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The Analysis of Multivariate Longitudinal Data: An Instructive Application of the Longitudinal Three-Mode Model

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ABSTRACT

Structural equation modeling is a common technique to assess change in longitudinal designs. However, these models can become of unmanageable size with many measurement occasions. One solution is the imposition of Kronecker product restrictions to model the multivariate longitudinal structure of the data. The resulting longitudinal three-mode models (L3MMs) are very parsimonious and have attractive interpretation. This paper provides an instructive description of L3MMs. The models are applied to health-related quality of life (HRQL) data obtained from 682 patients with painful bone metastasis, with eight measurements at 13 occasions; before and every week after treatment with radiotherapy. We explain (1) how the imposition of Kronecker product restrictions can be used to model the multivariate longitudinal structure of the data, (2) how to interpret the Kronecker product restrictions and the resulting L3MM parameters, and (3) how to test substantive hypotheses in L3MMs. In addition, we discuss the challenges for the evaluation of (differences in) fit of these complex and parsimonious models. The L3MM restrictions lead to parsimonious models and provide insight in the change patterns of relationships between variables in addition to the general patterns of change. The L3MM thus provides a convenient model for multivariate longitudinal data, as it not only facilitates the analysis of complex longitudinal data but also the substantive interpretation of the dynamics of change.

Introduction

Longitudinal studies in the life-sciences involve multiple observations at multiple measurement occasions, yielding multivariate longitudinal data sets. Structural equation modeling (SEM) offers a general and versatile framework for the analysis of such data. Compared with usual regression methods, SEM allows for the use of latent variables and measurement error of observed variables, and provides tests for overall goodness of fit, for specific hypotheses about relationships between variables and about longitudinal development. The longitudinal factor model (LFM; Oort, 2001; Tisak & Meredith, 1990) may include multiple latent variables, with multiple indicators from multiple measurement occasions, and thus enables investigation of complex longitudinal relations. However, the LFM becomes progressively large and unmanageable when

the number of measurement occasions increases. One of the methods that facilitates the investigation of longitudinal relations in more extensive data structures is the so-called longitudinal three-mode model (L3MM; Oort, 2001). In this paper, we provide an instructive description of the L3MM and illustrate how it can be used to test substantive hypotheses. It is our aim to facilitate applications of L3MMs for the investigation and interpretation of longitudinal dynamics, and thus help researchers and practitioners who are interested in developmental processes. In order to fully profit from the current tutorial, we recommend that the reader is familiar with the general SEM framework (cf. Bollen, 1989) and the LFM in particular.

The increased complexity of multivariate longitudinal data with larger numbers of measurement occasions can be best illustrated with an example. Imagine that we want to study the development of three

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KEYWORDS

Kronecker product; longitudinal factor model (LFM); longitudinal three-mode model (L3MM); multivariate longitudinal data



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constructs measured with three indicator variables each, which yields a data-structure consisting of nine indicators and three common factors. The LFM requires estimation of 78 parameters when it includes two measurement occasions, 228 parameters when it includes four measurement occasions, and 450 parameters when it includes six measurement occasions (see Appendix A for the calculation of the numbers of parameters). Estimation of model parameters may be difficult with such large models. Also, it has been argued that the trustworthiness of results decreases when the number of parameter estimates increases in relation to the sample size (Bentler & Chou, 1987; Jackson, 2003; Kline, 2011). Although the trustworthiness of results may also depend on many other model characteristics such as the number of variables per factor and the values of the factor loadings (Gagne & Hancock, 2006; Marsh, Hau, Balla, & Grayson, 1998), it seems plausible that convergence of estimation and stability of parameter estimates will be negatively affected with increasing model size because of increasing numbers of measurement occasions. More importantly, it becomes more difficult to arrive at a meaningful interpretation of findings when the number of model parameters is larger. For the interpretation of relations between the common factors from 2, 4, or 6 measurement occasions in the situation above, the LFM yields 15, 66, or 153 common factor covariances, respectively (see Appendix A). Such large numbers of parameter estimates complicate a meaningful interpretation of change in the relationships between common factors across time.

The increasing complexity of multivariate longitudinal models with multiple measurement occasions can be reduced by imposing additional restrictions on model parameters. Three-mode models are suited for the analysis of sets of data that are characterized by three modes. Multivariate longitudinal data are a kind of three-mode data, with the modes referring to the subjects, the variables, and the measurement occasions. Principal component and factor analysis techniques for three-mode data originate from Tucker's (1966) three-mode principal component analysis (e.g., the Tucker3 model), and include extensions of component analyses (e.g., the Candecomp/Parafac model; Carroll & Chang, 1970; Harshman, 1970), and of common factor analysis (Bentler & Lee, 1979; Bentler, Poon, & Lee, 1988; Bloxom, 1968). In the present paper, we focus on three-mode common factor analysis (see Kiers & van Mechelen, 2001; Smilde, Bro, & Geladi, 2004; Kroonenberg, 2008 for more general overviews of three-mode methods). The advantage of the common factor analysis framework is that it incorporates a versatile range of models and hypotheses to be tested. In addition, factor analysis techniques for multivariate longitudinal data are a special topic, as they offer unique opportunities for modeling the three-mode structure of the data (Oort, 1999) that not only greatly improve model parsimony but also facilitate interpretation of model parameters.

Specifically, the longitudinal three-mode model (L3MM; Oort, 2001) can be used for the analysis of multivariate longitudinal data, where the imposition of the so-called Kronecker product restrictions enables the decomposition of parameter matrices that describe the relationships between all variables from all measurement occasions into parameter matrices that describe the relationships between variables that apply to all measurement occasions, and parameter matrices that describe the relationships between measurement occasions that apply to all variables. Using this decomposition, the L3MM describes all relationships between all variables from all measurement occasions, but requires only the estimation of a much smaller number of parameters. In the example described above, imposition of Kronecker product restrictions on the relations between the common factors would require only 6, 13, or 24 estimates for the interpretation of 15, 66, or 153 correlations between common factors from 2, 4, or 6 measurement occasions, respectively. The L3MM thus substantially reduces the number of parameter estimates (i.e., leading to more parsimonious models), especially with larger numbers of measurement occasions. As a result, the L3MM parameters are easier to interpret. Instead of large matrices of parameters describing the relationships between all possible combinations of variables and measurement occasions, the L3MM yields two separate sets of much smaller matrices. One set of matrices has parameters describing the relationships between variables and another set of matrices has parameters describing the relationships between measurement occasions. In this way, relationships between variables can be modeled and interpreted separately from the relationships between measurement occasions. The underlying assumption of the imposed Kronecker product restriction is that longitudinal change in model parameters is proportional; the factor by which the relations between variables change over time applies to all variables equally. In the case of multiple indicators that measure the same underlying construct across time, this seems a plausible assumption. The imposes Kronecker product decomposition

multiplicative structure similar to longitudinal structures such as compound symmetry, autoregressive or latent curve models, which are often described as convenient models to simplify the correlation matrix of repeated measures (cf. Crowder & Hand, 1990; Lindsey, 1993). However, if the underlying assumption of the proposed Kronecker product restrictions does not hold, this may lead to biased estimates of effects. With the L3MM, the tenability of the Kronecker product restrictions can be tested, but the evaluation of (differences in) model fit is complicated due to the size of the models and increased parsimony of L3MMs. In the current tutorial, we will, therefore, also address the challenges of how to make appropriate decisions in evaluating the imposed L3MM restrictions, by using a combination of model fit statistics and substantive considerations.

The main benefit of increased model parsimony is that it enables the analysis of multivariate longitudinal data from many measurement occasions within the general SEM framework. Equally important, however, is that the substantive interpretation of changes in the relations between variables is facilitated as the L3MM yields separate estimates for the relationships between (observed and latent) variables and the relationships between the measurement occasions (i.e., the change in the relationships between the variables over time). The L3MM thus has the potential to improve our insight of longitudinal dynamics in the life sciences in general, and is especially suited to investigate and test the nature of these dynamics in multiple behavioral, cognitive, or psychophysiological measures (e.g., how the relationships between different neurological functionalities change with age, whether the variability in different mental abilities changes proportionally over time, or how the strength between several health outcomes is affected by therapeutic intervention). However, as of yet only few applications exist in the literature that take advantage of these unique characteristics of the L3MM. That is, applications are mostly limited to (technical) explanations (Kroonenberg & Oort, 2003; Oort, 2001) and are not yet used to address substantive research questions. There is thus a need to bridge the gap between the availability of L3MM model strategies for the analyses of longitudinal dynamics and their application.

The aim of the present paper is to provide an instructive description of L3MMs in order to stimulate their successful application. First, we will explain (1) how the imposition of Kronecker product restrictions can be used to take into account the multivariate

longitudinal structure of the data, (2) how to interpret the Kronecker product restrictions and the resulting L3MM parameters, and (3) how to test substantive hypotheses in L3MMs. Second, we will illustrate the application of L3MMs with an example of healthrelated quality of life (HRQL) data obtained from 682 patients with painful bone metastasis, with eight measurements at 13 occasions (104 variables); before and every week after treatment with radiotherapy. Part of these data have been analyzed before using simple repeated measures analyses to compare the development of HRQL between two different treatment regimens (Steenland et al., 1999), or using between group analyses to compare scores from only one specific measurement occasion (van der Linden et al., 2004). The latent variable model enables the analysis of changes in HRQL in much more detail, as it not only provides insight into changes in the means of variables, but also in changes in relations between variables over time. Using the example of bone metastases, we will illustrate how the L3MM can be successfully applied to provide a more comprehensive analysis of the multivariate longitudinal development of HRQL.

