



Published in final edited form as:

Appl Psychophysiol Biofeedback. 2013 June ; 38(2): 143–155. doi:10.1007/s10484-013-9217-6.

Dynamic Processes in Regulation and Some Implications for Biofeedback and Biobehavioral Interventions

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Abstract

Systems theory has long been applied in psychology, biology, and sociology. This paper applies newer methods of control systems modeling to the assessment of system stability in health and disease. Control systems can be characterized as open or closed systems with feedback loops. Feedback produces oscillatory activity, and the complexity of naturally occurring oscillatory patterns reflects the multiplicity of feedback mechanisms, such that many mechanisms operate simultaneously to control the system. Unstable systems, often associated with poor health, are characterized by absence of oscillation, random noise, or a very simple pattern of oscillation. This modeling approach can be applied to a diverse range of phenomena, including cardiovascular and brain activity, mood and thermal regulation, and social system stability. External system stressors such as disease, psychological stress, injury, or interpersonal conflict may perturb a system, yet simultaneously stimulate oscillatory processes and exercise control mechanisms. Resonance can occur in systems with negative feedback loops, causing high-amplitude oscillations at a single frequency. Resonance effects can be used to strengthen modulatory oscillations, but may obscure other information and control mechanisms, and weaken system stability. Positive as well as negative feedback loops are important for system function and stability. Examples are presented of oscillatory processes in heart rate variability, and regulation of autonomic, thermal, pancreatic and central nervous system processes, as well as in social/organizational systems such as marriages and business organizations. Resonance in negative feedback loops can help stimulate oscillations and exercise control reflexes, but also can deprive the system of important information. Empirical hypotheses derived from this approach are presented, including that moderate stress may enhance health and functioning.

Keywords

psychophysiology; heart rate variability; biofeedback; chaos; systems theory; stress

Introduction

Concepts of ‘cybernetics’, ‘systems’ (dynamic and otherwise), ‘complexity’, ‘chaos’, ‘catastrophe’, and ‘oscillation’ have long been part of discourse in the behavioral, social, and biological sciences. Without the ready availability of mathematical models for the social and biological sciences, a nonmathematical approach to systems theory was prominent in early biofeedback work, and was well articulated by Schwartz and colleagues (Schwartz, 1981; Schwartz et al., 1979). However, a mathematical approach for aircraft communication and control systems had been articulated by Wiener as early as the 1940s (Wiener, 1948, 1961). This model served as a heuristic for describing behavioral, economic, biological, and astronomical systems, among others (Mindell, Segal, & Gerovitch, 2003; Wiener, 1948). In

recent years, mathematical versions of systems theory have been applied in biology and behavioral science, as described below.

Because Wiener's theory had limited predictive power, it was soon combined and supplanted by theoretical systems with greater complexity, such as Shannon's *information theory* (Shannon & Weaver, 1949), which incorporates concepts of channel capacity, noise, and *entropy* (here reflecting the uncertainty in prediction rather than the conventional meaning implying dissolution of order), as well as *chaos theory* (Gleik, 1987; Kellert, 1993), which emphasizes deterministic nonrandom but complex nonlinear relationships among a large number of co-occurring processes, for which statistical prediction is possible. Applications of both information theory (Attneave, 1959; Yockey, 2005) and chaos theory (Elbert et al., 1994; Guastello, Koopmans, & Pincus, 2009; Rossler & Rossler, 1994; Weiss, Garfinkel, Speno, & Ditto, 1994) have been made to the biological and behavioral sciences, and the present article draws upon insights from this work.

For the purposes of this article, we define a system as a variety of elements that interact with one another to form a whole entity. A system's distinct parts are not isolated from each other, thus the characteristics of the whole entity cannot generally be deduced from each of its components. A stable system retains important characteristics even in the face of perturbations that disturb the behavior of specific elements. *Dynamical* systems show changing patterns of action over time, but retain characteristics of an integrated whole. *Cybernetics* is the study of such regulatory processes, including system characteristics like stability, feedback, adaptation, information, and the relationships among them. Dynamical systems have been described as "organized" or "disorganized" and of greater or lesser complexity. Simpler organized systems can often be described and modeled in linear terms, although nonlinear approaches usually are more appropriate for more complex systems.

A *control* system is defined as a stable system, in which system elements interact to preserve system stability, both for internal control and in response to perturbations caused by external influences. Systems can be modeled as *closed-loop* or *open-loop*. An open-loop system involves only a system response to an outside event. It does not provide an internal mechanism for monitoring system performance or preserving system stability. For example, open-loop models have been used to biomimetically describe the flying patterns of *drosophila melanogaster* after stimulation (Fry, Rohrseitz, Straw, & Dickinson, 2008). Behavioral and chemical interventions designed to change system behavior are examples of open loop control processes, such as when we teach a person to lower muscle tension or blood pressure directly, rather than by using natural feedback mechanisms.

A closed-loop model includes internal regulation, whereby system responses to outside stimulation are processed through internal feedback loops that monitor and adjust the response of the system itself. As a discipline, biofeedback is particularly concerned with closed-loop, internal regulatory systems, and tries to maximize their effectiveness. A good example of a closed loop system is the baroreflex (Vaschillo, Lehrer, Rische, & Konstantinov, 2002), as will be described in more detail later in this article. However, when biofeedback or relaxation is used as a strategic intervention to control acute symptomatology, the resultant process can be described as an open-loop system.

A common control mechanism leading to system stability in closed loop systems is the negative feedback loop. Negative feedback loops preserve stability by activating opposing processes that act to modulate change, though importantly, stable systems tend to contain both positive and negative feedback loops (Cinquin & Demongeot, 2002; Pigolott, Krishna, & Jensen, 2007), as will be described below. Negative feedback systems usually have inherent delays caused by the time needed for opposing system elements to effect change in

each other. Negative feedback loops with delays cause oscillations at particular frequencies (Cinquin & Demongeot, 2002). A very simple negative feedback system is the thermostat, which responds to changes in ambient temperature by switching a heating or air conditioning system on and off. Here the delay is dependent on such factors as outside air temperature, insulation, and volume of inside air. Thus, room temperature oscillates around the thermostatic setting, but is rarely constant. The negative feedback loop in the baroreflex system also contains delays, as will be described below.

Oscillatory properties have been observed in systems as small as the cell, and as large as societies. For example, at a cellular level, oscillations have been used to map various negative feedback loops in protein concentrations, and circadian gene expressions in bacteria (Pigolotti, Krishna, & Jensen, 2007). In social systems oscillations describe patterns of voting, marital relationships, and mood.

The baroreflex system is a good example of a stabilizing oscillatory process. It produces periodic fluctuations in blood pressure, heart rate, and vascular tone that reflect modulatory control. External influences on blood pressure (e.g., aperiodic stress or exercise) induce blood pressure changes that, without a control mechanism, could burst blood vessels or cause circulatory insufficiency, depending on the direction of blood pressure change. The baroreflex responds to blood pressure increases by slowing heart rate and causing vasodilatation, which, in turn, produces a decrease in blood pressure. Each time blood pressure decreases, the same mechanism produces vasoconstriction and heart rate acceleration, which causes blood pressure to rise again. Changes in blood pressure are delayed by a number of factors, including inertia in blood flow and blood vessel plasticity (Vaschillo et al, 2002; Vaschillo, Vaschillo, & Lehrer, 2006). The delay in blood pressure changes following heart rate change averages about five seconds, thus yielding the ubiquitous heart rate oscillation of about 10 seconds (Vaschillo et al., 2002), i.e., five seconds for the baroreflex to cause an increase in blood pressure when the baroreflex responds to a decrease in blood pressure, and five seconds to cause a decrease in blood pressure following a baroreflex response to a blood pressure increase. In actuality, however, the baroreflex system is composed of *two* closed linear systems with negative feedback loops – it also contains a slower vascular tone loop with a rhythm of approximately 0.02–0.03 Hz, which has a slower response time than the heart rate loop (Rahman, Pechnik, Gross, Sewell, & Goldstein, 2011; Vaschillo, Vaschillo, Buckman, Pandina, & Bates, 2012).

