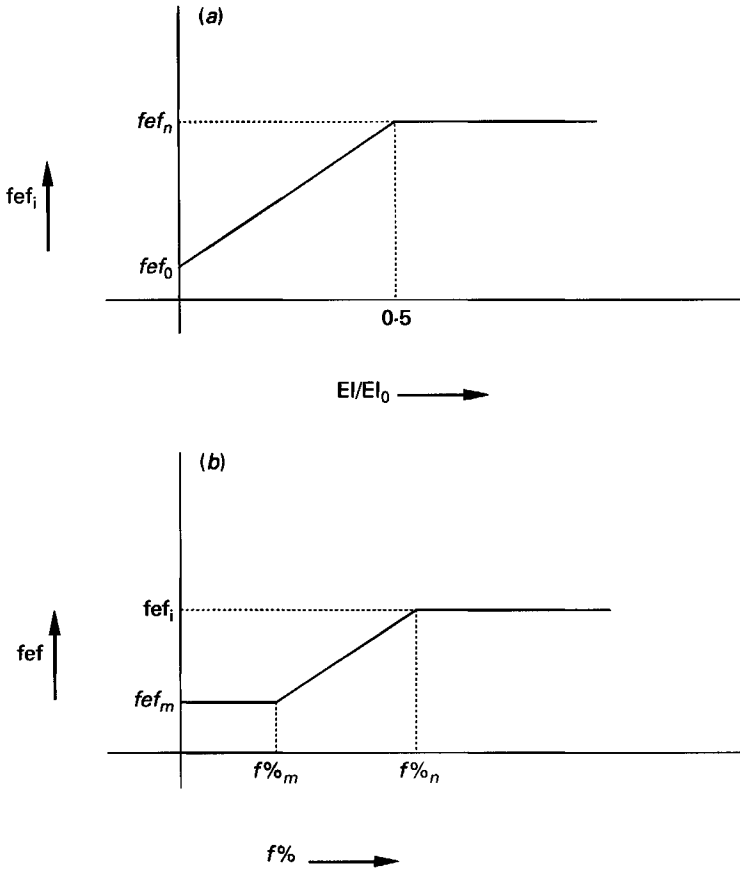


Fig. 1. Computation of (a) intermediate fat energy factor (fef_i) and (b) fat energy factor (fef). For definition of parameters and variables, see Table 1.

determines which part of the energy balance is transformed to FM, and fat mass energy density (fm_{ed}) and fat-free mass energy density (ffm_{ed}).

For the computation of the fat energy factor (fef) we need a measure for being out of equilibrium ($EB(t) = 0$). Let EI be the actual energy intake and $EI_0(t)$ the energy intake to achieve $EB(t) = 0$, which means that EI equals EE . From equations (1), (2) and (3) it follows that:

$$EI_0(t) = EE(t) = 0.1EI_0(t) + \alpha(t) BMR(t), \tag{11}$$

$$EI_0(t) = \alpha(t) BMR(t) / 0.9. \tag{12}$$

The fat energy factor (fef) is computed in a two-stage procedure (Fig. 1).

Stage 1: computation of an intermediate $fef(fef_i)$,

$$fef_i(t) = \begin{cases} fef_n & \text{for } (EI(t)/EI_0(t)) \geq 0.5 \\ 2(fef_n - fef_0) \cdot (EI(t)/EI_0(t)) + fef_0 & \text{for } 0 \leq (EI(t)/EI_0(t)) < 0.5, \end{cases} \tag{13}$$

where the factor fef_n is the normal fat energy factor and the factor fef_0 is the energy factor which is found for $EI = 0$.