The L3MM

In order to facilitate the explanation of the L3MM, we will first describe the longitudinal factor model (LFM) and show how Kronecker product restrictions can be applied to yield the L3MM. Suppose R latent traits are measured with K observed variables on J occasions, the means and covariances of the observed variables are given by

$$E(\mathbf{x}) = \boldsymbol{\mu} = \boldsymbol{\tau} + \boldsymbol{\Lambda}\boldsymbol{\kappa},\tag{1}$$

and

$$Cov(\mathbf{x}, \mathbf{x}') = \mathbf{\Sigma} = \mathbf{\Lambda} \mathbf{\Phi} \mathbf{\Lambda}' + \mathbf{\Theta}, \qquad (2)$$

where τ is a *JK*-vector of intercepts, Λ is a *JK*×*JR* matrix of common factor loadings, κ is a *JR*-vector of common factor means, Φ is a *JR*×*JR* symmetric matrix containing the variances and covariances of the common factors, and Θ is a *JK*×*JK* symmetric matrix containing the variances and covariances of the residual factors. To achieve identification of all model parameters, scales and origins of the common factors can be established by fixing the intercept of one indicator per common factor (e.g., at zero), and fixing one common factor loading per common factor (e.g., at one).

The L3MM can be described by restrictions on the parameter matrices that feature in the mean and covariance structures of the LFM. We will explain

Λ _(JKxJR)	_	(ريدل)	0	$\Lambda_{0(KxR)}$
$ \begin{array}{c} \Lambda_{1(KxR)} \\ & \Lambda_{2} \\ & & \Lambda_{3} \\ & & & \dots \\ & & & & \Lambda_{J} \end{array} $	=		8	λ ₁₁ λ _{1R} λ ₂₁ λ _{2R} λ ₃₁ λ _{3R} λ _{K1} λ _{KR}
Intercepts ($\tau = \mathbf{u} \otimes \tau_0$), assuming	g $ au_j = au_0$ for all j			
τ _(JKx1)	$\mathfrak{g} \ \mathfrak{r}_{\mathfrak{j}} = \mathfrak{r}_{\mathfrak{0}} \ ext{for all } \mathfrak{j} = \mathfrak{r}_{\mathfrak{0}}$	u _(Jx1) 1	~	τ _{0(Kx1)}
		u _(Jx1) 1 1	8	$\left \begin{array}{c} \tau_{0(\mathbf{Kx1})} \\ \tau_{1} \\ \tau_{2} \end{array} \right $
τ _(JKx1) τ _{1(Kx1)}		u _(Jx1) 1 1 1 1	⊗	τ _{0(Kx1)}
$\tau_{(JKx1)}$ $\tau_{1(Kx1)}$ τ_{2}		u _(Jx1) 1 1 1 	8	$\left \begin{array}{c} \tau_{0(\mathbf{Kx1})} \\ \tau_{1} \\ \tau_{2} \end{array} \right $

Table 1. Imposition of measurement invariance restrictions on factor loadings and intercepts using the Kronecker product.

Notes: Λ_0 and τ_0 contain invariant factor loadings and intercepts of one measurement occasion that are applicable to all measurement occasions, I and u are an identity matrix and unity vector with dimensions equal to the number of measurement occasions, and the symbol \otimes denotes the Kronecker product.

these restrictions on each of the parameter matrices, and show how further restrictions can be imposed to test substantive hypotheses. Specifically, we will explain the imposition of Kronecker product restrictions on (1) factor loadings and intercepts (Λ and τ) to comply with longitudinal measurement invariance; (2) residual factor variances and covariances (Θ) , and additional restrictions to test equality of variances, correlations and covariances across occasions; (3) common factor variances and covariances (Φ), and additional restrictions to test equality of variances, correlations and covariances across occasions; and (4) common factor means (κ) , and additional restrictions to test a linear trend of common factor means. This sequence of imposing Kronecker product restrictions was chosen because the Λ and τ restrictions (1) are the most logical starting point from the researcher's perspective, as longitudinal measurement invariance is required for the comparison of common factors means, the Θ and Φ restrictions (2, 3) are most effective in increasing model parsimony and facilitating parameter interpretation, while κ imposed restrictions (4) can be used to test specific hypotheses regarding the common factor means while profiting from the model parsimony yielded by the earlier restrictions.

Longitudinal measurement invariance

With longitudinal data, the structure of matrix Λ is a block diagonal matrix containing matrices of factor loadings of each measurement occasion on the diagonal $(\Lambda_1, \Lambda_2, \ldots, \Lambda_j, \ldots, \Lambda_j$; see Table 1), where each of the Λ_j is a $K \times R$ matrix containing the factor loadings of occasion *j*. Vector τ consists of stacked vectors of

intercepts from all measurement occasions ($\tau_1, \tau_2, ..., \tau_j, ..., \tau_j$; see Table 1), where each of the τ_j is of length *K*. To test substantive hypotheses about the common factors, it is required that the meaning of these factors is the same across occasions. The requirement of longitudinal measurement invariance entails that the common factor loadings (Λ_j) and the intercepts (τ_j) are invariant across occasions (i.e., $\Lambda_j = \Lambda_0$, and $\tau_j = \tau_0$ for all *j*). The usual longitudinal measurement invariance entails and intercepts across time, can be written as a Kronecker product constraint:

$$\Lambda = I \otimes \Lambda_0, \tag{3}$$

$$\tau = \mathbf{u} \otimes \tau_0, \tag{4}$$

where Λ_0 is a $K \times R$ matrix of invariant common factor loadings, τ_0 is a *K*-vector of invariant intercepts, **I** is a $J \times J$ identity matrix, **u** is a *J*-vector of ones, and the symbol \otimes denotes the Kronecker product (see Table 1). The Kronecker product is an operation that can be applied to two matrices **A** and **B** of arbitrary size, and results in a block matrix that contains the matrices **B** pre-multiplied by each element of **A** (see Appendix B). The Kronecker product operations in Eqs. (3) and (4) impose the restriction that factor loadings Λ_0 and intercepts τ_0 apply to all measurement occasions.

Residual factor variances and covariances

Matrix Θ is a symmetric $JK \times JK$ matrix, consisting of $K \times K \Theta_{jj}$ matrices that contain the covariances of the residual factors on occasion *j* with the residual factors on occasion *j*'. Residual factors do not correlate with other residual factors, but are allowed to correlate with the same residual factors across occasions. Thus,

	Table 2. Imposition	of Kronecker	product	restrictions	on	residual	factor	variances	and	covariances.
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$ \begin{array}{c c} \text{Residual factor variances and c} \\ \Theta_{(JKxJK)} \\ \hline \Theta_{11}(KxK) \\ \Theta_{21} \\ \Theta_{22} \\ \dots \\ \Theta_{J1} \\ \Theta_{J2} \\ \dots \\ \Theta_{J1} \\ \Theta_{J2} \\ \dots \\ \Theta_{JJ} \\ \end{array} \right $	covariances ($\Theta = \Theta_T \otimes \Theta_V$)	$\begin{array}{c} {\bf \Theta}_{{\sf T}({\sf J} \times {\sf J})} \\ \theta_{{\sf T} 11} & \\ \theta_{{\sf T} 21} & \theta_{{\sf T} 22} \\ \cdots & \cdots & \cdots \\ \theta_{{\sf T} J1} & \theta_{{\sf T} J2} & \cdots & \theta_{{\sf T} JJ} \end{array}$	\otimes	Ο _{V(KxK)} θ _{V11} θ _{V22} θ _{VKK}
Residual factor correlations (Θ	$\mathbf{*} = \mathbf{\Theta}_{T}^{\mathbf{*}} \otimes \mathbf{\Theta}_{V}^{\mathbf{*}}$			
$ \begin{array}{c c} \Theta^{*}_{(JKSJK)} & & \\ I_{(KxK)} & & \\ \Theta^{*}_{21} & I & \\ \dots & \dots & \dots \\ \Theta^{*}_{J1} & \Theta^{*}_{J2} & \dots & I \end{array} $	=	$ \begin{array}{c} \Theta_{T}^{*}(\mathbf{j}_{X},\mathbf{j}) \\ 1 \\ \theta^{*}_{T21} & 1 \\ \dots & \dots & \dots \\ \theta^{*}_{TJ1} & \theta^{*}_{TJ2} & \dots & 1 \end{array} $	\otimes	Ον [*] (K×K) 1 1 1 1 1
Residual factor standard deviat	ions ($\Delta=\Delta_{T}\otimes\Delta_{V}$)			
$ \begin{array}{ c c c c c c c c c c c c c c c c c c c$	=	$ \begin{vmatrix} \Delta_{T(J,S,J)} \\ \delta_{T11} \\ \delta_{T22} \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\$	\otimes	$\begin{array}{c c} & \Delta_{V(KxK)} \\ \hline \delta_{V11} & & \\ & \delta_{V22} & \\ & & \cdots & \\ & & & \delta_{VJJ} \end{array}$