It should be noted that a full description of baroreflex control would not be restricted to purely mechanical properties of the reflex. The baroreflex system shows characteristics of neuroplasticity. Dworkin (Dworkin, 1993) demonstrated that the baroreflex can adapt to changing environments through classical conditioning. Consistent with Dworkin's findings, Lehrer and colleagues demonstrated that regular stimulation of the baroreflex through biofeedback can increase baseline baroreflex gain, suggesting an improvement in regulatory capacity over blood pressure changes (Lehrer et al., 2003).

Complex control of systems

Most psychological and biological systems are much more complex than thermostats, with multiple control mechanisms, and multiple overlapping oscillations, the quantitative characteristics of which depend on the frequency and phase of each component oscillator, as well as the interactions among them (Hyndman, 1974). Changes to one system component (control mechanism) affect others, so the individual control mechanisms are interdependent, thus promoting biological vitality and complexity. Additionally, it should be noted that closed loop systems with negative feedback loops often require simultaneous open loop

processes in order to function properly, to provide sufficient stimulation to stimulate negative feedback loop activity (Feigl, 1998). This adds to the complexity.

Simple control, such as control of room temperature, can typically be described with linear mathematical models, but when multiple controllers operate, each with a characteristic frequency, attendant mathematical computations become more complex. Existence of multiple overlapping control systems assures system stability in the face of various perturbations by providing multiple “backups”. A high degree of entropy, reflecting a high degree of nonlinearity and/or a need for multiple linear formulas to describe the system, is often related to health and biological system stability. Conversely, simple periodicity or random variability (“white noise”) often underlies pathology in disordered systems (Goldberger, Peng, & Lipsitz, 2002), reflecting, respectively, either the presence of only a single control oscillator, or absence of any regulatory activity.

Oscillatory patterns with greater complexity, such as those that occur when a number of oscillatory patterns overlap, are described as “chaotic”. Chaos reflects the simultaneous operation of numerous control processes. Although the healthy cardiovascular system can be effectively modeled using linear statistics (Berntson, et al., 1997), it also has chaotic properties, which diminish when cardiac function is impaired or an individual is in danger of dying from physiological decompensation (Arzeno, Kearney, Eckberg, Nolan, & Poon, 2007). Similarly, healthy adjustment is characterized by chaotic rhythms in appetite (Berthoud, 2006) and mood (Haffen & Sechter, 2006), among other biobehavioral dimensions. Also, negative feedback loops often work in concert with open loops as described above, and with positive feedback loops and resonance characteristics, as described below, to maintain optimal functionality.

Oscillation, control, and health

Many oscillatory systems in the body, such as neural (Berthouze, James, & Farmer, 2010), cardiovascular (Berntson et al., 1997), and circadian (Garau, Aparicio, Rial, Nicolau, & Esteban, 2006) tend to degrade with age, reflecting a decrease in adaptive capacity. In addition, there are also pronounced age-related decreases in baroreflex gain (Brown, Hecht, Weih, Neundorfer, & Hilz, 2003), while blood pressure oscillations tend to increase with age (Cugini et al., 2003), perhaps reflecting degrading of control over this otherwise tightly regulated function. Generally speaking, oscillatory patterns in a variety of biological control systems tend to be a characteristic of youth, fitness, and health.

Oscillatory properties do not only represent activation of specific control reflexes (e.g., the baroreflex). They provide information as well. Periodic stimulation is required for system control in feedback networks. This principle has been proposed for cellular behavior (Klevecz, Li, Marcus, & Frankel, 2008) as well as macro biological and behavioral systems, as in discussions of open loop processes described above, and both resonance and stochastic processes, described below. External stimulation “perturbs” a control system causing it to oscillate. Without such oscillations, and hence, without noise (which can function as the perturbation that pushes the system to oscillate), systems may fail to get the information they need to function properly. The tendency for noise to stimulate system oscillation is called *stochastic resonance*. There is evidence that interoceptive and exteroceptive noise enhances the sensitivity of a variety of control processes, from micro to macro levels (Wiesenfeld & Moss, 1995). Studies of stochastic resonance show that noise can facilitate control functions, as in optimization of monosynaptic afferent reflexes in the motor system of the cat by noisy stretching of a synergistic muscle (Martinez, Perez, Mirasso, & Manjarrez, 2007). In humans, extreme absence of system perturbation, as in sensory deprivation, can cause

regulation go awry, resulting in psychotic-like sensory and emotional symptoms (Mason & Brady, 2009).

One could, therefore, speculate that psychological stress may have some beneficial effects, by introducing noise into neural regulatory processes, just as exercise helps maintain many physiological processes. Since, in biological systems, oscillations tend to reflect the operation of reflex systems involved in modulation and self-control, presumably a complete lack of stimulation would deny exercise to these reflexes, causing them to atrophy. Under normal circumstances, however, the environment provides sufficient perturbations in biological and behavioral systems to preserve their normal function. In psychophysiological theories of stress effects, this formulation overlaps with the more recent concept of “allostasis” and “allostatic overload” (Juster, McEwen, & Lupien, 2010). *Allostasis* refers to stability achieved through variability, and is analogous to the activity of oscillating control processes stimulated by stochastic noise, or general unorganized environmental stimulation. A well functioning personality or biological system responds well to moderate environmental stressors, and, in fact, may be energized by them, as in Yerkes and Dodson’s model of stress and performance, wherein moderate stress enhances performance in moderately difficult tasks, while very low or very high levels of stress can be detrimental (Yerkes & Dodson, 1908). *Allostatic overload* (McEwen, 2004) occurs when environmental stresses are either too great or too prolonged, or when various control reflexes become exhausted.

Resonance: A source of stimulation and simplicity in system oscillation

In addition to oscillation, another characteristic of negative feedback systems with a delay, is resonance (Grodins, 1963). When outside stimulation causes a system oscillation at a particular frequency, and no other forces are at work to dampen the oscillations, negative feedback loops can themselves destabilize the system by causing increasingly large resonance frequency oscillations, to the point where information from other frequencies no longer gets processed. A common example of a ‘run-away’ negative feedback loop, resonance effect is the “Larsen effect” that occurs when a microphone is placed near a speaker, causing a high-pitched squeal at a single frequency (Weaver & Lobkis, 2006). In such a case, a single high-amplitude oscillation, triggered either by noise (stochastic resonance) or by pulsatile stimulation close to the resonance frequency, can obscure the effect of otherwise meaningful perturbations. In the human body, resonance effects in the postural control loop have been implicated in postural impairment in Parkinson’s disease (Maurer, Mergner, & Peterka, 2004). Resonance has been similarly implicated in age-related impairment in gait control (Thurner, Mittermaier, & Ehrenberger, 2002).

In terms of psychophysiology, it is conceivable that resonance effects might inhibit the fight-or-flight reflex because a single homeostatic process may overwhelm reflexes needed to confront an environmental stressor. One could speculate that resonance effects, stochastic and otherwise, might deprive the system of information from various internal control mechanisms, as when resonance-frequency oscillations are at such a high amplitude that they obliterate information from reflexes operating at different frequencies. Thus, it is possible that resonant oscillations in the heart rate baroreflex system at 6/minute could weaken the effects of other control reflexes, by greatly diminishing their relative size and depriving the system of information. This is a hypothesis for future research. To prevent a ‘run-away’ resonance process, systems must be dampened (Siebert, 1986). Dampening effects on heart rate amplitude during heart rate variability biofeedback probably stem from inherent limitations in the ability of the heart to respond at extreme levels.

The literature on heart rate variability biofeedback, however, shows that not all resonance effects are detrimental. Resonance in the cardiovascular system produced by a rhythm in the baroreflex is the basis of the beneficial effects of heart rate variability biofeedback. The baroreflex provides a negative feedback loop for controlling blood pressure, such that heart rate falls when blood pressure rises, and vice-versa when blood pressure falls, thus modulating blood pressure fluctuations. The resonance frequency of this system appears to be related to the individual's blood volume (Vaschillo, et al., 2006), with a lower resonance frequency among people with a larger blood supply (men and taller people, vs. women and shorter people). The resonance frequency appears to be determined by changes in hemodynamic inertia resulting from fluctuating heart rate. When this resonance system is activated by a respiration rate close to the resonance frequency, oscillations in heart rate become very large. This high-amplitude stimulation of the baroreflexes appears to strengthen them (Lehrer et al., 2003), and is perhaps the cause of the various salutary effects of heart rate variability biofeedback.

