Research Paper

Optimizing future well-being with artificial intelligence: selforganizing maps (SOMs) for the identification of islands of emotional stability

Fedor Galkin¹, Kirill Kochetov¹, Michelle Keller¹, Alex Zhavoronkov^{1,2,3}, Nancy Etcoff⁴

¹Deep Longevity Limited, Hong Kong
²Insilico Medicine, Hong Kong
³Buck Institute for Research on Aging, Novato, CA 94945, USA
⁴Department of Psychiatry, Massachusetts General Hospital, Harvard Medical School, Boston, MA 02115, USA

Correspondence to: Alex Zhavoronkov; email: alex@insilico.comKeywords: self-organizing maps, well-being, aging, artificial intelligence, depressionReceived: February 5, 2022Accepted: April 25, 2022Published: June 20, 2022

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ABSTRACT

In this article, we present a deep learning model of human psychology that can predict one's current age and future well-being. We used the model to demonstrate that one's baseline well-being is not the determining factor of future well-being, as posited by hedonic treadmill theory. Further, we have created a 2D map of human psychotypes and identified the regions that are most vulnerable to depression. This map may be used to provide personalized recommendations for maximizing one's future well-being.

INTRODUCTION

Psychological well-being

With the surge in mental illness, well-being has attracted the attention of many government agencies and research groups worldwide. In 2017 "Our world in Data" estimated that 792 million people live with a mental health disorder, with depression and anxiety disorders leading the list [1]. In 2018 that estimate had risen to one billion globally. The COVID-19 pandemic has only exacerbated the mental health situation, according to several recent reports [2–5]. Growing psychological distress, boredom, and feelings of isolation and loneliness have led to a decrease in well-being that is hampering society's productivity and ability to endure these challenging times.

Decades of well-being research have identified psychological well-being as tightly linked to physical health, and to optimism and supportive social networks. Well-being can be conceptualized as a positive mood and outlooks or as a sense of flourishing or positive functioning. High well-being is associated with positive health behaviors, lower risk of cardiovascular events, and all-cause mortality [6–9]. Among almost one hundred scales used to measure well-being, the Ryff scale is one the most widely used ones [10, 11]. The six dimensions of well-being defined within this scale are (i) autonomy, (ii) environmental mastery, (iii) personal growth, (iv) positive relations, (v) purpose in life, and (vi) selfacceptance. The original scale comprises 20 questions for each aspect of well-being, although a truncated version with only three questions each is also available [12]. The truncated version of the scale was used in the Midlife in the United States study (MIDUS), which we explored to study personal well-being trajectories [13, 14].

MIDUS contains almost four thousand subjects with known psychological profiles at two time points (MIDUS1 collected at 1995-1996 and MIDUS2 collected in 2004-2006), including their well-being profiles (Table 1). This unique dataset allowed us to explore the age-dependent component of well-being and to address the questions of its stability. The U-shape hypothesis posits that people experience minimal well-being in

MIDUS1 variable	MIDUS2 variable	Description
A1SPWBR	B1SPWBR1	Positive relations
A1SPWBS	B1SPWBS1	Self-acceptance
A1SPWBA	B1SPWBA1	Autonomy
A1SPWBG	B1SPWBG1	Personal growth
A1SPWBE	B1SPWBE1	Environmental mastery
A1SPWBU	B1SPWBU1	Purpose in life
A1PAGE_M2	B1PAGE_M2	Age
A1PDEPAD	B1PDEPAD	Depressed affect
A1PDEPDX	B1PDEPDX	Depressed affect + anhedonia

Table 1. The variables of interest studied in this article.

middle age, a claim that is still being argued actively [15, 16]. The results of studies addressing this phenomenon are highly dependent on the data source and methodology. Some studies have reported that wellbeing is mostly static within a smaller timeframe [17]. In some cases, no age effects are detected on a lifetime scale [18]. The concept of static well-being is frequently referred to as the "hedonic treadmill," the idea that even after major events, people return to their baseline level of life satisfaction. This concept, however, is an inaccurate representation of reality, since well-being adaptation has been shown to be avoidable [19, 20]. While research has suggested that happiness levels are durable, withstanding sweeping changes in health and wealth, the extreme picture of the hedonic set point as inevitable for all is in need of revision. While the average person's happiness may return to baseline, the happiness of many individuals does not [21]. In part, the fluidity of overall life satisfaction can be explained by the multifaceted nature of well-being, and each aspect has a different response rate to life events. We show that there are quite distinct non-constant trajectories for the six well-being parameters.

mHealth apps and self-help

Data-driven approaches to mental health are a rapidly developing field of psychology. The widespread use of smartphones has enabled novel ways to monitor behaviors and interact with people in need of guidance. According to recent market research, the mHealth app industry will continue expanding at a 28% annual growth rate, indicating great demand [22]. Compared to face-to-face psychology sessions, the mHealth approach allows for more frequent interactions at a greatly reduced cost. Reviews of this technology suggest that internet-based cognitive behavioral therapy (CBT) reduces the frequency and severity of depressive symptoms [23].