Notes: Residual factor covariances (Θ), correlations (Θ^*), and standard deviations (Δ) are decomposed using the Kronecker product (\otimes), where Θ_T , Θ_T^* , and Δ_T represent relationships between measurement occasions of residual factor covariances, correlations, and standard deviations, respectively; and Θ_V , Θ_V^* , and Δ_V represent residual factor variances, correlations, and standard deviations, respectively.

all $\Theta_{jj'}$ matrices are diagonal. Imposition of the Kronecker product restriction entails

$$\boldsymbol{\Theta} = \boldsymbol{\Theta}_{\mathrm{T}} \otimes \boldsymbol{\Theta}_{\mathrm{V}}, \tag{5}$$

where the full $JK \times JK$ matrix (Θ) is decomposed into two smaller matrices that describe the relations between the measurement occasions (Θ_{T} ; a symmetric matrix of dimensions $I \times I$) and the variances of the residual factors (Θ_V is a diagonal matrix of dimensions $K \times K$, containing within occasion correlations between residual factors) (see Table 2). The subscripts "T" and "V" refer to "time" and "variable." To achieve identification at least one parameter of $\boldsymbol{\Theta}_{\mathrm{T}}$ or $\boldsymbol{\Theta}_{\mathrm{V}}$ needs to be fixed at a non-zero value. Fixing the first element of $\boldsymbol{\Theta}_{T}$ to unity is a convenient choice for the interpretation of parameter estimates. Matrix Θ_{V} then contains the residual factor variances at the first measurement occasion, and Θ_{T} contains the relationships between residual factors across time that apply to all residual factors. We refer to these estimates as coefficients of proportionate change. If useful, matrix Θ_T can be further restricted to conform to, for example, compound symmetry, autoregressive, or latent curve structures (Oort, 2001). Imposing the Kronecker product restriction implies that the changes in variances and covariances of the residual factors across occasions are proportionate for all residual factors.

To further facilitate interpretation of parameter estimates it is convenient to use a reparameterization that decomposes the residual factor variances and covariances of Θ into correlations Θ^* and standard deviations Δ :

$$\Theta = \Delta \Theta^* \Delta, \tag{6}$$

where Δ is a $JK \times JK$ diagonal matrix containing the standard deviations of the residual factors, and diag(Θ^*) = I, so that the off-diagonal elements of Θ^* contain the correlations between the residual factors. This, in turn, enables the imposition of Kronecker product restrictions on residual factor correlations, using

$$\boldsymbol{\Theta}^* = \boldsymbol{\Theta}_{\mathrm{T}}^* \otimes \boldsymbol{\Theta}_{\mathrm{V}}^*, \tag{7}$$

where the full correlation matrix (Θ^*) is decomposed into two smaller matrices that describe the correlations between the measurement occasions (Θ_T^*) and the correlations between residual factors (Θ_V^*). As residual factors do not correlate with other residual factors, $\Theta_V^* = I$ (see Table 2). The reparameterization, therefore, allows investigation of the Kronecker product restrictions on residual factor correlations, while allowing each residual factor to have a unique standard deviation.

In addition, Kronecker product restrictions can be imposed on the standard deviations of the residual factors, using

$$\Delta = \Delta_{\rm T} \otimes \Delta_{\rm V}, \tag{8}$$

where $\Delta_{\rm T}$ is a $J \times J$ diagonal matrix that describes the proportionate change in standard deviations across occasions, and $\Delta_{\rm V}$ is a diagonal $K \times K$ matrix that contains the standard deviations of the residual factors at the first occasion (see Table 2). Imposition of Kronecker product restrictions on both Θ^* and Δ is equivalent to the imposition of the Kronecker product restriction directly on Θ (as in Eq. (5)).

Table 3. Imposition of Kronecker product restrict	ons on common factor variances and covariances.
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⊅ _(JR×JR)	1		$\Phi_{T(JxJ)}$		Φ _{V(RxR)}
		=	φ_{T11} φ_{T21} φ_{T22}	\otimes	φ_{V11} φ_{V21} φ_{V22}
Φ_{J1} Φ_{J2}	···· ··· Φ _{JJ}		$\begin{vmatrix} \cdots & \cdots & \cdots \\ \varphi_{TJ1} & \varphi_{TJ2} & \cdots & \varphi_{TJJ} \end{vmatrix}$		$\begin{array}{cccccccccccccccccccccccccccccccccccc$
Common factor correlations ($\mathbf{\Phi}^{*} = {\mathbf{\Phi}_{T}}^{*} \otimes {\mathbf{\Phi}_{V}}^{*}$		*		*
$\mathbf{\Phi}^{*}_{(\mathbf{JRxJR})}$	1		Φ _T *(LxL) 1		$\Phi_{V (RxR)}^{*}$
$\Phi^{*}{}_{21}$ $\Phi^{*}{}_{22}$		=	φ^*_{T21} 1	\otimes	φ^*_{V21} 1
$\mathbf{\Phi}^*_{J1}$ $\mathbf{\Phi}^*_{J2}$	 Φ* _{IJ}		$\begin{bmatrix} \cdots & \cdots & \cdots \\ \varphi^*_{TJ1} & \varphi^*_{TJ2} & \cdots & 1 \end{bmatrix}$		$\varphi^*_{VR1} = \varphi^*_{VR2} = \dots = 1$
ommon factor standard devia	tions ($\Gamma=\Gamma_{ extsf{T}}\otimes\Gamma_{ extsf{V}}$)				
$\begin{bmatrix} \Gamma_{1(\mathbf{R}\mathbf{x}\mathbf{R})} \\ \Gamma_{1}(\mathbf{R}\mathbf{x}\mathbf{R}) \\ \Gamma_{2} \end{bmatrix}$	Г,	=	Γ _{T(אל)}) אדוז דבע יייי	\otimes	Γ _{V(RxR)} ^γ V11 ^γ V22
			ענד א		γ _{trr}

Common factor variances and covariances ($\Phi=\Phi_{\mathsf{T}}\otimes\Phi_{\mathsf{V}}$)

Notes: Common factor covariances (Φ), correlations (Φ^*) and standard deviations (Γ) are decomposed using the Kronecker product (\otimes), where $\Phi_{T_r} \Phi_{T_r}^*$, and Γ_T represent relationships between measurement occasions of common factor covariances, correlations, and standard deviations, respectively; and $\Phi_{V_r} \Phi_{V_r}^*$ and Γ_V represent common factor variances, correlations, and standard deviations, respectively; and

Substantive hypotheses: Further restrictions enable hypothesis tests about the equality of residual factor correlations, variances, and covariances.

Equality of residual factor correlations of the same lag is investigated by imposing a banded structure on Θ_T^* in Eq. (7) so that all elements of the same diagonal are equal. This restriction implies that correlations between residual factors at the first occasion and residual factors at the second occasion are equal to correlations between residual factors at the second and third occasion, and so on.

Equality of residual factor standard deviations is investigated by imposing:

$$\Delta = \mathbf{I} \otimes \Delta_0, \tag{9}$$

where **I** is a $J \times J$ identity matrix and Δ_0 contains the invariant standard deviations of the residual factors of one measurement occasion that are applicable to all measurement occasions.

Equality of residual factor covariances across occasions of the same lag is tested by imposing both restrictions described above. This is equivalent to the imposition of the Kronecker product to Θ (as in Eq. (5)), where the banded structure is imposed on $\Theta_{\rm T}$.

Common factor variances and covariances

The procedure of imposing Kronecker product restrictions on the matrix of common factor variances and covariances is largely similar to the procedure for imposing Kronecker product restrictions on the matrix of residual factor variances and covariances described above.

Matrix Φ is a $JR \times JR$ symmetric matrix, consisting of $R \times R \Phi_{jj'}$ matrices that contain the covariances of the common factors at occasion *j* with the common factors at occasion *j'*. Imposition of the Kronecker product restriction implies that the change in relations between the common factors across occasions is proportionate for all common factors¹:

$$\mathbf{\Phi} = \mathbf{\Phi}_{\mathrm{T}} \otimes \mathbf{\Phi}_{\mathrm{V}},\tag{10}$$

where $\mathbf{\Phi}_{T}$ is a $J \times J$ symmetric matrix that describes the relationships between the measurement occasions, and $\mathbf{\Phi}_{V}$ is a $R \times R$ symmetric matrix that describes the relationships between the variables (see Table 3). For the purpose of identification, the first element of $\mathbf{\Phi}_{T}$ can be fixed at unity so that $\mathbf{\Phi}_{V}$ contains the common factor variances and covariances at the first measurement occasion, and $\mathbf{\Phi}_{T}$ contains coefficients of proportionate change. If useful, matrix $\mathbf{\Phi}_{T}$ can be further restricted to conform to, for example, compound symmetry, autoregressive, or latent curve structures (Oort, 2001).