Heart rate variability biofeedback has been shown to restore autonomic control that has been acutely repressed by experimental exposure to inflammatory cytokines (Lehrer et al., 2010), and appears to ameliorate a number of disorders characterized by autonomic and/or emotional dysregulation (Lehrer, 2007), including hypertension (Lin et al., 2012; Nolan et al., 2010; Reineke, 2008), asthma (Lehrer et al., 2004), anxiety/stress (Hallman, Olsson, von Scheele, Melin, & Lyskov, 2011; Henriques, Keffer, Abrahamson, & Horst, 2011; Shenefelt, 2010), depression (Beckham, Greene, & Meltzer-Brody, 2013; Karavidas et al., 2007; Patron et al., in press; Siepmann, Aykac, Unterdorfer, Petrowski, & Mueck-Weymann, 2008), and chronic pain (Hallman et al., 2011; Sowder, Gevirtz, Shapiro, & Ebert, 2010; Strine, 2004; Yetwin, 2012), while improving athletic performance (Paul & Garg, 2012). We might, however, theorize that the effects of constant breathing at resonance frequency would not be advantageous. Chronic resonance frequency breathing could theoretically weaken or obstruct reflexes dependent on oscillations at other frequencies. Thus, individuals practicing heart rate variability biofeedback are instructed to practice for a relatively brief period of time daily (usually about 20 minutes) and to use the technique strategically when symptomatic.

Positive feedback loops

The body also contains multiple positive feedback loops, whereby change in a particular direction facilitates greater change in that direction. Although positive feedback can provide a stimulating effect in an open loop system, it also plays a role in closed loop systems. In psychophysiology, positive feedback loops often are involved in maladaptive processes (Thayer & Lane, 2000). A prime example is anxiety and depression's propensity to amplify their inherent symptomology by triggering greater sensitivity to anxious or depressive thoughts, thus creating a vicious cycle. On the cellular level, positive feedback loops may be necessary to stimulate modulatory negative feedback loops (Kurbel, 2012; Nishi et al., 2000; Pomeroy, Kim, & Ferrell, 2005; Prochazka, Gillard, & Bennett, 1997; Tsai et al., 2008).

However, positive feedback loops can also play an important role in enhancing emotional control. The psychological augmenting effects of positive feedback may add to oscillation amplitude during HRV biofeedback, or the sympatholytic effect of muscle relaxation. This may result from a spiraling relaxation effect. Just as anxiety begets more anxiety by sensitizing anxiogenic brain circuits (a positive feedback loop) (Bos, Hoenders, & de Jonge, 2012; Chemtob, Roitblat, Hamada, Carlson, & Twentyman, 1988; Krantz et al., 1987; Thayer & Lane, 2000), heart rate variability biofeedback or muscle relaxation therapy may compound relaxation effects.

Positive feedback loops can also play a role in creating system complexity, which may increase stability by creating a multiplicity of stable states (Plahte, Mestl, & Omholt, 1995), as well as by maintaining oscillation amplitude during frequency adjustment (Tsai et al., 2008). Positive feedback loops have been shown to promote load compensation in control of movement (Prochazka et al., 1997), and to be necessary to prevent dampening of oscillations in cellular function (Pomeroy et al., 2005). Positive feedback loops also are important for the maintenance of the ovulatory cycle (Kurbel, 2012), and for propagating dopaminergic signaling (Nishi et al., 2000). Without some form of regular perturbations, oscillations will gradually decline in amplitude and disappear. Positive feedback loops can provide these perturbations.

Simplicity, complexity, and randomness

The dimensions of simplicity and complexity are important for understanding psychobiological control, and can be usefully incorporated into theories of applied psychophysiology. Simplicity in a biological or behavioral control system can have multiple sources. In addition to reflecting the effects of resonance, simplicity may result from system fatigue due to allostatic overload (Juster et al., 2010), or from biological damage to system components, such as when heart failure leads to diminished entropy in heart rate (Ho, Lin, Lin, & Lo, 2011; Isler & Kuntalp, 2007; Liu et al., 2011). Simplicity in heart rhythms as a sign of cardiac pathology was discussed earlier. Emotional rigidity, or simplicity, is well described in DSM-IV (American Psychiatric Association, 2000) as a sign of psychopathology, which is characterized by a tendency to respond to a wide variety of situations with a stereotyped response: sadness, anxiety, suspiciousness, anger, etc. However, just as simplicity is a sign of poor adaptation, so is random variation, suggesting lack of modulatory control. Pathological examples of random fluctuations include manic or depressive episodes in bipolar disorder, as well as certain cardiac arrhythmias such as pre-ventricular or pre-atrial contractions, asthma exacerbations, and so on.

Work in the tradition of chaos and information theories has provided mathematical tools for describing organized complexity of temporal patterns in terms of entropy, as described in information theory, including Shannon's "spectral entropy" calculations (Shannon, 1948), Pincus' "approximate entropy" (Pincus, 1991, 1998), Lempel and Ziv's method for evaluating "randomness" (Lempel & Ziv, 1976). These measures represent various approaches to calculating relative unpredictability of fluctuation patterns in a time series, or the number of dimensions, or fractals (Mandelbrot, 1983) necessary to describe a data set mathematically.

Autonomic regulation

Healthy regulation is often characterized as sympathetic and parasympathetic activity converging in a limit cycle around a critical value, such as is the case of cortisol-vasopressin or acetylcholine-epinephrine reactions, where each process induces a compensatory reaction in the other (Bernard-Weil, 1986). Thus, sympathetic activity may simultaneously suppress parasympathetic activity, but increase parasympathetic *reactivity*, thereby producing an oscillation. During extremely stressful stimulation, an individual may show wild oscillations in various autonomic functions, both sympathetic and parasympathetic (e.g., bronchodilation and constriction, increased and decreased blood pressure, energy and fatigue, high versus low rates of peristalsis as in constipation and diarrhea, etc.), although these oscillations are sometimes paradoxically suppressed in severe emotional reactions (Gellhorn, 1969, 1970).

Heart rate changes in response to stimulation appear to be important modulators of systemic response. Reduced cardiac variability in response to stimulation is related to a variety of

pathophysiological states (Montano et al., 2009), while increased heart rate variability and vagal influence on the heart are positively correlated with recovery after physical exercise stress (Chen et al., 2011). Thus flexibility in cardiac response to stimulation, and higher vagal influence on the heart (measured as high-frequency heart rate variability, reflecting both baroreflex control and modulation of respiratory function) are related to better adaptation to interoceptive and exteroceptive demands (as evinced by rapid recovery). This is another example of ways in which complexity and oscillatory function are related to health and adaptability.

Thermoregulation

Thermoregulation is another system in which several systems converge around a critical value. Until the mid 20th Century, stability and circadian variability in core body temperature were considered homeostatic responses to environmental triggers, such as light intensity or daily routine. In fact, core body temperature is partially regulated by an endogenous clock in a central control mechanism, which integrates multiple systems including thermoreception of ambient temperature, systems for conserving or dissipating heat, as well as those for generating heat, and endogenous circadian regulators. As with heart rate variability, circadian thermal variability decreases with age (Weinert & Waterhouse, 2007). A relationship between circadian thermal oscillation amplitude and general homeostasis can be hypothesized, but further evidence is needed to confirm its existence.

Oscillation, complexity, and disease

Heart rate variability complexity is indicative of cardiac and neurocardiac flexibility and adaptability, and may be diminished significantly by pathology. A healthy heart can respond to various moment-to-moment physical and emotional demands, while the diseased heart does not exhibit such flexibility. Although oscillations may persist in the diseased heart, aging and disease cause increasingly more regular oscillatory patterns reflecting a decrease in mechanisms of cardiac control (Goldberger et al., 2002). Such noncomplex patterns of variability ultimately predict death in the critically ill (Arzeno et al., 2007; Norris, Anderson, Jenkins, Williams, & Morris, 2008; Norris, Stein, & Morris, 2008).