The models reported in this article may be used to develop online CBT tools for self-improvement, initial screening, or therapist-guided interventions.

The online mental health approach, however has serious shortcomings. First of all, mHealth CBTs tend to have double the dropout rate of clinician-based alternatives at 74%, thus demonstrating the necessity of therapist support [24]. In part, high dropout may be caused by insufficient personalization of self-help methods and a lack of professional oversight to achieve the desired change. This may not be an issue in academic studies with smaller samples and well-defined purposes, such as treating insomnia or anxiety [25–27]. However, the variability of psychological problems the general population encounters necessitates an online tool that can adapt to its users and their unique psychological profiles.

Deep learning approaches, such as neural networks and self-organizing maps (SOMs), can carry out initial assessment automatically and serve as recommendation engines to provide personalized daily tips. We expect that user interactions guided by the models described in this article can create better user satisfaction and lead to better retention.

The AI approach and aging clocks

Over the last decade, deep learning techniques have spread to the point of ubiquity. This blanket term encompasses a wide variety of data-driven algorithmic models used in synthetic data generation, image recognition, signal denoising, decision making, chatbots, recommendation engines, but most importantly to the scope of this article — in clinical settings.

Deep learning algorithms excel in tasks that require discerning non-linear dependencies, provided a sufficiently large training set is provided and the task is well formalized. In application to life sciences, deep learning has been demonstrated to successfully solve the tasks of pharmaceutical lead selection and drug design, clinical trial design, as well as diagnostic imaging and electrodiagnosis [28–32]. AI systems have a virtually unlimited bandwidth, compared to human professionals, and can deliver their reports in a matter of seconds, thus enabling more affordable and scalable healthcare [33].

The rapid development of deep learning has eventually reached the field of biogerontology in which neural networks were used to create digital models of aging aging clocks. Such models allow measuring the intensity of the aging-related processes in the human body and were introduced in a seminal paper by Steve Horvath in 2013 [34]. The original publication featured a linear model that predicted human chronological age based on their DNA methylation profile. Since then, aging clocks have been trained on practically any type of bio-relevant information, such as facial photos, transcriptomes, clinical blood tests and more [35-43]. The algorithms behind aging clocks have also become more sophisticated and now modern instances feature deep learning solutions as well as models trained to predict frailty or mortality risk directly [44-48].

Although the use of aging clocks is currently limited to research settings, this technology carries the promise of serving as the foundation for a disruptive model of global healthcare, longevity medicine [49–57]. The pace of aging detected with aging clocks has been associated with higher all-cause mortality risk, frailty, and the development of aging-related diseases [58–60].

Many factors of everyday life have been studied in the context of their effect on the pace of aging, including psychological factors such as stress. Childhood stress, traumatic experiences, intense competition, or experiencing violence have been shown to increase the biological age identified with epigenetic aging clocks [61–64]. The supposed mechanism linking stress and the pace of aging involves aberrant glucocorticoid signaling disrupting the DNA methylation patterns in key aging loci [65, 66]. Other studies also outline subjective age as major factor in overall well-being and all-cause mortality [67].

In this article, we continue to explore human psychology and its lifetime evolution. The longitudinal setting of the MIDUS dataset enabled us to create a deep learning predictor of future well-being and psychological age. Furthermore, we used SOMs to visualize human psychotypes and derive the paths of incremental lifestyle changes that should lead to an increase in current and future well-being (Figure 1). The deep learning approach allowed us to capture the complex, non-linear properties of the human psyche. The general approach described here can easily be implemented as the core of an mHealth application to help people improve their mental resilience.

RESULTS

Deep learning predictors of age and future well-being

We employed Boruta feature selection to identify all MIDUS features that may be associated with future wellbeing to obtain a list of 32 variables (Supplementary Figure 1A and Table 2), which were used to train an age



Figure 1. We have created the backbone of an Al-assisted recommendation engine to improve current and future psychological well-being based on self-organizing maps (SOMs). A person seeking self-improvement fills in a psychological test and is placed on a 2D representation of the multidimensional space containing all possible psychotypes. The map consists of regions associated with high (green) and low (red) well-being, which may be considered "mountains" and "pits". Distance metrics defined within a SOM allow finding the shortest path between a person's starting point and the point that maximizes their well-being. One's journey across the SOM may be interpreted as a chain of incremental changes leading to higher well-being. The SOM offers non-trivial, personalized paths towards improved well-being that can be followed and tracked within a self-help app or during therapy sessions.