In addition, it is convenient to use the following reparameterization:

$$\Phi = \Gamma \Phi^* \Gamma, \tag{11}$$

where Γ is a $JR \times JR$ diagonal matrix containing the standard deviations of the common factors, and

¹We note that the Kronecker product restriction of Eq. (10) is equivalent to a second order factor model $\Phi = \Lambda_2 \ \Phi_T \ \Lambda_2'$ in which Λ_2 contains the factor loadings of the (first-order) common factors on the second-order common factors, invariant across occasions ($\Lambda_2 = I_{(JxJ)} \otimes \lambda_{(Rx1)}$ so that $\Phi_V = \lambda_{(Rx1)} \lambda_{(Rx1)}$).

 Table 4. Imposition of Kronecker product restrictions on common factor means.

Common factor means ($\kappa = \kappa T \otimes \kappa V$)										
$ \begin{array}{c c} \kappa(KJx1) \\ & \mathbf{K}_{1}(Kx1) \\ & \mathbf{K}_{2} \\ & \cdots \\ & \mathbf{K}_{J} \end{array} $	=	$\kappa T(Jx1)$ $ \begin{matrix} \kappa_{T1} \\ \kappa_{T2} \\ \dots \\ \kappa_{TJ} \end{matrix}$	\otimes	кV(Кх1) _{Кү1} _{Кү2} _{КүJ}						
Linear tren	d of com	mon fact	or mea	ns ($\kappa =$	u \otimes a	$+ t \otimes $)			
к _(KJx1)	U	(Jx1)	ā	(Kx1)		t _(Jx1)	b (k	(x1)		
К _{1(Кх1)}		1		a ₁		0		b ₁		
κ ₂	=	1	\otimes	a ₂	+	1	\otimes	b ₂		
κ		1		a _K		J		b _K		

Notes: Common factor means ($\mathbf{\kappa}$) are decomposed using the Kronecker product (\otimes), where $\mathbf{\kappa}_T$ represents relationships between measurement occasions, $\mathbf{\kappa}_V$ represents common factor means of one measurement occasion, \mathbf{u} is a unity vector, \mathbf{a} is a vector of intercept parameters, \mathbf{t} is a vector with a time coding for the time of measurement, and \mathbf{b} is a vector of slope parameters.

diag(Φ^*) = I so that all off-diagonal elements of Φ^* are correlations between the common factors. This, in turn, allows for imposition of the Kronecker product restriction on the correlations between common factors (Φ^*) and the common factor standard deviations (Γ) separately

$$\mathbf{\Phi}^* = \mathbf{\Phi}_{\mathrm{T}}^* \otimes \mathbf{\Phi}_{\mathrm{V}}^*,$$

and

$$\Gamma = \Gamma_{\rm T} \otimes \Gamma_{\rm V}, \tag{13}$$

(12)

where Φ_T^* contains the correlations between measurement occasions, Φ_V^* contains the correlations between common factors irrespective of the measurement occasions, Γ_T contains coefficients of proportionate change in standard deviations across occasions (where the first element of Γ_T is fixed to unity for identification), and Γ_V contains the standard deviations of the common factors at the first measurement occasion (see Table 3). Imposition of Kronecker product restrictions on both Φ^* and Γ is equivalent to the imposition of the Kronecker product restriction directly to Φ .

Substantive hypotheses: Equality of common factor variances, correlations and covariances across occasions can be tested by further restricting the L3MM matrices.

The hypothesis of equal common factor correlations across occasions of the same lag is investigated by imposing a banded structure on Φ_T^* in Eq. (12) so that all elements of the same diagonal are equal. Because Φ_V^* is a symmetric matrix, this restriction entails that both the correlations between the common factors of one measurement occasion are equal across occasions, and that correlations between common factors at the first and second measurement occasions are equal to correlations between common factors at the second and third measurement occasions, and so on.

The hypothesis of equality of common factor variances across occasions is investigated by imposing

$$\Gamma = \mathbf{I} \otimes \Gamma_0, \tag{14}$$

where **I** is a $J \times J$ identity matrix and Γ_0 is an $R \times R$ matrix that contains the invariant standard deviations of the common factors of one measurement occasion that apply to all measurement occasions.

Equality of common factor covariances across occasions of the same lag is tested by imposing both restrictions described above, which is equivalent to imposing a banded structure directly on $\Phi_{\rm T}$ in Eq. (10).

Common factor means

The *JR*-vector $\mathbf{\kappa}$ consists of stacked $\mathbf{\kappa}_j$ vectors of length *R*, containing the common factor means of occasion *j*. The imposition of the Kronecker product restriction requires only estimation of $\mathbf{\kappa}_T$ and $\mathbf{\kappa}_V$:

$$\mathbf{\kappa} = \mathbf{\kappa}_{\mathrm{T}} \otimes \mathbf{\kappa}_{\mathrm{V}}, \tag{15}$$

where $\mathbf{\kappa}_{T}$ is the *J*-vector that contains coefficients of proportionate change in common factor means across occasions, and $\mathbf{\kappa}_{V}$ is a *R*-vector that contains the common factor means at the first measurement occasion (see Table 4).

Substantive hypotheses: To test and facilitate interpretation of (possible) changes in common factor means across time, we may impose various restrictions on κ . For example, to test for linear development of common factor means, we can impose

$$\boldsymbol{\kappa} = \boldsymbol{u} \otimes \boldsymbol{a} + \boldsymbol{t} \otimes \boldsymbol{b}, \tag{16}$$

where **u** is a unity *J*-vector, **a** is a *K*-vector of intercepts, **t** is a *J*-vector with some coding for the time of the occasions (for example 0, 1, 2, ... *J*), and **b** is a *K*-vector of slope parameters (see Table 4). The slope parameters then give an indication of the change across time for each common factor (instead of having to interpret all separate estimates of common factor means). To test invariance of common factor means we can fix the slopes at zero (i.e., $\mathbf{b} = \mathbf{0}$).

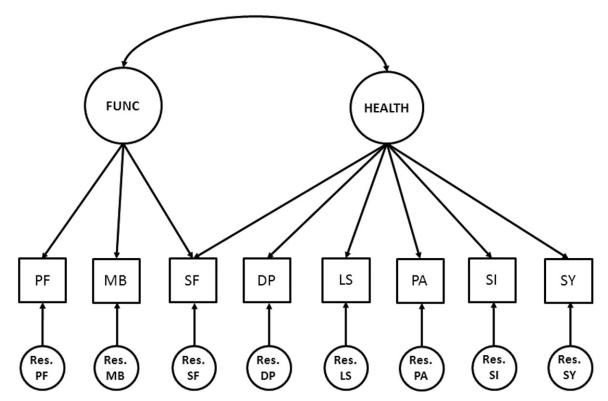


Figure 1. The measurement model. *Notes*: Circles represent latent variables (common and residual factors) and squares represent observed variables (the scale scores). FUNC: functional limitations; HEALTH: health impairments; PF: physical functioning; MB: mobility; SF: social functioning; DP: depression; LS: listlessness; PA: pain; SI: sickness; SY: treatment related symptoms; Res.: Residual factors.

Longitudinal structures of common factor

To further test substantive hypotheses regarding the common factors, one can apply longitudinal structures to both the covariance and mean structures of the common factors. Examples of these longitudinal structures are the autoregressive model and the latent curve model, which are common extensions within the SEM framework. For an explanation of these models, including possible variations and examples of hypotheses testing, the reader is referred to Oort (2001).

Illustrative example

Sample

The sample in the current study is a subset from the sample from the Dutch Bone Metastasis Study (DBMS; Steenland et al., 1999; van der Linden et al., 2004, 2006). In the DBMS, a total of 1157 patients (533 women) with painful bone metastases from a solid tumor were enrolled from 17 radiotherapy institutes in The Netherlands. The purpose of the study was to investigate the effectiveness of single fraction

versus multiple fraction radiation therapy for patients with painful bone metastases; the primary endpoint of the study was response to pain. The Medical Ethics Committees of all participating institutions approved the study and all patients gave their informed consent. For the present study, only patients who survived at least 13 weeks were included, which resulted in a total sample size of 682 patients (354 women). Patients' primary tumor was either breast cancer (n=321), prostate cancer (n=181), lung cancer (n=106), or other (n=74). Ages ranged from 33 to 90, with a mean of 64.2 (standard deviation 11.5).

Measures

Health-related quality of life questionnaires were administered before treatment (T0), and during the first 12 weeks of follow-up, patients completed weekly HRQL questionnaires by mail (T1 through T12). Questionnaire items were grouped into scales based on the results of principal component analysis of data from the first measurement occasion and substantive considerations (for more information see Verdam, Oort, van der Linden, & Sprangers, 2015). This resulted in the computation of eight health indicators: physical functioning (PF; four items), mobility (MB; five items), social functioning (SF; two items), depression (DP; eight items), listlessness (LS; six items), pain (PA; four items), sickness (SI; six items), and treatment-related symptoms (SY; 11 items). All scale scores were calculated as mean item scores, ranging from 1 to 4, with higher scores indicating more symptoms or more dysfunctioning. Analysis at the scale level is consistent with the usual evaluation of the original questionnaires that are still (partly) represented in the current scales, and was required to yield a manageable number of variables.