Similarly, a disturbance of complementary oscillatory patterns in pancreatic islet beta cells is associated with impaired glucose tolerance and diabetes mellitus (Polonsky, Sturis, & Van Cauter, 2000). Under normal conditions, oscillations in nervous output from pancreatic ganglia cause pancreatic islet beta cells to secrete insulin in a coordinated pulsatile manner, at two distinct periodicities, 1) in small amplitude oscillations with a periodicity of 5–15 minutes, and 2) in larger amplitude ultradian oscillations ranging from 80 to 150 minutes, particularly in response to the ingestion of food or intravenous glucose perfusion (Polonsky et al., 2000). This oscillatory rhythm operates through an insulin-glucose negative feedback loop, with a delay between plasma insulin and hepatic glucose production. Compared with steady-state infusion, this oscillation promotes more efficient glucose utilization (Tolic, Mosekilde, & Sturis, 2000). When these oscillatory rhythms are disturbed, such as in diabetes mellitus type II, decreased oscillatory complexity is associated with system decompensation (Bertram, Sherman, & Satin, 2010).

Mental illness

Mood and behavior appear to be governed by many of the same concepts used to describe oscillatory behavior in physiological processes. As with physiological systems, organized complexity is the hallmark of healthy psychological behavior. Both hyper and hypostabilities may characterize a breakdown in normal mental functioning, and leave a person

psychologically vulnerable to the deleterious effects of stress. Bauer and Whybrow (1995) found that while participants with bipolar disorder showed some brief periods of fairly well defined cycling in mood, their overall pattern of mood changes were less complex than controls. Paulus et al. compared patients with schizophrenia to healthy individuals in a binary choice study, and found that participants with schizophrenia showed a simpler and more predictable pattern of response, apparently reflecting decreased adaptability in cognitive processing (Paulus & Braff, 2003; Paulus, Geyer, & Braff, 1996). A study of laterality in electrodermal activity found decreased activity and complexity in the left hand, compared with the right, among depressed patients, and decreased activity and complexity in the right hand, compared with the left, among patients with schizophrenia, while there were no laterality differences among healthy individuals (Bob, 2007). These results are consistent with theories of right hemisphere dysfunction in depression (represented by decreased adaptiveness in this instance), and left hemisphere dysfunction in schizophrenia. However, even some psychopathological states appear to possess chaotic organization. Tschacher, et al. (Tschacher, Scheier, & Hashimoto, 1997) recorded occurrence of psychotic symptoms among 14 patients with schizophrenia over time, and found a complex, nonlinear time course in eight of them. Others showed more random psychotic behavior, suggesting that the patients were responding to environmental cues, not modulated by control processes.

EEG oscillations reflect system functions

Systems theory also applies to neurons' activity in the brain, whose functioning has been shown to be interdependent with other nearby neurons and even neurons in different brain regions. The complexity and chaotic characteristics of brain function, particularly as studied by electroencephalogram (EEG), has been thoroughly reviewed elsewhere (Buzsaki, 2006). Briefly, EEG reflects summations of excitatory postsynaptic potentials that neurons emit as discrete events. When neurons fire synchronously, relatively high-amplitude slow waves occur, in a relatively simple pattern; when they fire desynchronously, the numerous potentials can both augment and cancel each other out, generally resulting in lower-amplitude, more complex signals at higher frequencies (Onton, Westerfield, Townsend, & Makeig, 2006). Fast-firing cells in the cortex may contribute to these effects, as does the interaction between excitatory and inhibitory processes. Faster, low-amplitude oscillations tend to predominate during states of arousal, attention, and cognitive processing (Timofeev & Chauvette, 2011), and in general, during periods of greater cortical activity (Klimesch, 1996).

Oscillations in EEG signal reflect the interplay of various brain processes involved in cortical integration, such that specific kinds of mental activity may be reflected in definable patterns of activity at specific frequencies, and at specific electrode locations (Gottesmann, 1999; Jacobs, 2001). Slower waves from the cortex reflect decreased arousal at the measured EEG sites (i.e., fewer simultaneous processes producing potentials at various frequencies). Low levels of alpha and beta activity (the faster, higher-frequency ranges) and/or increased waking slow-wave activity are associated with a lack of inhibitory control over behavior (Knyazev, 2007). Similarly a greater frontal theta beta ratio is related to decreased emotional response inhibition (Putman, van Peer, Maimari, & van der Werff, 2010). Greater delta and theta activity and lower alpha activity have been found in the EEGs of children with fetal alcohol syndrome (Kaneko, Phillips, Riley, & Ehlers, 1996). This pattern is also found in attention deficit disorder, lower cognitive abilities, impaired motor control, hypoglycemia, and antisocial behaviors. It is even found during deep relaxation and sleep, while exposed to hypoxia, fasting, and in sexual arousal and orgasm (Knyazev, 2007). Conversely, delta, theta, and lower-frequency alpha activity are inhibited when people are awake and cognitively active. A recent study found that, during tasks with increased cognitive demand,

increases occurred in the EEG frequency with the greatest activity across electrode sites, but a decrease in EEG entropy (Zarjam, Epps, & Chen, 2011).

Oscillations are not smoothly distributed across the frequency spectrum. Resonance structures have been identified at frequencies of 4, 10, 20, and 40 Hz (Erol Basar, 1999) with the largest body of research on the 10 Hz rhythm, the center of the alpha frequency band (8 – 12 Hz). Resonance properties can be determined by a variety of methods. For instance, one can stimulate the system at specific frequencies with photic stimulation, and determine the frequencies at which high-amplitude oscillations are obtained (Fedotchev, Bondar, & Konovalov, 1990; Herrmann, 2001; Spiegler, Knösche, Schwab, Haueisen, & Atay, 2011). One may also use computer simulations based on known frequency characteristics of particular nerve cells (Kasevich & LaBerge, 2011) or from evoked potentials (Basar, Gönder, & Ungan, 1976; Bayram et al., 2011; Winterer et al., 1999). Alternatively, one can stimulate an individual with audible noise and measure the obtained frequency peaks (stochastic resonance) (Ward, MacLean, & Kirschner, 2010). Resonances at these frequencies suggest the existence of specific positive feedback reflex arcs at each resonance frequency, related to particular neural processes contributing to each resonance frequency.

Thalamocortical processes have been studied and modeled, and rhythm bands containing each of these frequencies have been described in a large literature (theta or delta for 4 Hz, alpha for 10 Hz, beta for 20 Hz, and gamma for 40 Hz) (Knyazev, 2007). It has been theorized that delta oscillations reflect activity of motivational systems, while theta oscillations reflect emotional regulation, and alpha oscillations reflect inhibitory processes (Knyazev, 2007). Some theorists have related specific resonance frequencies to the period length of experimentally evoked action potentials, event-related potentials, synchronies, or asynchronies (Pfurtscheller & Lopes da Silva, 1999). Patterns of theta and alpha-frequency resonances triggered by memory consolidation processes have been identified (Klimesch, Schack, & Sauseng, 2005). It is also known that memory consolidation occurs during sleep (Fogel & Smith, 2011), which is characterized by high-amplitude slow EEG waves primarily stemming from the brain stem, with cortical modulation of lower centers, perhaps from a single oscillator (Crunelli & Hughes, 2010).

Phase relationships in EEG activity among various brain areas have also been studied, appearing to vary systematically. For example, occipital phase synchrony in the alpha rhythm tends to be greatest during relaxation with eyes closed, and decreases with greater arousal (Gengerelli, 1978). Experience of “pure consciousness” in transcendental meditation appears to be characterized by alpha phase synchrony from the frontal and central areas (Orme-Johnson & Haynes, 1981). Although phase synchrony may reflect tight coordination among brain sites, more complex forms of coordination have also been modeled, often involving varying frequencies, varying orderly phase relationships, and nonlinear relationships, where frequent changes in phase relationships among brain areas reflect complexity of cognitive processing (Basar, 2006; Panzeri, Brunel, Logothetis, & Kayser, 2010; Tognoli & Kelso, 2009).