Selected features	Description			
A1PD1	Satisfied with life at present			
A1SF1C	Some wander aimlessly, but not me			
A1SF1D	Demands of everyday life often get me down			
A1SF1F	Maintaining close relationships difficult			
A1SF1I	Good managing daily responsibilities			
A1SF1K	Life process of learning/changing/growth			
A1SF1L	Experience challenge how think important			
A1SF1M	Others describe me as giving/share time			
A1SF1U	Do just about anything I set my mind to			
A1SF1X	When really want something, find way			
A1SF1Y	Many things interfere with what I want do			
A1SF1Z	Whether I get what want is in own hands			
A1SF3B	Do what can to change for better			
A1SF3P	Know what I want out of life			
A1SF3Q	I live one day at a time			
A1SF3T	Helpful to set goals for near future			
A1SF3W	No use in thinking about past because nothing can be done			
A1SF4A	Outgoing describes you how well			
A1SF4D	Organized describes you how well			
A1SF4Y	Broad minded describes you how well			
A1SF4Z	Sympathetic describes you how well			
A1SK10A	Give spouse/partner emotional support (hours/month)			
A1SK17A	World is too complex for me			
A1SK17F	Feel close to others in community			
A1SK17G	Daily activities not worthwhile for community			
A1SK17J	People do not care about others problems			
A1SK17M	Society not improving for people like me			
A1SK17N	Believe people are kind			
A1SK7I	Serve on a jury if called			
A1SK7Q	Volunteer for social causes			
A1SM13	Rely on friends for help with problem			
A1SM5	Open up to family about worries			

Table 2. Features used to train the SOM and the

predictor and a well-being predictor. Since the selected features are not, by design, associated with age, the final model only marginally outperformed the median age assignment, displaying mean absolute error (MAE) of 9.06 years in the test set (Supplementary Table 1). The predicted psychological age may be further corrected for chronological age to make it representative of the typical aging rate in each particular age group. This procedure boosted the accuracy of the model to an MAE of 4.30 years, nevertheless completely transforming the meaning of its output. Thus, the prediction is no longer a chronological age estimate but a metric of psychological similarity to age peers.

Future well-being predictions were not adjusted, and the accuracy of predictions ranged between mean absolute percentage error (MAPE) of 15.81% and 22.53%

for different aspects of well-being in 10 years (Supplementary Table 1). The most accurate predictions were obtained for personal growth, while estimates of future self-acceptance were the least accurate.

Relative effect of deep learning features on well-being

To illustrate the reliability of the generated future wellbeing estimates, we performed a relative importance analysis using an elastic net (EN) method. Actual wellbeing features in the follow-up MIDUS2 wave were regressed as a linear combination of sex, age, the corresponding predicted well-being variable, and all six MIDUS1 well-being features (Supplementary Table 2). In all such models, model-derived estimates showed the largest coefficients, thus identifying them as more important determinants of future well-being than current well-being.

SOM-based recommendations for improving wellbeing

The 32 selected variables form a space of psychotypes that is representative of the general US population. To visualize and navigate these psychotypes, we used SOMs—a method of dimensionality reduction that may be interpreted as a 2D projection that preserves the distances between samples.

The SOM we obtained contains 625 cells organized in a 25×25 grid. Each psychological profile matches only one cell, while each cell may have multiple matching profiles. By design, SOMs ensure that subjects with similar psychotypes occupy the same cell. Conversely, more dissimilar psychotypes will be farther apart on an SOM.

SOMs provide a way to explore metadata and assess whether people with similar psychotypes (the same SOM region) share any properties not included in the initial 32 variables. Within the scope of this article, we have focused on the well-being structure of SOMs. Originally, we intended to describe all six well-being parameters with SOMs. However, for the sake of brevity, we have used a proxy of well-being in the main text, and the SOM distributions for all six parameters are available in the Supplementary Materials.

Our results showed that the odds of depression for people mapped to the same SOM cell were significantly negatively correlated with all current and four out of six future well-being parameters (Supplementary Table 3). Thus, we established depression odds as a reliable proxy for well-being.

Although the SOM was trained on a cohort of nondepressed people, it contained three distinct clusters with significantly different affinity for depressed people from the test set (Figure 2, Table 3 and Supplementary Figure 1B). Respondents mapped to the smallest Cluster-1 had 2.15 odds of being depressed, while in the two other clusters, non-depressed people outnumbered the depressed. Similarly, future well-being in all six



Figure 2. The SOM trained on a cohort of non-depressed people separates the depressed and the non-depressed. (A) Hierarchical clustering of SOM's nodes identified three clusters; (B) Dendrogram of the clusters displayed in section A. Distance is Euclidean distance between clusters (complete linkage); numbers in brackets mark the number of leaves below the pruned branches; (C) Cluster-1 displayed in section A coincides with SOM's cells to which more depressed, rather than non-depressed, respondents from the test cohort (N = 1173) are mapped. NA (dark green) marks the cells to which no respondents from the test cohort were mapped; (D) SOM colored by the average distance between a cell and its neighbors (U-matrix). The green dotted line is the shortest path between the cell with the most depressed respondents (top-right) and with the most non-depressed respondents (bottom-left).