Table 1. Summary of parameters and variables

Parameter	Meaning	Value	Unit
fef_m	minimal fat energy factor	0	—
fef_n	normal fat energy factor	0.95	—
fef_0	fef for energy intake = 0	0.9	—
ffm_{ed}	fat-free mass energy density	6	MJ/kg
fm_{ed}	fat mass energy density	38	MJ/kg
f^0_m	minimal fat percentage	3	—
f^0_n	normal fat percentage	men 7.5, women 15	—
Variable	Meaning	Definition (equation number)	Unit
α	physical activity factor	input variable	—
e	BMR-reduction factor	(6)	—
fef	fat energy factor	(14)	—
fef_i	intermediate fat energy factor	(13)	—
$f\%$	fat percentage	$100\% \cdot FM/(FM + FFM)$	—
h	conversion cost factor	(8)	—
t	time	—	d
BMR	basal metabolic rate	(4)	kJ
DEE	diet-induced energy expenditure	(2)	kJ
EB	energy balance	(9)	kJ
EC	conversion cost	(7)	kJ
EE	energy expenditure	(1)	kJ
EER	energy expenditure reduction	(5)	kJ
EI	energy intake	input variable	kJ
EI_0	'equilibrium' energy intake	(12)	kJ
FFM	fat-free mass	(10 <i>b</i>)	kg
FM	fat mass	(10 <i>a</i>)	kg
PA	physical activity	(4)	kJ

Stage 2: computation of fef ,

$$fef(t) = \begin{cases} fef_i(t) & \text{for } f\% > f^0_n \\ fef_m + (fef_i(t) - fef_m)(f\% - f^0_m)/(f^0_n - f^0_m) & \text{for } f^0_m \leq f\% \leq f^0_n \\ fef_m & \text{for } f\% \leq f^0_m \end{cases} \quad (14)$$

where $f\%$ is the fat percentage, f^0_n the normal fat percentage, f^0_m the minimal fat percentage and fef_m the minimal fat energy factor. A summary of all variables and parameters is given in Table 1.

RESULTS

A simulation usually starts from energy balance ($EI = EE$). When EI or EE is measured, EE and EI respectively can be matched to reach energy balance. When both EI and EE are measured there is usually a discrepancy and energy balance can only be reached by a manipulation of the data. A realistic starting point for a manipulation of the data is the knowledge that EE normally ranges between 1.2 BMR (Apfelbaum, 1978; Waterlow, 1986) and 2.2 BMR (Westerterp *et al.* 1992*a*), and has a value of 1.7 BMR for a subject with a moderate level of physical activity (World Health Organization, 1985 and Appendix 1). Subjects in the upper or lower parts of the range should know themselves as being, respectively, somebody with an above average or below average level of physical activity.

Theoretical predictions of the model can be compared with experimental evidence.

Table 2. *Body mass (BM) and fat mass (FM) in seven men before and after 21 d energy restriction (by 3.4 MJ/d) as reported by Heyman et al. (1992) compared with the results of the simulation model*

	After 21 d energy restriction					
	Baseline		Observed		Simulated	
	BM (kg)	FM (kg)	BM (kg)	FM (kg)	BM (kg)	FM (kg)
Mean (n 7)	69.2	9.1	67.2	8.0	67.5	7.8

Manipulations of energy balance inducing weight loss include studies with weight-reducing diets. On the other hand there are a number of studies describing weight gain after an experimental increase in EI. Studies describing the long-term effect of a manipulation of the level of physical activity on energy balance are scarce. Very few studies include observations on all three components of the energy balance equation: EI, EE, and FM and FFM. EI can be measured with sufficient accuracy with a fully weighed dietary record or with full provision of the diet. EE can be measured with sufficient accuracy in a respiration chamber or with doubly-labelled water. Acceptable methods for measuring changes in FM and FFM include hydrodensitometry and isotope dilution. Using these criteria there is currently one study on the effect of energy restriction, one study on overfeeding, and one study on the effect of an increase in physical activity.