The eight health indicators were modeled to be reflective of two common factors: functional limitations and health impairments (see Figure 1). The squares represent observed variables (scale scores), the circles on the top represent the common factors, and the circles on the bottom represent residual factors. Functional limitations are measured by three observed variables, health impairments are measured by six observed variables, with one observed variable in common.

Statistical analyses

The program OpenMx (Boker et al., 2011) was used to run the statistical analyses. OpenMx is free and open source software for use within R that allows estimation of a wide variety of advanced multivariate statistical models. It was used because it allows for an operation of the structural equation model using matrix specifications, and therefore the Kronecker product restrictions can be easily applied. The variance-covariance matrix and mean vector that were used for statistical analyses, and syntaxes of all analyses that are reported in this paper, are provided as online Supplementary material.

Evaluation of Goodness of Fit: To evaluate goodness of fit the chi-square test of exact fit (CHISQ) was used, where a significant chi-square indicates a significant difference between model and data. As an alternative, the root mean square error of approximation (RMSEA; Steiger, 1990; Steiger & Lind, 1980) was used as a measure of approximate fit, where an RMSEA value below .05 indicates "close" approximate fit, and values below .08 indicate "reasonable" approximate fit (Browne & Cudeck, 1992). Additionally, expected cross-validation index the (ECVI; Browne & Cudeck, 1989) can be used to compare different models for the same data, where the model with the smallest ECVI indicates the model with the best fit. The ECVI is linearly related to the Akaike Information Criteria (AIC; Akaike, 1987) and

thus provides the same ranking of competing models (Browne & Cudeck, 1992). However, the ECVI has the advantage that confidence intervals are available for the differences between ECVI values of nested models. For both the RMSEA and ECVI 95% confidence intervals were calculated using the program NIESEM (Dudgeon, 2003). We also calculated the Comparative Fit Index (CFI; Bentler, 1990), where the model of interest is compared to a model of independence, i.e., a model where all covariances in Σ are assumed zero. The CFI ranges from zero to one, and as a general rule of thumb values above 0.95 are indicative of relatively "good" model fit (Hu & Bentler, 1999).

With different tests and indices to evaluate model fit, providing decision rules on whether the fit of a model is "good" is complicated by the fact that one might find inconsistent results (e.g., a significant exact chi-square test, but close approximate fit according to the RMSEA). The researcher then has to make a decision on which fit index is most appropriate for the data and hypotheses under study. For example, although the chi-square test of exact fit is the most commonly used, it is also generally acknowledged that it tends to become significant in larger samples and favors highly parameterized models. The described indices of approximate fit are less dependent on sample size and reward model parsimony, but they usually do not provide a test of model fit. In our example of bone metastasis the sample size is large and the model has many degrees of freedom. As a result, the chisquare test of exact fit has high power to detect small, but clinically meaningless, differences between model and data. Therefore, in this paper, we will base our evaluation of overall model fit on indices of approximate fit and will substantiate decisions on model fit evaluation in case of inconsistent results. A more extensive discussion about model fit evaluation follows in the discussion paragraph at the end of this paper (see also Verdam, 2017).

Evaluation of differences in Model Fit: To evaluate differences between hierarchically related models the chi-square difference test (CHISQ_{diff}) can be used, where a significant chi-square indicates a significant difference in model fit. The ECVI difference (ECVI_{diff}) can be used to test equivalence in approximate model fit, where a value that is significantly larger than zero indicates that the more restricted model has significantly worse approximate fit. In addition, it has been proposed that the difference between CFI values (CFI_{diff}) can be used to evaluate measurement invariance and more generally, the difference in

Table 5.	Goodness o	f overall fit	t of the	longitudinal	three-mode	models.

Model		Р	Df	CHISQ	CFI	RMSEA [95% CI]	ECVI [95% CI]
Measureme	nt model						
1.1	No restrictions	1274	4290	7002.34	0.975	0.031	14.71
						[0.029; 0.032]	[14.30; 15.13]
L3MM restri							
2.1	$\Lambda =$ I $^{\otimes}$ Λ_{0} ; $\tau =$ u $^{\otimes}$ τ_{0}	1118	4446	7661.17	0.971	0.033	15.13
	* ~					[0.031; 0.034]	[14.70; 15.59]
2.2	$_{a} \Theta = \Delta (\Theta_{T}^{*} \otimes I) \Delta$	572	4992	9471.75	0.959	0.036	15.90
						[0.035; 0.038]	[15.40; 16.41]
	$_{b} \Theta \!=\! \Theta_{T} ^{\otimes} \Theta_{V}$	488	5076	9829.93	0.956	0.037	16.13
						[0.036; 0.038]	[15.62; 16.66]
	_c Equal ε variances	476	5088	9846.17	0.956	0.037	16.11
		422	5140	10(00.4	0.040	[0.036; 0.038]	[15.60; 16.64]
	_d Equal ε correlations	422	5142	10690.4	0.949	0.040	17.16
	\mathbf{s} Equal ε covariances	410	E1E4	11/12 6	0.943	[0.039; 0.041]	[16.62; 17.72]
	e Equal & covariances	410	5154	11413.6	0.945	0.042 [0.041; 0.044]	18.18 [17.62; 18.77]
2.3	$_{a} \Phi = \Gamma(\Phi_{T}^{*} \otimes \Phi_{V}^{*})\Gamma$	242	5322	10486.0	0.953	0.038	16.24
2.5	$a \Phi = I (\Phi^{\uparrow} \Phi^{\downarrow}) I$	272	JJZZ	10400.0	0.755	[0.037; 0.039]	[15.71; 16.78]
	$_{b} \Phi \!=\! \Phi_{T} ^{\otimes} \Phi_{V}$	230	5334	10515.3	0.952	0.038	16.24
	P = = = A	250	5551	10515.5	0.752	[0.037; 0.039]	[15.71; 16.79]
	$_{\epsilon}$ Equal ξ variances	218	5346	10552.9	0.952	0.038	16.25
						[0.037; 0.039]	[15.73; 16.80]
	_d Equal ξ correlations	166	5398	10820.4	0.950	0.038	16.47
	u					[0.037; 0.040]	[15.93; 17.02]
	_e Equal ξ covariances	152	5412	10854.5	0.950	0.038	16.47
						[0.037; 0.040]	[15.93; 17.02]
2.4	$_{\rm a} \kappa = \kappa_{\rm T}^{\otimes} \kappa_{\rm V}$	218	5346	10579.3	0.952	0.038	16.29
						[0.037; 0.039]	[15.76; 16.84]
	_b Linear trend κ	208	5356	10611.4	0.952	0.038	16.30
						[0.037; 0.039]	[15.77; 16.85]
	_c Equal κ	206	5358	10637.2	0.952	0.038	16.34
Final Model						[0.037; 0.039]	[15.80; 16.89]
2.5	All tenable restrictions	184	5380	10664.7	0.951	0.038	16.30
			2.500			[0.037; 0.039]	[15.77; 16.85]

Notes: N = 682; P = number of free parameters in the model; Df = degrees of freedom; κ , Φ , and Θ are common factor means, common factor covariances, and residual factor covariances, respectively, and the subscripts "T" and "V" refer to matrices that contain relationships between different measurement occasions or variables respectively, Δ and Γ are standard deviations of residual factors and common factors, and Λ and τ are common factor loadings and intercepts.

model fit between two nested models (Cheung & Rensvold, 2002). As a rule of thumb CFI_{diff} values larger than 0.01 are taken to indicate that the more restricted model should be rejected. As confidence intervals are not available for CFI values, the CFI_{diff} cannot be used to test whether the difference in model fit is significant.

Evaluation of differences in model fit is complicated for similar reasons as described above. When comparing different models for the same data, one has to decide on the tradeoff between a deterioration of model fit and a gain in model parsimony. Such decisions should be guided by the evaluation of differences in model fit, but depend also on the substantive considerations with regard to interpretation of the model or model parameters. For example, the imposition of Kronecker product restrictions to take into account the multivariate longitudinal structure of the data generally leads to a large gain in model parsimony. When one considers the assumption of the multivariate structure of the data to be reasonable, one might not want to have too much power to detect small, but trivial, differences between model and data. However, when testing specific substantive hypotheses, one might consider high power to detect small differences to be beneficial. In this paper we will report results of all tests for differences in model fit that are explained above and will provide a rationale for the decision that is being made.

Results

The measurement model of Figure 1 was the basis for a structural equation model for baseline and follow-up measurements without any across occasion constraints. This model yielded a chi-square test of exact fit that was significant but the RMSEA measure and CFI indicated close fit (see Table 5, Model 1.1). The number of model parameters to be estimated was 1274.