	Cluster-1	Cluster-2	Cluster-3	Total
Number of cells	53	244	328	625
Number of depressed people in the test set	71	179	144	394
Number of non-depressed people in the test set	33	261	485	779
Depression odds	2.15	0.69	0.30	0.51
% male	31.73	40.00	41.34	39.98

Table 3. Overview of the three clusters identified within the SOM.

aspects was significantly different among the three clusters, with Cluster-1 showing the lowest future wellbeing scores.

Further, we examined the features that made Cluster-1 different from the other two clusters. To compare people from different clusters, rather than the depressed versus the non-depressed, we considered only the non-depressed 779 respondents from the test cohort in this section. We observed people with certain mindset features were more likely to be categorized into the depression-prone Cluster-1 (Supplementary Figures 2-4). For example, people in Cluster-1 were less likely to seek personal growth (A1SF3B), had a more vague understanding of life (A1SF3P), were more narrow-minded (A1SF4Y), and were less outgoing (A1SF4A). Other variables in cluster-1 and Cluster-2 that differed significantly (p <0.001) include: life satisfaction (A1PD1), being organized (A1SF4D), and being open with friends (A1SM13) and family (A1SM5) about one's problems.

Some differences between clusters showed that people in Cluster-1 felt lost and out of control (A1SF1C, A1SF1U, A1SF1X, and A1SF1Z) and were less likely to use problem-focused coping mechanisms (A1SF1K, A1SF1L, A1SF1X, and A1SF1Z). See Supplementary Figures 2–4.

The geometric nature of an SOM allowed us to treat it as an actual map to mark paths connecting regions of interest. In this particular case, we chose to mark the path between the depression hotspot in Cluster-1 and the most depression-resilient cell in Cluster-3 (Figure 2D). This path might be interpreted as a sequence of gradual changes that a depressed individual should go through to overcome their condition (Figure 3).

To complete the transition, a respondent would need to change certain attitudes and behaviors. Some of the identified features were trivial and difficult to manipulate; for example, overall life satisfaction



Figure 3. The features that change the most along the path between the respondents mapped to the "island of depression" and those mapped to the "island of mental stability." (A) Top-five attitudes that are generally not shared by mentally stable people but are prevalent among depressed people; (B) Top-five attitudes that are prevalent in mentally stable people but generally not shared by depressed people; (C) The shortest path connecting the SOM cell with the highest prevalence of depressed people (cell-0) to the cell with the highest prevalence of non-depressed people (cell-18). The curves in Panel A and Panel B represent the feature vectors stored in the SOM cells on the path from cell-0 to cell-18. The path displayed in Panel C is also marked in Figure 1D.

(A1PD1) increased toward Cluster-3. Other features offered more convenient therapy targets, such as increasing community involvement (A1SK17F) or openness with family (A1SM5).

A similar approach may be used for a person projected onto any other cell of the SOM to derive primary CBT goals and quantify their effects on well-being or depression symptoms in advance.

DISCUSSION

In this study, we explored the MIDUS dataset to describe the age-dependent component of well-being and to develop a framework for helping people improve their long-term life satisfaction.

To address the first part of our research goal, we preselected 32 psychological variables associated with well-being and created an age and a future well-being predictor. The age predictor, quite expectedly, showed poor performance at MAE = 9.06 years, since the features it was built on were not chosen to be representative of age. On the contrary, as part of our feature selection process, we deliberately removed any variables that might have leaked age. For example, the number of children, income, and the number of comorbidities are expected to be good predictors of age. However, they provide few actionable items and do not represent one's psychological state, and any analysis of these features in the context of aging will offer only trivial results.

The 9.1-year accuracy is marginally better than the baseline but still falls short of the 6.7-year MAE, another similar aging clock, PsychoAge, demonstrated in MIDUS [68]. Thus, we chose not to use the new age predictor to describe the connection between age and well-being.

Well-being as a function of age can be easily visualized to verify the U-shaped hypothesis (Supplementary Figure 5). Since the well-being score distribution in MIDUS is heavily skewed to the right, the mean or median well-being levels were not representative of its age trajectory. Instead, we used the portion of people above 90% of the maximal score as a function of age. Within this definition, the predicted and the real age trends closely resembled each other (MAPE <5.4%). However, neither one of them have been U-shaped, as described elsewhere [16]. This may be due to the inherent differences in well-being scales, or due to MIDUS using a truncated version of the Ryff scale. In this study, certain well-being parameters were observed to trend upward with age (positive relations, environmental mastery, and autonomy), whereas others trended downward (purpose in life and personal growth). In the context of psychological aging, this outcome indicates that there is no "optimal age" at which people reach the apex of wellbeing. Similarly, there is no "age of unhappiness" at which people experience all-round low well-being. Since different aspects of well-being do not uniformly depend on one's psychological age, using the AI-derived metrics of age in practice requires caution. Psychological rejuvenation as a target should be considered within the context of personal priorities and be adjusted according to which well-being aspects a person considers most important.