Measures of chaos and complexity applied to EEG data suggest a simpler pattern of brain organization, with, fewer control processes, among patients with schizophrenia (Roschke, Fell, & Beckmann, 1995) and depression (Nandrino et al., 1994; Pezard et al., 1996). Pezard et al. found that EEG patterns became more complex among depressed patients, whose depression improved, becoming indistinguishable from those among healthy individuals, while the pattern of decreased complexity persisted among individuals whose depression did not improve (Pezard et al., 1996). Other studies have found risk for autistic spectrum disorder (Bosl, Tierney, Tager-Flusberg, & Nelson, 2011) and the presence of attention

deficit hyperactivity disorder (Sohn et al., 2010) to be related to decreased average EEG complexity.

Social systems

Humans live as couples, in families and in neighborhoods or larger social and economic networks. Systems theory has been applied to social systems, and, indeed, oscillatory patterns of social system behavior can reflect negative feedback loops, which could contribute to stability. Stability in a social system generally represents a stable pattern of interaction, implying the action of mechanisms that resist change. Oscillatory patterns of openness and closedness to new ideas have been described in societies (Klapp, 1975), as have variations in societal norms about expression of emotion (Cancian & Gordon, 1988), although presumably extremes in either of these characteristics trigger a consequent tendency in the opposite direction. Voter patterns appear to have oscillatory characteristics, which, in the United States, have been quantified as thirty-year oscillatory patterns in party alignment (Mayhew, 2002). The cause for this oscillation period may reflect generational change. Perhaps people see the problems caused by results of their parents' voting patterns, and react to this by changing course. Although delays, chaos, and resonance characteristics in social feedback loops have not been measured, the existence of specific frequency patterns in some of them suggest that these characteristics may be present.

Dyadic systems may also have oscillatory qualities. Gottman and his colleagues have taken a mathematical approach to assessing marital systems (Gottman, Murray, Swanson, Tyson, & Swanson, 2002). They have applied concepts of nonlinear dynamics and catastrophe theory, which posit complete periodic restructuring of systems when perturbed by certain forces. They propose that marriages have one or more "set points" for nature and style of interaction, some of which may be favorable to marital satisfaction and stability, and others unfavorable. From this perspective, they have developed formulas that predict marital stability and divorce. They record verbatim transcripts of marital interactions, where they note the number of times that individuals respond to each other's content with specific affect, including such variables as threshold for negativity, and frequency of positive and negative interactions. In marriages characterized by reciprocal negativity and paucity of positivity in interactions, couples can be trained to modify their individual behavior. Thus, based on positive feedback loops, marital systems, wherein negativity begets negativity, may be altered so that positivity begets positivity. The overarching goal is to move the interaction system from a negative steady state to one that is more positive. Repair and dampening functions have been calculated, which can modify the positive or negative influence of one partner on another in various steady state conditions. The investigators report conflicting data about whether rigidity in behavior predicts poor marriage outcome, but show how intervention can affect the system of interaction and, ultimately, marital satisfaction. Their model includes the interaction of perception of well-being in the relationship, the flux over of negative and positive behaviors, and physiological responses.

Systems theory has also been applied in analysis of corporate structures (Weick, 2009), but concepts of oscillation, delay, and influences of positive and negative feedback loops have not been studied. Corporations are systems, just as those defined above. They are constantly bombarded by changes in market conditions, competition, etc., and need to maximize the adaptiveness with which they respond to changing conditions, while still maintaining their structure. Information and communication, both from outside of and within the organization, has been proposed as a medium for achieving these ends. Utilization of feedback from information allows an organization to monitor both external events and internal processes. For example, if an organization's norms do not allow for transmission of bad news to its CEO, then poor, uninformed decisions are likely to be made. Various feedback mechanisms

can prevent change from happening (e.g., not heeding information identified as coming from the “wrong channel”). Although we have found little research using time series analysis to discover patterns of oscillations in negative feedback loops comprising organizational structures, examples of feedback loops in organizational behavior have been described (Weick, 2009). Presumably oscillations occur in organizational communication as well as in other control systems, and might be quantifiable on a number of dimensions, such as commands to change operational procedures to adapt to external pressures, oscillating with instructions to follow internal protocols, pressured work versus relaxation, and supportive versus critical communications from superiors.

Implications for research

In psychophysiology, linear models are frequently employed to predict one variable from another. However, because physiological systems are so complex, such prediction does not truly characterize physiological states. Time series models have become available that can be used to better model these systems. The complexity of biological and behavioral systems and the fact that complex systems can be mathematically modeled present a challenge to psychobehavioral research. This paper proposes that we re-examine the systems approach, in light of methodologies recently proposed. Cybernetic systems approaches in biology and psychology present practical challenges such as in the assessment and measurement of multiple systems, and large quantities of time series data. Nevertheless, quantification of complex systems involved in personality, illness, marital or social harmony, and stress reactivity may give us greater insight into ways of promoting health and managing disease, and are thus amply justified.

Hypotheses generated by this review

This review would lead us to posit certain hypotheses. These include:

1. Any stable system can be modeled using time series analysis.
2. Internal control mechanisms can be modeled as closed negative feedback loop systems with oscillatory properties. Decreases in oscillatory properties will be associated with less adaptability to perturbations (stressors), and greater vulnerability to system failure.
3. Stable systems are complex, indicating a multiplicity of control mechanisms; however, they are not random.
4. Adaptability of a system can be modeled from the effect of perturbations on its various closed-loop systems of internal control. Changes in any elements due to system perturbations will return toward a pattern of stable equilibrium.
5. A stable system does not necessarily have a single level of stability. This may change depending on demands placed on the system.
6. Disease, injury, stress, and older age all reduce adaptability, and will be reflected in decreases in oscillations of control systems, complexity of oscillations, and ability to return to equilibrium after perturbation. They may also be characterized by greater periods of randomness.
7. This model can be applied to all systems: biological, behavioral, cognitive, emotional, social, organizational, etc. Elements of systems could be quite varied, and include such diverse phenomena as behaviors, communications, and biochemical or physiological functions.

Particular implications for applied psychophysiology and biofeedback

The present analysis indicates that further investigation of patterns of oscillation in various psychobiological systems is warranted. Better identification of oscillatory systems may lead to the elucidation of homeostatic or allostatic control mechanisms in such systems, and may ultimately lead to the development of ways to exercise and strengthen reflexes that strengthen allostatic capacity. Ideally, these exercises could render an individual more resilient to effects of environmental stressors, improve performance, and strengthen resistance to psychological and physical disease. Systematic use of various quantitative approaches to assessing oscillation and complexity could contribute to advances in this field.

Work in this direction has already begun in studies of heart rate variability and EEG biofeedback. Heart rate variability biofeedback is based on the explicit mechanism involving stimulation and exercise of modulatory reflexes by stimulating resonance characteristics of an oscillating system. For this method, the theory proposed in this paper predicts that only *periodic* practice of the technique would be beneficial, while *constant* practice of the technique might actually be iatrogenic, since the resonance frequency oscillations would prevent chaotic variability in heart rate, and possibly decrease adaptability, if they were *continuously* present. However, strengthening of homeostatic reflexes, with a consequent increase in chaotic variability, appears to be beneficial. Periodic stimulation of a reflex seems to improve its efficiency. We have found this to be the case with the baroreflex (Lehrer et al., 2003). As yet, no studies have reported changes in chaotic variability in the resting state after a course of heart rate variability biofeedback; although other measures of variability do appear to increase.

It is possible that biofeedback used to increase other sources of periodic variability in the cardiovascular system might also be explored. One such possibility is biofeedback targeting the vascular tone baroreflex, following research by Vaschillo and colleagues (Vaschillo et al., 2012; Vaschillo, Vaschillo, Pandina, & Bates, 2011).

Neurofeedback is another biofeedback modality that explicitly addresses oscillations by stimulating and exercising modulatory reflexes in the central nervous system. Alpha rhythm and theta: beta rhythm biofeedback have been used to enhance relaxation, memory consolidation, and attention, and appear helpful for such disorders as attention deficit hyperactivity disorder (Ramirez, Desantis, & Opler, 2001; Rossiter, 2004), and anxiety disorders (Moore, 2000), as well as intellectual performance enhancement (Angelakis et al., 2007; Gruzelier, Egner, & Vernon, 2006; Moore, 2000; Ramirez, Desantis, & Opler, 2001; Rossiter, 2004). This feedback may strengthen reflexes involved in conscious attention and memory consolidation, depending on the particular frequencies and brain regions that are trained. Pulsatile stimulation by electricity (Schindler, Elger, & Lehnertz, 2007) and light flicker have also been used (Jain, Woodruff, & Bissessar, 2001).