Heyman *et al.* (1992) fed seven men at a level of 3.4 MJ/d below baseline requirements for 21 d. Baseline EI to maintain BM was determined over 10 d. The resulting mean EI of 15.3 MJ/d was very close to the measured EE with doubly-labelled water of 15.4 MJ/d over the 10 d interval. Subsequently EI was decreased by 3.4 MJ/d resulting in an EE of 14.7 MJ/d and 13.6 MJ/d over the first and the second 10 d intervals of energy restriction respectively. The level of PA expressed as EE/BMR showed a non-significant decrease from 2.1 (SE 0.1) to 2.0 (SE 0.2). Table 2 shows the relevant parameters before the intervention and at 21 d after the start of the intervention as measured and simulated with the model.

Diaz *et al.* (1992) overfed nine men by 50% above baseline requirements with a mixed diet for 42 d. Subjects were in energy balance in the baseline period of 3 weeks. Mean EI, calculated from the food supplied using food composition tables, was 13.3 MJ/d and mean EE, measured over the second and third weeks with doubly-labelled water, was 13.2 MJ/d. EE minus EI was 1 (SD 9)%. Subsequently EI was increased to 19.6 MJ/d, resulting in an EE of 15.0 MJ/d over weeks 6 and 7 of the 7-week overfeeding interval. The level of PA expressed as EE/BMR was 1.8 (SD 0.2) in the baseline period and remained unchanged. Table 3 shows the relevant parameters before the intervention and at 42 d after the start of the intervention as measured and simulated with the model.

Bingham *et al.* (1989) trained sedentary subjects, two women and three men, keeping EI at the level for maintenance of BM as measured in a 3–5-week control period before the start of the training. At the end of the 9-week training the subjects were capable of running for 1 h/d, 5 d/week. The level of PA expressed as EE/BMR, measured with a doubly-labelled water (EE) and in a respiration chamber (BMR) in the baseline period and over the last 2 weeks of the training, increased from 1.6 (SD 0.1) to 1.7 (SD 0.1) for the women and from 1.6 (SD 0.1) to 2.2 (SD 0.2) for the men. Table 4 shows the relevant parameters before

Table 3. *Body mass (BM) and fat mass (FM) in eight men before and after 42 d overfeeding (by 6.6 MJ/d) as reported by Diaz et al. (1992) compared with the results of the simulation model*

Subject no.	After 42 d overfeeding					
	Baseline		Observed		Simulated	
	BM (kg)	FM (kg)	BM (kg)	FM (kg)	BM (kg)	FM (kg)
1	75.4	13.1	85.9	20.7	85.0	20.3
2	71.3	11.7	76.3	12.9	77.2	16.1
3	68.5	14.2	76.5	18.7	77.9	21.3
4	57.9	6.5	66.7	13.6	67.7	13.9
5	89.8	26.9	97.3	32.1	96.5	31.9
6	84.7	25.6	92.6	30.6	89.9	29.5
7	70.2	11.8	76.1	15.3	76.5	16.5
8	68.2	14.7	76.1	18.0	76.3	20.8
Mean	73.4	16.1	80.4	20.2	80.9	21.2
SD	9.0	6.7	9.5	7.3	9.1	6.4

Table 4. *Body mass (BM) and fat mass (FM) in two women and three men before and after 63 d exercise training (increasing the average daily metabolic rate by 1 MJ/d in women and 4 MJ/d in men) as reported by Bingham et al. (1989) compared with the results of the simulation model*

Subject no.	Sex	After 63 d exercise training					
		Baseline		Observed		Simulated	
		BM (kg)	FM (kg)	BM (kg)	FM (kg)	BM (kg)	FM (kg)
1	f	51.2	15.4	52.3	15.6	53.4	16.8
2	f	66.5	20.0	66.7	18.0	62.6	17.1
3	m	64.4	10.6	63.2	10.0	64.7	10.7
4	m	68.4	7.3	68.0	4.9	59.7	2.4
5	m	63.8	8.2	60.4	3.7	60.0	5.9
Mean		62.9	12.3	62.1	10.4	60.1	10.6
SD		6.8	5.3	6.2	6.3	1.3	6.5

the intervention and at 63 d after the start of the intervention as measured and simulated with the model.