As we have observed, the "positive relations" aspect of well-being is higher in older people while more exploratory, future-oriented aspects such as "personal growth" and "purpose in life" are higher in the youth (Supplementary Figure 5). This age-related "conversion" of well-being is well described within the socioemotional selectivity theory by Laura Carstensen [69]. The theory posits that as people grow older they "shift from preparatory to consumptive goals". The choices dedicated to exploration or long-term benefits, such as building a career, self-improvement, or a quest for the meaning of life, are ranked higher than the choices promising emotional comfort by younger people. As people grow older and their time horizon dwindles, they start to value positive emotions more, especially when choosing social partners. Interestingly, a series of studies has shown that a similar effect is observed in people presented with the necessity to relocate or people primed with death reflection [70-72]. This suggests that psychological age, as defined in this article, should be malleable and represents one's longevity optimism as well as one's chronological age. Interestingly, new research suggests that in individuals 85 to 90 years old, all measures of optimism were associated with improved 5 year survival [73].

To verify whether the hedonic adaptation hypothesis holds true, we performed a relative importance analysis with EN. In this experiment, we compared whether future well-being estimates (provided by our predictor) or current well-being were a better predictor of actual future well-being. For all six well-being parameters, the deep-learning-derived predictions were significantly closer to actual future well-being than current wellbeing levels (Supplementary Table 2). If the hedonic adaptation hypothesis was true, the opposite should have been observed, with future well-being being influenced more by baseline well-being levels.

Lastly, we set out to develop an SOM-based recommendation engine that provides personalized selfimprovement goals. SOMs are a type of unsupervised learning model that shares technological similarities to neural networks. An untrained SOM stores a vector of length N in each of its cells, also called neurons. Then, it is trained by presenting it with data vectors of length N, each one slightly adjusting the values in the neurons according to an activation function so that the topological structure of the N-dimensional dataset is preserved. In the end, a 2D representation of a multidimensional space is obtained. The basic operation for an SOM is matching or finding a neuron that stores a vector that is the most similar to a newly presented data instance. Based on the position of the best-matching unit (BMU), useful insights may be derived for the new instance.

SOMs have already been used to describe mental health in humans. For example, an SOM classifier has been trained to determine mental disorders in based on patients' transcribed speech [74]. SOMs do not require sample labels to be trained which greatly facilitates data collection and preparation stages of a research project. The cited model has reached high (97%) accuracy while predicting the type of a mental diseases. SOMs have also been employed to detect psychological stress based on biometric data from wearables and phone usage [75]. These models, however, are not based on easily interpretable and actionable features, and thus, are not fit to serve as an mHealth application or a therapy tool.

In this study, an SOM was trained on a collection of non-depressed people and validated on a smaller set of both depressed and non-depressed MIDUS participants. When presented with a new sample, the SOM can return future well-being levels and depression risk to it based on the metadata of any known samples mapped to the same BMU. Most importantly, the SOM can be used for initial screening and highlighting mindset aspects that should first be addressed to optimize a person's longterm well-being.

Applying hierarchical clustering to the SOM yielded three clusters, aggregating significantly different people in

terms of well-being and depression status. Interestingly, the clusters also had different sex distributions, with the depression-prone Cluster-1 being predominantly female, which is in line with existing studies of depression [76].

Features contributing to one's placement in Cluster-1 may be considered primary therapy targets. For example, feeling burdened by close relationships (A1SF1F) is the feature that differentiates depressionsusceptible and mentally stable people the most, according to our model (Figure 3 and Supplementary Figure 3). Thus, a therapy focused primarily on building deep, valuable relationships might be more effective in general compared to a therapy focused on goal-setting (A1SF3T) or social involvement (A1SF4A, Figure 3, and Supplementary Figure 2). The demonstrated path to mental stability would only apply to some people, while the primary therapy targets may be different for individuals who belong to other BMUs. Moreover, different well-being densities across the SOM suggest that the path may be adjusted based on the priorities of a particular person. It is expected that the efficiency of certain therapeutic techniques may also be influenced by factors not present in our model. For example, physical attractiveness is positively correlated with psychological well-being and social warmth [77, 78]. A multitude of lifestyles, habits, and other unaccounted factors can affect how fast a person progresses along an SOM-defined path. Thus, a follow-up study is required to estimate the effect such factors have on one's placement in the SOM.

We also propose that these psychological trajectories may be modeled as a Markov process containing clusters of well-connected states and "bridge" states connecting different clusters (Figure 4). Some attempts to model major depressive disorder as a Markov process have been made before [79, 80]. The cited





works explored the cases of two-state and four-state chains, as well as constant or periodic transition probabilities. In some works, depression is conceptualized as a partially-observable Markov decision process [81]. We believe that our findings can serve as a starting point for a practically applicable mathematical model to unite our understanding of well-being, aging, and mental health. Such modeling would require continuous data collection to observe the fluidity of inmodel psychotypes and approximate state transition probabilities.