L3MMs were applied to the 104 variables from 13 measurement occasions to investigate change in HRQL. Kronecker product restrictions were imposed on (1) factor loadings and intercepts (Λ and τ) to

comply with longitudinal measurement invariance; (2) residual factor variances and covariances (Θ), (3) common factor variances and covariances (Φ), and (4) common factor means (κ). Substantive hypotheses were tested at each consecutive step. We illustrate the application of each of the L3MMs that were described above, and explain why each model could be of substantive interest.

Longitudinal measurement invariance: Longitudinal measurement invariance is required in order to test substantive hypotheses about the common factors. The model with both factor loadings and intercepts restricted to be equal across occasions yielded a chi-square test of exact fit that was significant, but the RMSEA and CFI measures indicated close and good fit respectively (Model 2.1, see Table 5). To test whether the assumption of longitudinal measurement invariance holds, the model fit of this model can be compared to the model fit of the LFM without restrictions. Both the chi-square difference test and the ECVI difference test are significant (CHISQ_{diff} (156) = 658.8, p < .001; ECVI_{diff} = 0.43 95% CI: 0.29-0.59), indicating that the restrictions of invariant factor loadings and intercepts across occasions may not be tenable. However, the difference in CFI values indicates that the hypothesis of invariance should not be rejected (CFI_{diff} = 0.005). Moreover, overall model fit of the measurement invariance model is considered to be close (RMSEA =0.03). Inspection of parameter estimates showed no obvious deviational pattern between invariant factor loadings and intercepts (Model 2.1) as compared with the free factor loadings and intercepts (Model 1.1). As an example, the invariant factor loading of MB was estimated to be 0.71 (SE =0.02), where some of the freely estimated factor loadings were lower and others were higher, varying between 0.67 and 0.78 (a full overview of estimated factor loadings and intercepts of both models are provided in Appendix C). We therefore retain the model with measurement invariance restrictions, also in view of the overall model fit statistics. We consider the measurements "practically invariant." Nevertheless, it may be of interest to investigate possible violations of measurement invariance (i.e., measurement bias). However, because the invariant factor loadings and intercepts are a function of parameter estimates, the detection of measurement bias in Kronecker product restricted models requires alternative methods. A procedure for measurement bias detection in Kronecker product restricted models has been proposed elsewhere (Verdam & Oort, 2014). Here, we will retain the model with measurement invariance restrictions

on both factor loadings and intercepts for practical purposes. The invariance restrictions entail that the interpretation of both common factors is stable across time. The number of free parameters in the longitudinal measurement invariance model is 1118.

Residual factor variances and covariances: In our example of bone metastasis, it may be reasonable to assume that the residual variances, and the covariances of the same observed indicators at different occasions, change proportionately over time (e.g., patients may show more or less variability and co-variability over time, where this change is proportionally equal for all observed variables). Moreover, the complete matrix of residual factor variances and covariances has dimensions 104×104 and contains 728 free parameters, thus adding a large number of parameters to the model. Kronecker product restrictions on the residual factor variances and covariances are therefore most effective in increasing model parsimony and will therefore facilitate parameter interpretation. The imposition of the Kronecker product restriction on the residual factor correlations (Model 2.2a, see Table 5) and residual factor standard deviations (Model 2.2b) yielded close fit according to the RMSEA and CFI. Although the overall model fit is considered to be good, the deterioration in model fit as compared to the measurement invariance model (Model 2.1) significant (CHISQ_{diff} (630) = 2168.8, p < .001; is $ECVI_{diff} = 1.00$ 95% CI: 0.74–1.27; $CFI_{diff} = 0.015$). The number of degrees of freedom that is gained with these L3MM restrictions is considerable (630), with a total number of 488 parameter estimates. Therefore, in spite of the significant difference in fit, we will retain these L3MM restrictions because the overall model fit is good and the gain in model parsimony is substantial, and use this model as the reference model in subsequent model comparisons below. The imposed structure indicates that residual variances change proportionally over time, and that the longitudinal covariances apply to all residual factors.

Substantive hypotheses: In our example of bone metastases, it may be of interest to test whether the variances of the residual factors are invariant across time (i.e., showing equal reliability), and whether the relations between residual variables are stable across time. Models 2.2c, 2.2d, and 2.2e were used to test hypotheses about equality of residual factor variances, correlations and covariances, respectively. These restrictions have been imposed, one at a time (see Table 5). It appears that the residual factor variances are invariant across occasions (CHISQ_{diff} (12) = 16.2, p = .18; ECVI_{diff} = -0.02; CFI_{diff} < 0.001), but the

Table 6. Three-mode model parameter estimates for the final model (Model 2.5).

Longitudinal I Invariant facto			nce, where	$\Lambda = \mathbf{I} \otimes \mathbf{I}$	Λ_0 , and $\tau =$	= $\mathbf{u} \otimes \mathbf{\tau}_0$							
	i loaulitys	PF		MB	SF	DP		LS	PA		SI		SY
FUI	NC	1.00		0.76	0.32								
HEA					0.69	1.00		1.09	0.91		0.82		0.48
Invariant intere	cepts (τ_0)												
		PF		MB	SF	DP		LS	PA		SI		SY
		0.00		-0.19	-0.09	0.00		0.10	0.69		-0.03		0.51
Residual facto					$(\Theta_T^* \otimes I)$	Δ , and $\Delta =$	$I\otimes \Delta_{0}$						
Invariant stanc	dard deviat			$(diag(\Delta_0))$									
		PF	MB		SF		DP	LS		PA	SI		SY
		0.44	0.43		0.68		0.43	0.3	5	0.61	0.48	0	.25
Correlations be	etween me	asurement	occasions (Θ * _T)									
	T0	T1	T2	T3	T4	T5	T6	T7	T8	T9	T10	T11	T12
TO	1.00												
T1	0.64	1.00											
T2	0.57	0.71	1.00										
T3	0.52	0.65	0.74	1.00									
T4	0.49	0.61	0.67	0.76	1.00	1.00							
T5	0.48	0.61	0.65	0.71	0.78	1.00	1 00						
T6	0.48	0.60	0.63	0.68	0.73	0.80	1.00	1.00					
T7 T8	0.48 0.47	0.59	0.62 0.59	0.65	0.71 0.67	0.77 0.72	0.81 0.75	1.00 0.81	1 00				
T8 T9	0.47	0.56 0.55	0.59	0.62 0.61	0.67	0.72	0.75	0.81	1.00 0.81	1.00			
T10	0.40	0.53	0.56	0.60	0.64	0.68	0.73	0.78	0.81	0.81	1.00		
T10 T11	0.43	0.53	0.56	0.59	0.62	0.65	0.72	0.74	0.74	0.81	0.82	1.00	
T12	0.43	0.55	0.54	0.58	0.62	0.63	0.66	0.69	0.74	0.74	0.77	0.82	1.0
									0	•	•	0102	
Common factor Invariant stance						$P_V^*)I$, and I	$I = I \otimes I$	L ₀					
				5 (ulug(1 ()	/	FUNC			HEA	ITH			
						0.83			0.				
Correlations be	etween cor	nmon facto	ors (Φ_v^*)										
			•			FUNC			HEA	LTH			
		FUNC				1.00							
		HEAL	TH			0.44			1.0	00			
Correlations be	etween me	asurement	occasions (Φ _T *)									
	TO	T1	T2	T3	T4	T5	T6	T7	T8	T9	T10	T11	T12
Т0	1.00												
T1	0.89	1.00											
T2	0.86	0.93	1.00										
Т3	0.83	0.88	0.92	1.00									
T4	0.80	0.86	0.89	0.93	1.00								
T5	0.80	0.83	0.87	0.90	0.93	1.00							
T6	0.77	0.80	0.85	0.87	0.90	0.93	1.00						
T7	0.74	0.78	0.82	0.86	0.89	0.91	0.95	1.00					

0.73

0.70

0.68

0.67

0.64

Τ8

Т9

T10

T11

T12

2.58			

0.78

0.76

0.74

0.73

0.72

0.81

0.80

0.76

0.75

0.74

0.84

0.82

0.80

0.78

0.76

1.80

0.87

0.85

0.82

0.80

0.79

Slope parameters common factor means (b) FUNC HEALTH

0.04 -0.06

Notes: The variance-covariance structure of the final model, i.e., $\Sigma = \Lambda \Phi \Lambda' + \Theta$, is modeled by imposing the following Kronecker product restrictions: $\Sigma = (I \otimes \Lambda_0)((I \otimes \Gamma_0) (\Phi_T^* \otimes \Phi_V^*) (I \otimes \Gamma_0)) (I \otimes \Lambda_0)' + (I \otimes \Lambda_0)(\Theta_T^* \otimes I) (I \otimes \Lambda_0).$

0.90

0.89

0.86

0.84

0.81

0.92

0.90

0.88

0.86

0.83

0.94

0.91

0.90

0.87

0.85

1.00

0.95

0.93

0.92

0.89

1.00

0.95

0.92

0.90

1.00

0.95

0.92

1.00

0.95

1.00

The mean structure of the final model, i.e., $\mu = \tau + \Lambda \kappa$, is modeled using: $\mu = (u \otimes \tau_0) + (l \otimes \Delta_0)$ ($u \otimes a + t \otimes b$).

hypotheses about equal correlations and thus covariances across occasions must be rejected according to the chi-square difference and ECVI difference tests (CHISQ_{diff} (66) = 860.4, p < .001; ECVI_{diff} = 1.05 95% CI: 0.88–1.24). The CFI difference indicates that the

hypothesis of equal correlations might be tenable (CFI_{diff}=0.007), but that the hypothesis of equal covariances must be rejected (CFI_{diff}=0.014), although the overall model fit for both models is not considered to be good (CFI <0.95). Therefore, in our

example of bone metastases, only the hypothesis of equal residual variances seems tenable. This indicates that the unique variance of each indicator stays stable across time.