Chaotic and oscillatory properties of various psychophysiological measures suggest that there may be many more potential control reflexes that can be trained, perhaps one on each fractal plane. This raises a host of other possibilities for future biofeedback research. Future research may bear fruit by examining various sources of oscillation in the body and in behavior, to determine whether periodic stimulation of these oscillations may similarly exercise modulatory reflexes, and improve health. In particular, resonance and positive feedback loops could be considered, such as in heart rate variability biofeedback, wherein positive feedback can promote complexity and maintain oscillatory activity caused by negative feedback, helping individuals adapt to environmental demands.

Applications of systems theories to social and organizational structures, other than for Gottman's approach to marital interactions, have tended to be nonmathematical, focusing

primarily on norms, beliefs, and behaviors that preserve a particular pattern of social interaction. Time series analysis of these interaction patterns may yield important information that could be used as instruments for increasing social stability or social change.

Acknowledgments

This work was supported in part by Grant # 5R01HL089495-02 from the National Institutes of Health. For helpful comments in preparing this manuscript, the authors are indebted to Dr. Jaye Derrick of the State University of Buffalo and to the patient and thorough editorial reviewers for *Applied Psychophysiology and Biofeedback*.

REFERENCES

- Angelakis E, Stathopoulou S, Frymiare JL, Green DL, Lubar JF, Kounios J. EEG neurofeedback: a brief overview and an example of peak alpha frequency training for cognitive enhancement in the elderly. *Clinical Neuropsychologist*. 2007; 21:110–129. [PubMed: 17366280]
- Arzeno NM, Kearney MT, Eckberg DL, Nolan J, Poon C-S. Heart rate chaos as a mortality predictor in mild to moderate heart failure. *Conference Proceedings: Annual International Conference of the IEEE Engineering in Medicine & Biology Society*. 2007; 2007:5051–5054.
- Attneave, F. *Applications of Information Theory to Psychology: A Summary of Basic Concepts, Methods, and Results*. New York: Holt; 1959.
- Basar E. The theory of the whole-brain-work. *International Journal of Psychophysiology*. 2006; 60:133–138. [PubMed: 16563537]
- Basar E, Gönner A, Ungan P. Important relation between EEG and brain evoked potential. *Biological Cybernetics*. 1976; 25:27–40. [PubMed: 999965]
- Basar, E. *Brain function and oscillation*. New York: Springer; 1999.
- Bayram A, Bayraktogly Z, Karahan E, Erdogan B, Bilgic B, Özker M, Kasikci I, Duru AD, Ademoglu A, Oztürk C, Arikan K, Tarhan N, Demiralp T. Simultaneous EEG/fMRI analysis of the resonance phenomena in steady-state evoked responses. *Clinical EEG and Neuroscience*. 2011; 42:98–106. [PubMed: 21675599]
- Beckham AJ, Greene TB, Meltzer-Brody S. A pilot study of heart rate variability biofeedback therapy in the treatment of perinatal depression on a specialized perinatal psychiatry inpatient unit. *Archives of Women's Mental Health*. 2013; 16:59–65.
- Bernard-Weil E. A general model for the simulation of balance, imbalance and control by agonistic antagonistic biological couples. *Mathematical Modelling*. 1986; 7(9–12):1587–1600.
- Berntson GG, Bigger JT Jr, Eckberg DL, Grossman P, Kaufmann PG, Malik M, Nagaraja HN, Porges SW, Saul JP, Stone PH, van der Molen MW. Heart rate variability: origins, methods, and interpretive caveats. *Psychophysiology*. 1997; 34:623–648. [PubMed: 9401419]
- Berthoud, Hans-Rudolf. Homeostatic and non-homeostatic pathways involved in the control of food intake and energy balance. *Obesity*. 2006; Suppl 5(14):197S–200S. [PubMed: 17021366]
- Berthouze, Luc; James, Leon M.; Farmer, Simon F. Human EEG shows long-range temporal correlations of oscillation amplitude in Theta, Alpha and Beta bands across a wide age range. *Clinical Neurophysiology*. 2010; 121:1187–1197. [PubMed: 20346732]
- Bertram R, Sherman A, Satin LS. Electrical bursting, calcium oscillations, and synchronization of pancreatic islets. *Advances in Experimental Medicine & Biology*. 2010; 654:261–279. [PubMed: 20217502]
- Bob P. Chaos, brain and divided consciousness. *Acta Universitatis Carolinae - Medica - Monographia*. 2007; 153:9–80. [PubMed: 17867519]
- Bos EH, Hoenders R, de Jonge P. Wind direction and mental health: a time-series analysis of weather influences in a patient with anxiety disorder. *BMJ Case Reports*. 2012 pii: bcr2012006300.
- Bosl W, Tierney A, Tager-Flusberg H, Nelson C. EEG complexity as a biomarker for autism spectrum disorder risk. *BMC Medicine*. 2011; 9:18. [PubMed: 21342500]
- Brown CM, Hecht MJ, Weih A, Neundorfer B, Hilz MJ. Effects of age on the cardiac and vascular limbs of the arterial baroreflex. *European Journal of Clinical Investigation*. 2003; 33:10–16. [PubMed: 12492447]

- Buzsaki, Gyorgy. Rhythms of the brain. New York, NY: Oxford University Press; US; 2006.
- Cancian FM, Gordon SL. Changing emotion norms in marriage: Love and anger in U.S. women's magazines since 1900. *Gender & Society*. 1988; 2:308–342.
- Chemtob CM, Roitblat H, Hamada RS, Carlson JG, Twentyman CT. A cognitive action theory of Post-Traumatic Stress Disorder. *Journal of Anxiety Disorders*. 1988; 2:253–275.
- Chen J-L, Yeh D-P, Lee J-P, Chen C-Y, Huang C-Y, Lee S-D, Chen CC, Kuo TB, Kao CL, Kuo C-H. Parasympathetic nervous activity mirrors recovery status in weightlifting performance after training. *Journal of Strength & Conditioning Research*. 2011; 25:1546–1552. [PubMed: 21273908]
- Cinquin O, Demongeot J. Roles of positive and negative feedback in biological systems. *Comptes Rendus Biologies*. 2002; 325:1085–1095. [PubMed: 12506722]
- Crunelli, Vincenzo; Hughes, Stuart W. The slow (<1 Hz) rhythm of non-REM sleep: a dialogue between three cardinal oscillators. *Nature Neuroscience*. 2010; 13:9–17.
- Cugini P, Ferrari P, De Rosa R, Caliumi C, Delfini E, Colotto M, Fontana S, Mandolini C, Manetti L, Letizia C. Severity of human hypertension in relation to the age in which high blood pressure makes its presumptive appearance. *Clinica Terapeutica*. 2003; 154:21–26. [PubMed: 12854280]
- Dworkin, Barry R. Learning and physiological regulation. Chicago: University of Chicago Press; 1993.
- Elbert T, Ray WJ, Kowalik ZJ, Skinner JE, Graf KE, Birbaumer N. Chaos and physiology: deterministic chaos in excitable cell assemblies. *Physiological Reviews*. 1994; 74:1–47. [PubMed: 8295931]
- Fedotchev AI, Bondar AT, Konovalov VF. Stability of resonance EEG reactions to flickering light in humans. *International Journal of Psychophysiology*. 1990; 9:189–193. [PubMed: 2228753]
- Feigl EO. Neural control of coronary blood flow. *Journal of Vascular Research*. 1998; 35:85–92. [PubMed: 9588871]
- Fogel, Stuart M.; Smith, Carlyle T. The function of the sleep spindle: a physiological index of intelligence and a mechanism for sleep-dependent memory consolidation. *Neuroscience & Biobehavioral Reviews*. 2011; 35:1154–1165. [PubMed: 21167865]
- Fry, Steven N.; Rohrseitz, Nicola; Straw, Andrew D.; Dickinson, Michael H. TrackFly: virtual reality for a behavioral system analysis in free-flying fruit flies. *Journal of Neuroscience Methods*. 2008; 171:110–117. [PubMed: 18405978]
- Garau C, Aparicio S, Rial RV, Nicolau MC, Esteban S. Age related changes in the activity-rest circadian rhythms and c-fos expression of ring doves with aging. Effects of tryptophan intake. *Experimental Gerontology*. 2006; 41:430–438. [PubMed: 16564149]
- Gellhorn E. Further studies on the physiology and pathophysiology of the tuning of the central nervous system. *Psychosomatics: Journal of Consultation Liaison Psychiatry*. 1969; 10:94–104.
- Gellhorn E. The emotions and the ergotropic and trophotropic systems. *Psychologische Forschung*. 1970; 34:48–94. [PubMed: 4929035]
- Gengerelli JA. Wave coherence in the human EEG. *Journal of Psychology*. 1978; 99(2d Half):203–223. [PubMed: 671377]
- Gleik, J. Chaos: Making a new science. New York: Penguin Books; 1987.
- Goldberger AL, Peng CK, Lipsitz LA. What is physiologic complexity and how does it change with aging and disease? *Neurobiology of aging*. 2002; 23:23–26. [PubMed: 11755014]
- Gottesmann C. Neurophysiological support of consciousness during waking and sleep. *Progress in Neurobiology*. 1999; 59:469–508. [PubMed: 10515665]
- Gottman, JM.; Murray, JD.; Swanson, CC.; Tyson, R.; Swanson, KR. The mathematics of marriage: Dynamic nonlinear models. Cambridge, MA: The MIT Press; 2002.
- Gottschalk A, Bauer MS, Whybrow PC. Evidence of chaotic mood variation in bipolar disorder. *Archives of general psychiatry*. 1995; 52:947. [PubMed: 7487343]
- Grodins, FS. Control theory and biological systems. New York: Columbia University Press; 1963.
- Gruzelier J, Egner T, Vernon D. Validating the efficacy of neurofeedback for optimizing performance. *Progress in Brain Research*. 2006; 159:421–431. [PubMed: 17071246]
- Guastello, SJ.; Koopmans, M.; Pincus, D., editors. Chaos and complexity in psychology. New York: Cambridge University Press; 2009.