CONCLUSION

In this article, we presented a deep learning model based on a psychological questionnaire that can be used to estimate one's psychological age, future well-being, and risk of depression. Based on the same questionnaire, we described an SOM that contained three distinct clusters characterized by different well-being levels and propensity for depression. The SOM can be employed for the purposes of cognitive behavioral therapy and online mental health approaches. Such a tool when implemented may be used as an initial and follow-up assessment tool in face-to-face therapy, or as a standalone consumer application to improve well-being. We also hypothesize that the transitions between the three clusters of psychological profiles may be modeled as a Markov chain for future practical applications.

MATERIALS AND METHODS

Data

The samples used in this article were obtained from MIDUS1 (1995-1996) and MIDUS2 (2004-2006) datasets. Their participants provided information over phone interviews and self-administered questionnaires.

Only the participants present in both MIDUS iterations were included in the study. The target variables used to train models and detect aging-related psychological trends are listed in Table 1. Subjects with missing A1PDEPDX (depression or anhedonia) values were excluded.

The six MIDUS well-being variables are based on the Ryff scale [11]. While the original Ryff scale allocates 20 questions to each aspect of psychological well-being, the MIDUS version of the well-being scale uses only three questions. These questions did not change between MIDUS1 and MIDUS2, apart from a slight change in one of the autonomy questions.

The 3891 participants were separated into training (N = 2718) and test (N = 1173) sets. The test set contained

779 non-depressed individuals and 394 marked as depressed in MIDUS1. No depressed individuals were included in the training set.

Any Supplementary Files related to this study have been deposited to a public Open Science Framework repository [82].

Feature selection

To create the psychological aging clock and the six future well-being predictors, a multi-stage feature selection procedure was employed (Supplementary Figure 1A).

During the first stage, all 2097 MIDUS1 variables were considered. All features relevant to either one of the MIDUS2 well-being features were selected using Boruta v0.3 for Python 3.9 [83]. The union of all relevant features contained 135 variables.

During the second stage, we filtered all the selected features. All non-modifiable features were excluded, and all composite features were deconvoluted into their constituent questions.

For example, A1SEFA ("Paternal affection") was excluded because it represents historical information that cannot be changed. A1SPIWOR ("Perceived inequality in work") was excluded because modifying this feature for a respondent might mean quitting their job to find another—an action associated with the financial risk most people are unwilling to take. A1SKINPO ("Support from family") is a score calculated from four other questions, so it was replaced with A1SM2, A1SM3, A1SM4, and A1SM5.

After filtering the non-modifiable features, 101 variables remained. Boruta was applied once again to determine the constituents of the relevant composite features. All features from Boruta's output were subjected to a collinearity test, and all features with a variance inflation factor above 10 were excluded. The variance inflation factor was calculated with the statsmodels v1.16.0 package for Python 3.9.

Eventually, 32 variables remained, as listed in Table 2. Among these, only seven were from the MIDUS1 wellbeing survey (out of 18). The flow of our feature selection procedure is presented in Supplementary File 1.

Model training

We trained two separate regression models, one for age prediction and one for future well-being prediction. The age model returned only one prediction as output, but the well-being model returned all six parameters simultaneously and was trained in a multilabel manner. We used the 32 features remaining after our feature selection for both predictors.

Feed-forward neural networks with more than three hidden layers were used as predictors with Python 3.9 implementation, using the Keras library (<u>https://keras.io/</u>) with TensorFlow backend (<u>https://www.tensorflow.org</u>) to build and train neural networks. We used the grid search optimization technique over the space of model parameters to find the best performing network architecture separately for both prediction tasks. The MAE loss function was used as an objective for deep neural networks.

The best performing model for age prediction had 4 hidden layers with 256 neurons each, Leaky ReLU activation function [84] after each layer, dropout [85] with 25% probability, and L2 with coefficient 1e-6 after each layer for the purposes of regularization.

The best performing model for future well-being prediction had 4 hidden layers with 400 neurons each, Leaky ReLU activation function after each layer, dropout with 25% probability, and L2 with coefficient 1e-7 after each layer for the purposes of regularization.

Neural networks were trained using five-fold crossvalidation to compensate for overfitting and to obtain more robust performance metrics.

Relative importance analysis

To compare whether the current well-being or wellbeing predicted by MindAge2 was a better determinant of future (MIDUS2) well-being, we used the elastic net (EN) method and treated the variable coefficients as a measure of importance.

EN models were trained to predict either one of the six MIDUS2 well-being features and contained age and sex variables, the corresponding predicted future well-being, and all six MIDUS1 well-being features.