Common factor and variances covariances: Imposition of the Kronecker product restriction on the matrix of common factor variances and covariances yields 94 estimates to compute 351 parameters (i.e., the total number of common factor variances and covariances), thus having a large impact on model parsimony. Moreover, in our example of bone metastasis, it might be interesting to investigate whether the change in the variances and covariance of the underlying factors is proportionate over time. The model with Kronecker product restrictions imposed on both the common factor correlations and the common factor standard deviations yielded close fit according to the RMSEA and CFI (Model 2.3b, Table 5) and can be considered to show equivalent approximate fit compared to the model with free common factor variances (CHISQ_{diff} (258) = 685.3,p < .001; $ECVI_{diff} = 0.11 95\%$ CI: -0.03 to 0.26; $CFI_{diff} = 0.004$). Therefore, this model is retained and used as the reference model in subsequent model comparisons below. This result indicates that the standard deviations of the common factors functional limitations and health impairments and their correlation, change proportionately over time. The total number of free parameters in this model is 230.

Substantive hypotheses: It might be of interest to test whether the variances of functional limitations and health impairments are invariant across time, or whether the relationship between functional limitations and health impairments is invariant across time. Models 2.3c, 2.3d, and 2.3e are used to test equality of common factor variances, correlations, and covariances, respectively. The hypothesis of equal common factor variances across occasions should be rejected based on the chi-square difference test, but based on the ECVI difference test and the CFI difference the model with equal common factor variances can be retained (CHISQ_{diff} (12) = 37.6,.001; p < $ECVI_{diff} = 0.01$, CI: -0.010.06; 95% to $CFI_{diff} < 0.001$). Moreover, the overall model fit of this model is still considered to be good. The hypotheses about equal correlations and thus covariances across occasions must be rejected based on the chi-square difference and ECVI difference tests, but might be retained based on the CFI difference (CHISQ_{diff} $(64) = 305.2, p < .001; ECVI_{diff} = 0.23$ 95% CI: 0.13–0.34; CFI_{diff}=0.002). Taken together, these results indicate that only the hypothesis of equal

common factor variances is tenable. This entails that both the individual variability in the common factors and the covariance between the two common factors are stable across time.

Common factor means: In order to investigate the longitudinal development of the underlying factors functional limitations and health impairments, Kronecker product restrictions were imposed to test (possible) changes in common factor means across time. The model with Kronecker product restrictions on the common factor means yielded close fit according to the RMSEA, and good fit according to the CFI (Model 2.4a, see Table 5). The model that imposes a linear trend on the means of the common factors (Model 2.4b) can be considered to show equivalent approximate fit (CHISQ_{diff} (10) = 32.1, p < .001; $ECVI_{diff} = 0.01 95\%$ CI: -0.01 to 0.05; $CFI_{diff} < 0.001$), whereas the model that imposes equality of common factor means across occasions (Model 2.4c) shows a significant deterioration in model fit according to the chi-square difference and ECVI difference tests, but should not be rejected based on the CFI difference (CHISQ_{diff} (12) = 57.9, p < .001; ECVI_{diff} = 0.04 95% CI: 0.01–0.09; CFI_{diff} < 0.001). These results indicate that there is a significant change in the common factor means across time, and that this change can be described using a linear trend.

The final model: The model that includes all Kronecker product restrictions deemed tenable based on the results reported above would be a plausible final model. The final L3MM thus includes Kronecker product restrictions on the factor loadings and intercepts to comply with measurement invariance (Model 2.1), on the residual variances and covariances (Model 2.2b), on the common factor variances and covariances (Model 2.3b), and on the common factor means (Model 2.4a). In addition, the final model includes equality restrictions on residual factor variances (Model 2.2c) and common factor variances (Model 2.3c), and imposes a linear trend on the common factor means (Model 2.4b). The resulting final model yielded close fit according to the RMSEA, and good fit according to the CFI (Model 2.5, see Table 5). This L3MM required only 184 parameter estimates, and gained 1090 degrees of freedom as compared to the measurement model (Model 1.1).

Interpretation of parameter estimates: To illustrate the interpretation of L3MM parameters, we will examine the parameter estimates of the final model (Model 2.5) that are given in Table 6.

Factor loadings and intercepts: In the final L3MM, the factor loadings and intercepts are invariant over

time, i.e., $\Lambda = \mathbf{I} \otimes \Lambda_0$, and $\tau = \mathbf{u} \otimes \tau_0$ respectively (see Table 6). For example, the element λ_{052} is the estimated factor loading of the observed indicator listlessness, that applies to all measurement occasions (i.e., $\lambda_{052} = 1.09$). The element τ_{05} is the invariant intercept value that was estimated for the same indicator.

Residual factor variances and covariances: In the final L3MM, the Kronecker product restrictions imposed on Θ imply that residual factor variances are equal across time, and that the correlations between residual factors change proportionately, i.e., $\Theta = \Delta(\Theta_{T}^{*} \otimes I)\Delta$, and $\Delta = I \otimes \Delta_{0}$ (see Table 6). Thus, the first element of Δ_0 is the estimate of the invariant standard deviation of the residual factor of physical functioning ($\Delta_{011} = 0.44$), that is used for the computation of the residual variance of 0.20 (= 0.44^2) for all indicators of physical functioning across time. Estimates of Θ_{T}^{*} are the correlations between measurement occasions. The factor by which the relationships between the residual factors change across occasions is equal for all residual factors, but the actual covariances between the residual factors across occasions may differ because they are dependent on the standard deviations of the indicators. Also, it is now easy to see that correlations between measurement occasions decrease as the lag between the occasions becomes larger (i.e., the correlation between the first and the second measurement occasion is larger than the correlation between the first and third measurement occasion, and so on). In addition, correlations between measurement occasions of the same lag increase over time, i.e., the correlation between the first and the second measurement occasion is smaller than the correlation between the second and the third measurement occasion, and so on. This pattern of correlations explains why the restriction of equal correlations of the same lag (i.e., Model 2.2d) was not tenable. It might be, for example, that patients get used to the repeated assessments and therefore answer the questions in a more homogenous way.

Common factor variances and covariances: In the final L3MM, the imposed Kronecker product restrictions on Φ imply that common factor variances are equal across time, that the covariance between the common factors of the same measurement occasion is equal across time, and that the correlations between common factors of different measurement occasions change proportionately, i.e., $\Phi = \Gamma(\Phi_T^* \otimes \Phi_V^*)\Gamma$, and $\Gamma = I \otimes \Gamma_0$ (see Table 6). The estimates of the invariant standard deviations of the common factors functional limitations and health impairments (Γ_0) are

0.83 and 0.51, respectively, and the correlation between the two common factors at the same measurement occasion ($\Phi_{\rm V}^{*}$) is 0.44. The invariant covariance between the two common factors of one measurement occasion is thus 0.19 (i.e., 0.44 * 0.83 * 0.51). Correlations between measurement occasions $(\Phi_{\rm T}^{*})$ show the change in correlations between measurement occasions across time, e.g., correlation between measurement occasions decrease as the lag between measurement occasions becomes larger. Although correlations between measurement occasions apply to both common factors, actual covariances between common factors across occasions differ as they are dependent on the standard deviations of the common factors. Similar to the pattern of correlations between measurement occasions of residual factors, the result of the common factors shows a decrease in correlations between measurements as the lag between the occasions becomes larger, while correlations between measurement occasions of the same lag increase over time. This indicates that patients become more homogenous in their answers to the observed variables of physical limitations and health impairments over time.

Common factor means: In the final L3MM, the longitudinal development of common factor means is described by a linear trend, i.e., $\kappa = u \otimes a + t \otimes b$ (see Table 6). The intercept parameters $(\mathbf{a}_1 \text{ and } \mathbf{a}_2)$ are equal to the common factor means at the first measurement occasion. The slope parameters (\mathbf{b}_1 and \mathbf{b}_2) represent the linear change in common factor means across occasions, where the means of the common factor functional limitations increase across occasions ($\mathbf{b}_1 = 0.04$), while the means of the common factor health impairments decrease across occasions $(\mathbf{b}_2 = -0.06)$. However, only the decrease in health impairments was statistically significant based on the associated standard error of the slope parameter. The complete vector of common factor means is computed as a function of time of measurement. These results may indicate possible effects of patients' coping strategies, as the relatively objective indicators of functional limitations remain stable (e.g., physical functioning, mobility), whereas the more subjective indicators of health impairments (e.g., depression, pain) decrease over time.