- Haffen E, Sechter D. Disruption of biological rhythms and mood disorders. *Encephale*. 2006; 32(5 Pt 2):S795–S801. [PubMed: 17119474]
- Hallman DM, Olsson EM, von Scheele B, Melin L, Lyskov E. Effects of heart rate variability biofeedback in subjects with stress-related chronic neck pain: a pilot study. *Applied Psychophysiology & Biofeedback*. 2011; 36:71–80. [PubMed: 21365308]
- Henriques G, Keffer S, Abrahamson C, Horst SJ. Exploring the effectiveness of a computer-based heart rate variability biofeedback program in reducing anxiety in college students. *Applied Psychophysiology & Biofeedback*. 2011; 36:101–112. [PubMed: 21533678]
- Herrmann CS. Human EEG responses to 1–100 Hz flicker: resonance phenomena in visual cortex and their potential correlation to cognitive phenomena. *Experimental Brain Research*. 2001; 137(3–4): 346–353.
- Ho Y-L, Lin C, Lin Y-H, Lo M-T. The prognostic value of non-linear analysis of heart rate variability in patients with congestive heart failure--a pilot study of multiscale entropy. *PLoS ONE*. 2011; 6:e18699. [PubMed: 21533258]
- Hyndman BW. The role of rhythms in homeostasis. *Biological Cybernetics*. 1974; 15:227–236.
- Isler Y, Kuntalp M. Combining classical HRV indices with wavelet entropy measures improves to performance in diagnosing congestive heart failure. *Computers in Biology & Medicine*. 2007; 37:1502–1510. [PubMed: 17359959]
- Jacobs GD. The physiology of mind-body interactions: the stress response and the relaxation response. *Journal of Alternative & Complementary Medicine*. 2001; Suppl 1(7):S83–S92.
- Juster RP, McEwen BS, Lupien SJ. Allostatic load biomarkers of chronic stress and impact on health and cognition. *Neuroscience and biobehavioral reviews*. 2010; 35:2–16. [PubMed: 19822172]
- Kaneko W, Phillips E, Riley E, Ehlers C. EEG findings in fetal alcohol syndrome and Down syndrome children. *Electroencephalography and clinical neurophysiology*. 1996; 98:20–28. [PubMed: 8689990]
- Karavidas MK, Lehrer PM, Vaschillo E, Vaschillo B, Marin H, Buyske S, Malinovsky I, Radvanski D, Hassett A. Preliminary results of an open label study of heart rate variability biofeedback for the treatment of major depression. *Applied Psychophysiology & Biofeedback*. 2007; 32:19–30. [PubMed: 17333315]
- Kasevich RS, LaBerge D. Theory of electric resonance in the neocortical apical dendrite. *PLoS ONE*. 2011; 6:e23412. [PubMed: 21853129]
- Kellert, Stephen H. *In the Wake of Chaos: Unpredictable Order in Dynamical Systems*. Chicago: University of Chicago Press; 1993.
- Klapp OE. Opening and closing in open systems. *Behavioral Science*. 1975; 20:251–257.
- Klevecz RR, Li CM, Marcus I, Frankel PH. Collective behavior in gene regulation: the cell is an oscillator, the cell cycle a developmental process. *FEBS Journal*. 2008; 275:2372–2384. [PubMed: 18410382]
- Klimesch W. Memory processes, brain oscillations and EEG synchronization. *International Journal of Psychophysiology*. 1996; 24(1–2):61–100. [PubMed: 8978436]
- Klimesch W, Schack B, Sauseng P. The functional significance of theta and upper alpha oscillations. *Experimental Psychology*. 2005; 52:99–108. [PubMed: 15850157]
- Knyazev GG. Motivation, emotion, and their inhibitory control mirrored in brain oscillations. *Neuroscience & Biobehavioral Reviews*. 2007; 31:37–395.
- Krantz DS, Contrada RJ, LaRiccia PJ, Anderson JR, Durel LA, Dembroski TM, Weiss T. Effects of beta-adrenergic stimulation and blockade on cardiovascular reactivity, affect, and type A behavior. *Psychosomatic Medicine*. 1987; 49:146–158. [PubMed: 3575603]
- Kurbel S. A phase plane graph based model of the ovulatory cycle lacking the "positive feedback" phenomenon. *Theoretical Biology and Medical Modelling*. 2012; 9:35. [PubMed: 22870942]
- Lehrer, PM. Principles and practice of stress management. 3rd ed. New York, NY: Guilford Press; US; 2007. Biofeedback training to increase heart rate variability; p. 227–248.
- Lehrer PM, Karavidas MK, Lu S-E, Coyle SM, Oikawa LO, Macor M, Calvano SE, Lowry SF. Voluntarily produced increases in heart rate variability modulate autonomic effects of endotoxin induced systemic inflammation: An exploratory study. *Applied Psychophysiology and Biofeedback*. 2010; 35:303–315. [PubMed: 20635134]