EN was implemented using sklearn v.0.24.2 for Python 3.9. To find the optimal coefficients, a grid search was performed for the L1 ratio (from 0 to 1 with 0.01 step), and penalty weight (1E-5, 1E-4, 1E-3, 1E-2, 1E-1, 0.0, 1.0, 10, 100).

Model statistics

The metrics used to measure the performance of the models in this article are mean absolute error (MAE),

mean absolute percentage error (MAPE), and Person's r. These metrics are defined as follows:

$$MAE = \frac{1}{N} \sum_{i=1}^{N} |Age_{true,i} - Age_{predicted,i}|,$$

where N is the total number of samples

$$MAPE = \frac{100}{N} \sum_{i=1}^{N} \frac{1}{Age_{true,i}} \Big| Age_{true,i} - Age_{predicted,i} \Big|,$$

where N is the total number of samples

Self-organizing maps (SOMs)

We used the SOM approach to map our feature set into a 2D space to visualize the data and provide recommendations through SOM paths [86].

We found the best-performing SOM in terms of quantization error on the presented data using a grid search optimization technique over the space of SOM parameters. Quantization error was the main measurement used to assess SOM quality; it is an average difference between coordinates of SOM-mapped input samples and their corresponding BMUs. An SOM was trained using 32 features remaining after feature selection. We used the MiniSom library for Python 3.9 to train the SOM.

We found that the following parameters performed better for our task: sigma equal to 1.8, learning rate equal to 0.4, Gaussian neighborhood function, Euclidian activation distance, and an SOM size equal to 25×25 .

The Sigma parameter defines the radius around an updated neuron, in which neighbors should also be updated. The neighborhood function is used to update neighbor neurons and determine the rate of change around the winner neuron. Activation distance is the distance used to activate the map or as a main distance function.

The SOM was trained until convergence, when the quantization error stopped decreasing.

We used the Bellman–Ford algorithm to find the weighted shortest path between two points on the SOM; distances between neurons were used as weights. Any other graph-based shortest path algorithm can be used, for example, the Dijkstra algorithm.

Abbreviations

AI: Artificial intelligence; BMU: Best matching unit; CBT: Cognitive behavioral therapy; EN: Elastic net; MAE: Mean absolute error; MAPE: Mean absolute percentage error; MIDUS: Midlife in the United States study; MIDUS1: MIDUS wave 1995-1996; MIDUS2: MIDUS wave 2004-2006; SOM: Selforganizing map.

AUTHOR CONTRIBUTIONS

AZ, Conceptualization, supervision, resources; FG, Methodology, writing (original draft), formal analysis, project administration; KK, Methodology, software, formal analysis; MK, Visualization, data curation, formal analysis; NE, Writing (review and editing), validation.

CONFLICTS OF INTEREST

AZ, FG, KK, MK are affiliated with Deep Longevity, part of Endurance Longevity, a publicly-traded (HK:0575) company.

FUNDING

Publically available data from the MIDUS study was used for this research. Since 1995 the MIDUS study has been funded by the following: John D. and Catherine T. MacArthur Foundation Research Network. National Institute on Aging (P01-AG020166). National institute on Aging (U19-AG051426).

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SUPPLEMENTARY MATERIALS

Supplementary Figures



present in both MIDUS1 and MIDUS2 US-wide survey were considered. Using Boruta, a method of feature selection, we selected only the variables whose values in MIDUS1 were representative of the six Ryff scale well-being parameters in MIDUS2. The initial list of 135 variables was reduced down to 32 features by reapplying Boruta, removing non-modifiable and strongly correlated features. After the first round of Boruta filtering, some selected features represented linear scores. Such scores were decomposed into their elemental variables before the second round of feature reduction. (B) Data partition and models built. Only participants assessed in both MIDUS waves were considered. This cohort was divided into the training set (N=2718) containing only non-depressed individuals and the test set. The same training set was used to train both the aging clock and the SOM. The test set was further subdivided into the cohort of non-depressed (N=394) people, as described in MIDUS1. The SOM was hierarchically clustered prior to mapping the participants from the test set. The independently defined clusters, however, display regions with an increased presence of depressed people and people with low psychological well-being.

1=A lot; 4=Not at all



Cluster no

Supplementary Figure 2. Feature distribution across the three clusters identified within the SOM. Only MIDUS features with a four-point scale are shown. Asterisks mark the variables with significant (p < 0.01, U-test) differences in mean feature value between cluster-1 (depression odds = 2.15) and cluster-2 (depression odds = 0.69). Boxes represent the interquartile region, whiskers protrude no further than 1.5 times the IQR, solid horizontal lines represent the median, and dotted lines represent the mean.

1=Agree strongly; 7=Disagree strongly

Response



Cluster no

Supplementary Figure 3. Feature distribution across the three clusters identified within the SOM. Only MIDUS features with a seven-point scale are shown. Asterisks mark the variables with significant (p < 0.01, U-test) differences in mean feature value between cluster -1 (depression odds = 2.15) and cluster-2 (depression odds = 0.69). Boxes represent the interquartile region, whiskers protrude no further than 1.5 times the IQR, solid horizontal lines represent the median, and dotted lines represent the mean.