The model is applicable to predict the effect of a planned intervention to change EI and EE on FM and FFM. The other way around, it allows calculation of EI and EE to reach a planned value of FM and FFM at a given time interval. Finally, the model can provide information on the energy balance under normal circumstances. Two examples are given, one for the consequences of a planned intervention to change EI and EE on FM and FFM, the other one for the information this model can provide on energy balance under normal circumstances. In one study it was planned to reduce dietary fat intake by one third in

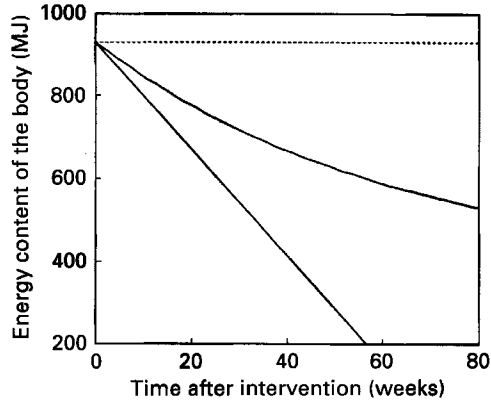


Fig. 2. Changes in the energy content of the body for a man weighing 75 kg, with 20% body fat at a daily energy intake equal to a daily energy expenditure of 13 MJ, after reduction of energy intake by 1.8 MJ/d, assuming no changes in physical activity. Upper line: result of the simulation model; lower line: result without adaptive changes in energy expenditure. For details, see pp. 343–344.

subjects consuming a Western diet with 40% energy as fat. For a man age 30 years, FFM 60.0 kg, FM 15.0 kg and $EI = EE = 13.0$ MJ/d, this implies a reduction of EI by 1.8 MJ/d. The results of the simulation model and a linear model, assuming no changes in PA, are after 26 weeks: FFM and FM of 58.4 and 10.2 kg, and 54.8 and 6.1 kg respectively. After 52 weeks the respective values are: FFM and FM of 57.3 and 7.1 kg, and 43.3 and 0.0 kg. A nearly twofold difference between the changes of FFM and FM and thus the energy content of the body calculated with the linear model and, the more realistic, simulation model (Fig. 2). Thus the simulation model can teach subjects about real EI and EE and about consequences of changes in EI and EE for FM and FFM. Starting a simulation with data on EI from a dietary record and data on EE from an activity budget usually results in a negative energy balance. Most subjects, especially overweight subjects underestimate EI (Schoeller, 1990; Westerterp *et al.* 1992b). In a group of students (twenty-four females and seven males) starting a simulation run with the model with individual information on recorded intake and recorded physical activity, EI was significantly lower than EE ($P < 0.001$; Wilcoxon signed rank). Additionally, the discrepancy between EI and EE was higher in subjects with a higher body mass index (Pearson r 0.56, $P < 0.001$).

DISCUSSION

The agreement of the simulation with observed values for BM and FM after a manipulation of EI or EE can be judged in units of weight or in units of energy. The latter allows combination of the two variables using the energy equivalents of the model, i.e. a change in FM has an energy equivalent of 38 MJ/kg and a change in BM minus FM is a change in FFM with an energy equivalent of 6 MJ/kg. Table 5 shows the discrepancy between predicted and observed changes related to EI and EE over the observation interval. Mean discrepancies for the three studies between simulated and observed results are within the measurement precision of EI, EE and body composition (see below).

Obviously there are two possible explanations for discrepancies between observed and simulated results. First, the input data on FFM, FM, EI and EE have a limited degree of accuracy. The accuracy of body composition data is difficult to assess as *in vivo* comparison with true values is impossible. The estimated precision of changes in FM and consequently

Table 5. Differences between measured and simulated energy content of the body (ΔE) in (A) the study of Heyman *et al.* (1992), (B) the study of Diaz *et al.* (1992) and (C) the study of Bingham *et al.* (1989)

Study	ΔE (MJ)		
	Mean	SD	Range
A	4	—	—
B	8	82	-67 to 128
C	-36	59	-109 to 52

in FFM is at best between 1 and 2 kg (Murgatroyd & Coward, 1989). All three studies included provisioning of a fixed diet over the entire observation interval. Thus, EI was calculated assuming that subjects consumed the food provided and did not consume additional food. EE was calculated as the mean of the value measured at the start and at the end of the observation interval on the assumption that any changes like the decrease through energy restriction or the increase through overfeeding or training were gradual and not instantaneous from the start of the intervention. Second, the model is based on assumptions and general equations including biological variation. It is difficult to estimate the effect of uncertainties in the separate components of the model. The resulting estimates are possibly the maximum one can obtain.