Discussion

The L3MM is a valuable tool for the assessment of change in situations where there are many measurement occasions. Kronecker product restrictions yield very parsimonious models, enabling the application of SEM to large longitudinal data sets. In the present paper we explained and illustrated the imposition of Kronecker product restrictions on the parameter matrices of (1) factor loadings and intercepts to comply with the assumption of longitudinal measurement invariance; (2) residual factor covariances and correlations, and additional restrictions to test equality of variances, correlations and covariances across occasions; (3) common factor covariances and correlations, and additional restrictions to test equality of variances, correlations, and covariances across occasions; and (4) common factor means, and additional restrictions to test a linear trend of common factor means. In addition, we explained how the resulting parameter estimates can be interpreted. This paper, therefore, serves as an instructive description of L3MMs in order to facilitate their applications for complex longitudinal data and to enhance the substantive interpretation of model parameters. However, the illustration in the current tutorial also showed that model fit evaluation is not straightforward for these types of parsimonious models. Therefore, in the following, we address the challenges with regard to evaluation of (differences) in model fit in more detail, suggest areas for future research, and provide recommendations for researchers and practitioners who wish to apply the L3MM in an informative way.

The main benefit of the L3MM is that it enables the modeling of multivariate longitudinal data from many measurement occasions within the general and versatile SEM framework. When applying L3MMs one can thus profit from general SEM features and developments, such as methods for handling missing data (e.g., using alternative estimators), and diagnostics of possible misfit in (parts of) the model (e.g., using correlation residuals or modification indices) (cf. Bollen, 1989; Kline, 2011). The L3MM is especially suited for analyses of data from many measurement occasions (i.e., > 2) with fixed intervals between occasions. However, the L3MM can become of unmanageable size with very large numbers of occasions (e.g., with 50 occasions, the decomposition will yield a matrix of dimensions 50×50) and alternative modeling strategies are more appropriate (cf. Hamaker, Ceulemans, Grasman, & Tuerlinckx, 2015). Moreover, some general SEM guidelines may not be applicable to the specific L3MM context, such as sample size requirements or model fit evaluation, the latter of which we elaborate on below.

L3MMs are applied to assess change in multivariate longitudinal data with many measurement occasions.

The size of these types of models is usually large in terms of observed variables and model parameters. For example, in our sample of 682 patients with bone metastases we modeled 104 observed variables measured over 13 measurement occasions, which resulted in a measurement model that required estimation of 1274 model parameters with 4290 degrees of freedom. Evaluation of model fit is complicated by the fact that the chi-square test of exact fit is dependent on sample size and number of degrees of freedom (i.e., with increasing sample size and equal degrees of freedom the chi-square value increases) and tends to favor highly parameterized models (i.e., the chi-square value decreases when parameters are added to the model). The RMSEA and CFI indices of approximate fit are less dependent on sample size and reward model parsimony. In our illustration with the L3MMs the evaluations of overall model fit indicated that none of the models showed exact fit according to the chi-square test, while all models showed close approximate fit (RMSEA <0.05). The CFI index seemed to be somewhat more discriminative as not all models showed good fit (CFI >0.95), but without confidence intervals for these values the precision of the index is unknown. Therefore, this raises the question of how informative these overall model fit measures are in the case of highly parsimonious (longitudinal) models. Existing guidelines are based on simulation studies that have addressed the performance of overall goodness of fit measures, with relatively simple, singleoccasion examples (cf. Hu & Bentler, 1999). It may be the case that highly parsimonious models with large numbers of degrees of freedom require alternative fit indices or decision rules for an accurate evaluation of overall goodness of fit. For example, the RMSEA may not have enough discriminative power when models are very parsimonious (i.e., have many degrees of freedom). The CFI may not be informative with longitudinal data analyses, as the null model (i.e., a model without correlations between variables) is too unrealistic in the case of repeated measures of the same variables across time. As complex longitudinal models will only become more prevalent in the presence of large data sets, it would be worthwhile to investigate the behavior of overall model fit indices as a topic of future research.

The imposition of Kronecker product restrictions leads to more parsimonious models, and thus to deterioration in model fit. To test whether the restrictions are tenable we can test differences in model fit. As the imposition of Kronecker product restrictions usually leads to a large gain in number of degrees of freedom, evaluation of difference in model fit is complicated for the same reasons as described above. As an alternative to the chi-square difference test, we used the difference in ECVI and CFI values to evaluate differences in model fit. An advantage of the ECVI difference is that the associated confidence interval provides information about the precision of the estimate and allows to test the equivalence in approximate model fit. In our application, the chisquare difference test showed the highest power, rejecting all the L3MM restrictions (i.e., Kronecker product restrictions on parameter matrices), and all but one of the additional restrictions on L3MM parameter matrices to test substantive hypotheses. The ECVI difference test showed that some of the L3MM restrictions and substantive hypotheses could be retained based on the evaluation of equivalence in approximate fit, whereas the CFI difference showed the least discriminative power as almost all models could be retained based on the rule of thumb for this index (CFI_{diff} = 0.01). Thus, our illustration seems to indicate that the ECVI difference may be more informative for the evaluation of differences in model fit than the differences in chi-square or CFI values. However, stringent evaluation of the performance of the ECVI for the comparison of nested models has not yet been performed. Future research is needed to address the appropriateness of the ECVI difference in various contexts. Although the Kronecker product restrictions represent the data well and aid interpretation, it is difficult to provide decision rules for when the assumption of proportional change does not hold. Simulation studies are required to provide guidelines on how to address model fit evaluation in these circumstances.

Due to the problems in evaluating overall model fit and differences in model fit there is a risk of retaining an incorrect solution. Although Kronecker product restrictions lead to simpler - and thus more favorable - models according to the parsimony principle, one should be aware that very restrictive models may lead to biased parameter estimates. A rigid adherence to the parsimony principle could thus lead to bias and misinterpretation in model evaluation and selection. Unfortunately there is a lack of studies about when and to what extent parameters may be biased when the Kronecker product restrictions do not hold, thus complicating decisions on model fit evaluation. Therefore, we want to emphasize that statistics alone are not sufficient to guide decisions regarding these types of model evaluations, and that such decisions require substantive guidance as well. For example, the evaluation of difference in model fit can be used to test the tradeoff between model fit and model parsimony, but may also be affected by interpretability of results. In our illustration we incorporated Kronecker product restrictions on residual factor variances and covariances, even though these restrictions yielded deterioration in model fit. In part, we chose to incorporate these restrictions in favor of model parsimony and interpretability of results. Instead of yielding 104 estimates of residual factor variances and 624 estimates of residual factor covariances, the L3MM yielded eight estimates of residual factor variances, and 90 estimates that represent the proportional change in residual factor variances and covariances over time. These restrictions thus facilitate the substantive interpretation of model parameters - not only in terms of their number but also in terms of their meaningfulness due to the specifics of the L3MM decomposition. The argument to favor parsimonious models only when they facilitate interpretation has been made previously by others as well (e.g., Mulaik, 1998; McDonald & Marsh, 1990), and should help to avoid rigid applications of the parsimony principle (Raykov & Marcoulides, 1999). Therefore, the decision of whether to incorporate Kronecker product restrictions might not only be guided by evaluation of differences in model fit, but also by the improved substantive evaluation of findings. With regard to the imposition of additional restrictions on L3MM parameter matrices however, the decision of whether to incorporate these restrictions might not be guided by the same considerations of model parsimony and interpretability of findings. As these restrictions are imposed to test specific substantive hypotheses, it might be more desirable to have a high power to detect differences in parameter estimates. Therefore, substantive decisions play an important role in the evaluation of differences in model fit. As there is no single fit-index that can be appropriately applied for the evaluation of (differences) in model fit under these different circumstances, this might even require the use of different fit indices or different decision rules that are guided to the purpose of the analysis. This does not mean that one is free to apply the fit index or decision rule that is most convenient, but rather that fit indices and decisions rules should not be applied without critical evaluation of both the substantive and statistical considerations of the model and hypotheses under investigation. Future research may provide new fit indices or new guidelines that are better suited for the different purposes of L3MMs.

As a recommendation for researchers and practitioners that apply these types of models, we would suggest to (1) use several tests and indices of model fit in order to find support for the robustness of the result in their commonalities, (2) keep in mind that some fit indices are more appropriate in certain circumstances than others (e.g., specifically developed to take into account model parsimony), and (3) take into account substantive considerations when making decisions on model fit evaluation (e.g., using theory to establish an appropriate measurement model in addition to relying on model fit tests or indices to guide the specification of a measurement model) (see also Verdam, 2017).

To conclude, this paper provides an instructive application of the L3MM for multivariate longitudinal data from many measurement occasions. Kronecker product restrictions are used to model the multivariate longitudinal structure of the data, which yields models that are more parsimonious and have attractive interpretation. Application of the L3MM therefore facilitates the analysis of complex longitudinal data and can provide meaningful interpretation of the dynamics of change. However, future research is needed in order to support statistical decision rules for the tenability of Kronecker product restrictions and other substantive hypotheses in general. Such research will facilitate future applications of the L3MM and thus further our understanding of longitudinal dynamics within the life sciences.

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