3 · 2 · 1 ·

1

2

3

How much obligation would you feel...

0=None;



Supplementary Figure 4. Feature distribution across the three clusters identified within the SOM. Only MIDUS features with a ten-point scale are shown. Asterisks mark the variables with significant (p < 0.01, U-test) differences in mean feature value between cluster -1 (depression odds = 2.15) and cluster-2 (depression odds = 0.69). Boxes represent the interquartile region, whiskers protrude no further than 1.5 times the IQR, solid horizontal lines represent the median, and dotted lines represent the mean.



Supplementary Figure 5. The deep learning model replicates age-associated changes in different well-being aspects. "Top score" is defined as 19–21 points out of 21 for each well-being aspect. MAPE is the mean absolute percentage error between the predicted and chronological age lines. All plots are obtained from the training set (N = 2718), and predicted age values were obtained in five-fold cross-validation.

Supplementary Tables

Taugat mariable	Variable description	MAPE, %		MAE, units		Pearson's r	
l'arget variable	variable description	Training	Test	Training	Test	Training	Test
A1PAGE_M2(adjusted)		9.50	9.79	4.29	4.30	0.91	0.92
A1PAGE_M2	Current age	20.57	21.39	8.97	9.06	0.46	0.47
A1PAGE_M2(baseline)		23.79	23.31	10.31	10.32	0.00	0.00
B1SPWBA1	Autonomy in 10 years	15.84	15.81	2.36	2.36	0.35	0.34
B1SPWBE1	Environmental mastery in 10 years	14.38	18.51	2.10	2.28	0.51	0.48
B1SPWBG1	Personal growth in 10 years	13.91	15.97	2.08	2.25	0.52	0.53
B1SPWBR1	Positive relations in 10 years	16.52	18.47	2.31	2.46	0.58	0.61
B1SPWBS1	Self-acceptance in 10 years	18.42	22.53	2.39	2.62	0.54	0.53
B1SPWBU1	Purpose in life in 10 years	17.25	19.27	2.42	2.59	0.43	0.39

Supplementary Table 1. Predictor accuracy metrics for all target variable

Training metrics were obtained in five-fold cross-validation. MAE is measured in years for age and in points for well-being. The baseline model for age prediction was the median age assignment.

Supplementary Table 2. Relative importance analysis shows that future predicted well-being is more closely associated with actual future well-being than current well-being, age, or sex.

MIDUS2	³² Target Chronol			Predicted	MIDUS1 actual well-being variables					
target variable	description	Is male	age	MIDUS2 variable	A1SPWBR	A1SPWBS	A1SPWBA	A1SPWBG	A1SPWBE	A1SPWBU
B1SPWBR1	Close relations	-0.26 *	0.02	0.43 *	0.23 *	0.08	-0.02	0.02	0.01	0.05
B1SPWBS1	Self-acceptance	0.00	0.01	0.52 *	0.02	0.38 *	0.00	0.00	-0.01	-0.01
B1SPWBA1	Autonomy	0.15 *	0.01	0.47 *	0.01	0.00	0.35 *	0.00	0.00	0.01
B1SPWBG1	Personal growth	-0.32 *	-0.01	0.55 *	0.01	0.07	0.00	0.19 *	-0.02	0.04
B1SPWBE1	Environmental mastery	0.25 *	0.02	0.63 *	0.01	0.09	0.00	0.01	0.10 *	0.00
B1SPWBU1	Purpose in life	0.00	-0.03	0.39 *	0.00	0.00	0.01	0.09	0.02	0.28 *

EN linear regressions were trained to approximate each MIDUS2 well-being variable using sex (binary "Is male"), age, predicted counterpart of the MIDUS2 variable, and all six MIDUS1 well-being variables. The largest coefficients in all cases belonged to model predictions, attesting to them being more important determinants of future well-being than current well-being. Asterisks mark the EN coefficients larger than 0.1 in absolute terms. EN is elastic net (see Methods).

Supplementary Table 3. Correlation between mean well-being scores and depression odds for people mapped to the same SOM cells.

Time point	Well-being parameter	Pearson's r
	Close relationships	-0.24*
-	Self-acceptance	-0.36*
SU	Autonomy	-0.24*
41D	Personal growth	-0.16*
2	Environmental mastery	-0.33
	Purpose in life	-0.15*
lIDUS2	Close relationships	-0.21*
	Self-acceptance	-0.25*
	Autonomy	-0.12*
	Personal growth	-0.11
2	Environmental mastery	-0.25*
	Purpose in life	-0.06

"*" marks the coefficients with p <0.01.

Supplementary File

Please browse Full Text version to see the data of Supplementary File 1.