Unfortunately there are very few well-controlled studies on the consequences of a manipulation of EI or EE for FM and FFM in man. Measuring EI always interferes with a subject's behaviour under normal living conditions. Measuring EE is even more demanding when subjects are observed in a respiration chamber. The alternative, measuring EE with doubly-labelled water, is prohibitively expensive for long-term or large-scale application. Thus, three studies were available to validate the model with an intervention ranging from 3 to 9 weeks. The longer the study the lower the precision of the data on EE. Heyman *et al.* (1990) measured EE over a 10 d baseline and the full 21 d intervention interval. The other two studies measured EE over a baseline interval and the last 2 weeks of the 6- and 9-week intervention intervals and figures for the other weeks had to be calculated by interpolation.

The vast literature on the effect of energy restriction on FM and FFM is unfortunately not suitable to validate the model. Only the referred study of Heyman *et al.* (1992), using mild and short-term energy restriction, measured EE. Surprisingly up until now none of the slimming studies have included measurements on EE. If any, separate components of EE are measured. Keys *et al.* (1950), restricting EI in thirty-two men from 14.6 to 6.6 MJ/d for 6 months in the classical Minnesota experiment, did not measure EE either. Only since the introduction of the doubly-labelled water method to measure EE in man (Schoeller & Van Santen, 1982) is it feasible to measure EE with sufficient accuracy under normal living conditions. It is a matter of time before more studies on the effect of energy restriction on energy balance including EE measurements with doubly-labelled water appear.

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APPENDIX

Calculation of basal metabolic rate

Basal metabolic rate (BMR) is calculated from fat-free mass (FFM) and fat mass (FM), and FFM is calculated from sex, age, height and body mass (BM) with equations generated by combining studies on adult healthy subjects presenting individual data on sex, age, height, BM, FM, FFM, BMR and energy expenditure (EE) (see References). Table A1 summarizes the relevant data.

Data are combined in a multiple regression analysis resulting in the equations:

$$\text{BMR (MJ/d)} = 0.102\text{FFM (kg)} + 0.024\text{FM (kg)} + 0.85 \quad (n \ 190, r^2 \ 0.89),$$

for women:

$$\text{FFM (kg)} = -12.47 - 0.074 \text{ age (years)} + 27.392 \text{ height (m)} + 0.218 \text{ BM (kg)} \\ (n \ 105, r^2 \ 0.81),$$

for men:

$$\text{FFM (kg)} = -18.36 - 0.105 \text{ age (years)} + 34.009 \text{ height (m)} + 0.292 \text{ BM (kg)} \\ (n \ 85, r^2 \ 0.79).$$

Table A1. Characteristics of the subjects

	Women (n 105)			Men (n 85)		
	Mean	SD	Range	Mean	SD	Range
Age (years)	42	20	20-95	42	19	19-80
Height (m)	1.63	0.08	1.38-1.81	1.77	0.08	1.60-2.05
Body mass (kg)	62	16	38-130	80	24	55-215
Fat mass (kg)	19	12	3-76	19	17	2-128
Fat-free mass (kg)	42	6	29-60	61	10	44-93
BMR (MJ/d)	5.6	0.9	3.4-8.2	7.6	1.4	4.7-12.7
EE (MJ/d)	8.8	2.1	4.5-15.1	12.7	3.0	7.8-21.5
EE:BMR	1.57	0.25	1.07-2.25	1.70	0.30	1.13-2.60

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