- Lehrer PM, Vaschillo EG, Vaschillo B, Lu S-E, Eckberg DL, Edelberg R, Shih WJ, Lin Y, Kausela TA, Tahvanainen KU, Hamer R. Heart rate variability biofeedback increases baroreflex gain and peak expiratory flow. *Psychosomatic Medicine*. 2003; 65:796–805. [PubMed: 14508023]
- Lehrer PM, Vaschillo EG, Vaschillo B, Lu S-E, Scardella A, Siddique M, Habib RH. Biofeedback treatment for asthma. *Chest*. 2004; 126:352–361. [PubMed: 15302717]
- Lempel A, Ziv J. On the Complexity of Finite Sequences. *IEEE Transactions on Information Theory*. 1976; 22:75–81.
- Lin G, Xiang Q, Fu X, Wang S, Wang S, Chen S, Shao L, Zhao Y, Wang T. Heart rate variability biofeedback decreases blood pressure in prehypertensive subjects by improving autonomic function and baroreflex. *Journal of Alternative & Complementary Medicine*. 2012; 18:143–152.
- Liu, Chengyu; Liu, Changchun; Shao, P.; Li, L.; Sun, X.; Wang, X.; Liu, F. Comparison of different threshold values r for approximate entropy: application to investigate the heart rate variability between heart failure and healthy control groups. *Physiological Measurement*. 2011; 32:167–180. [PubMed: 21178247]
- Mandelbrot, BB. *The fractal geometry of nature*. New York: Macmillan; 1983.
- Martinez L, Perez T, Mirasso CR, Manjarrez E. Stochastic resonance in the motor system: effects of noise on the monosynaptic reflex pathway of the cat spinal cord. *Journal of Neurophysiology*. 2007; 97:4007–4016. [PubMed: 17428901]
- Mason OJ, Brady F. The psychotomimetic effects of short-term sensory deprivation. *Journal of Nervous & Mental Disease*. 2009; 197:783–785. [PubMed: 19829208]
- Maurer C, Mergner T, Peterka RJ. Abnormal resonance behavior of the postural control loop in Parkinson's disease. *Experimental Brain Research*. 2004; 157:369–376.
- Mayhew, DR. *Electoral Realignments: A Critique of an American Genre*. New Haven, CT: Yale University Press; 2002.
- McEwen BS. Protection and damage from acute and chronic stress: allostasis and allostatic overload and relevance to the pathophysiology of psychiatric disorders. *Annals of the New York Academy of Sciences*. 2004; 1032:1–7. [PubMed: 15677391]
- Mindell, D.; Segal, J.; Gerovitch, S. Cybernetics and information theory in the United States, France and the Soviet Union. In: Walker, M., editor. *Science and Ideology: A Comparative History*. London: Routledge; 2003. p. 66-95.
- Montano N, Porta A, Cogliati C, Costantino G, Tobaldini E, Casali KR, Iellamo F. Heart rate variability explored in the frequency domain: a tool to investigate the link between heart and behavior. *Neuroscience & Biobehavioral Reviews*. 2009; 33:71–80. [PubMed: 18706440]
- Moore NC. A review of EEG biofeedback treatment of anxiety disorders. *Clinical Electroencephalography*. 2000; 31:1–6. [PubMed: 10638346]
- Nandrino JL, Pezard L, Martinerie J, el Massioui F, Renault B, Jouvent R, Allilaire JF, Widlocher D. Decrease of complexity in EEG as a symptom of depression. *NeuroReport*. 1994; 5:528–530. [PubMed: 8003689]
- Nishi A, Bibb JA, Snyder GL, Higashi H, Nairn AC, Greengard P. Amplification of dopaminergic signaling by a positive feedback loop. *Proceedings of the National Academy of Sciences of the United States of America*. 2000; 97:12840–12845. [PubMed: 11050161]
- Nolan RP, Floras JS, Harvey PJ, Kamath MV, Picton PE, Chessex C, Hiscock N, Powell J, Catt M, Hendrickx H, Talbot D, Chen MH. Behavioral neurocardiac training in hypertension: a randomized, controlled trial. *Hypertension*. 2010; 55:1033–1039. [PubMed: 20194302]
- Norris PR, Anderson SM, Jenkins JM, Williams AE, Morris JA Jr. Heart rate multiscale entropy at three hours predicts hospital mortality in 3,154 trauma patients. *Shock*. 2008; 30:17–22. [PubMed: 18323736]
- Norris PR, Stein PK, Morris JA Jr. Reduced heart rate multiscale entropy predicts death in critical illness: a study of physiologic complexity in 285 trauma patients. *Journal of Critical Care*. 2008; 23:399–405. [PubMed: 18725047]
- Onton J, Westerfield M, Townsend J, Makeig S. Imaging human EEG dynamics using independent component analysis. *Neuroscience & Biobehavioral Reviews*. 2006; 30:808–822. [PubMed: 16904745]

- Orme-Johnson DW, Haynes CT. EEG phase coherence, pure consciousness, creativity, and TM--Sidhi experiences. *International Journal of Neuroscience*. 1981; 13:211–217. [PubMed: 7026478]
- Panzeri S, Brunel N, Logothetis NK, Kayser C. Sensory neural codes using multiplexed temporal scales. *Trends in Neurosciences*. 2010; 33:111–120. [PubMed: 20045201]
- Patron E, Benvenuti SM, Favretto G, Valfre C, Bonfa C, Gasparotto R, Palomba D. Biofeedback assisted control of respiratory sinus arrhythmia as a biobehavioral intervention for depressive symptoms in patients after cardiac surgery: a preliminary study. *Applied Psychophysiology & Biofeedback*. in press
- Paul, Maman; Garg, Kanupriya. The effect of heart rate variability biofeedback on performance psychology of basketball players. *Applied Psychophysiology and Biofeedback*. 2012; 37:131–144. doi: <http://dx.doi.org/10.1007/s10484-012-9185-2>. [PubMed: 22402913]
- Paulus, Martin P.; Braff, David L. Chaos and schizophrenia: does the method fit the madness? *Biological Psychiatry*. 2003; 53:3–11. [PubMed: 12513940]
- Paulus MP, Geyer MA, Braff DL. Use of methods from chaos theory to quantify a fundamental dysfunction in the behavioral organization of schizophrenic patients. *American Journal of Psychiatry*. 1996; 153:714–717. [PubMed: 8615422]
- Pezard L, Nandrino JL, Renault B, el Massioui F, Allilaire JF, Muller J, Varela FJ, Martinerie J. Depression as a dynamical disease. *Biological Psychiatry*. 1996; 39:991–999. [PubMed: 8780833]
- Pfurtscheller G, Lopes da Silva FH. Event-related EEG/MEG synchronization and desynchronization: basic principles. *Clinical Neurophysiology*. 1999; 110:1842–1857. [PubMed: 10576479]
- Pigolott, iS; Krishna, S.; Jensen, MH. Oscillation patterns in negative feedback loops. *Proceedings of the National Academy of Sciences of the United States of America*. 2007; 104:6533–6537. [PubMed: 17412833]
- Pincus SM. Approximate entropy as a measure of system complexity. *Proceedings of the National Academy of Sciences*. 1991; 88:2297–2301.
- Pincus, SM. Applications of nonlinear dynamics to developmental process modeling. Mahwah, NJ: Lawrence Erlbaum Associates Publishers; US; 1998. Approximate entropy (ApEn) as a regularity measure; p. 243-268.
- Plahte E, Mestl T, Omholt SW. Feedback loops, stability, and multistationarity in dynamical systems. *Journal of Biological Systems*. 1995; 3:409–413.
- Polonsky KS, Sturis J, Van Cauter E. Temporal profiles and clinical significance of pulsatile insulin secretion. *Hormone Research in Paediatrics*. 2000; 49:178–184.
- Pomerening JR, Kim SY, Ferrell JE Jr. Systems-level dissection of the cell-cycle oscillator: bypassing positive feedback produces damped oscillations. *Cell*. 2005; 122:565–578. [PubMed: 16122424]
- Prochazka A, Gillard D, Bennett DJ. Implications of positive feedback in the control of movement. *Journal of Neurophysiology*. 1997; 77:3237–3251. [PubMed: 9212271]
- Putman P, van Peer J, Maimari I, van der Werff S. EEG theta/beta ratio in relation to fear-modulated response-inhibition, attentional control, and affective traits. *Biological Psychology*. 2010; 83:73–78. [PubMed: 19897008]
- Rahman F, Pechnik S, Gross D, Sewell L, Goldstein DS. Low frequency power of heart rate variability reflects baroreflex function, not cardiac sympathetic innervation. *Clinical Autonomic Research*. 2011; 21:133–141. [PubMed: 21279414]
- Ramirez PM, Desantis D, Opler LA. EEG biofeedback treatment of ADD. A viable alternative to traditional medical intervention? *Annals of the New York Academy of Sciences*. 2001; 931:342–358. [PubMed: 11462752]
- Reineke, Anke. The effects of heart rate variability biofeedback in reducing blood pressure for the treatment of essential hypertension. *Dissertation Abstracts International: Section B: The Sciences and Engineering*. 2008; 68(7-B):4880.
- Roschke J, Fell J, Beckmann P. Nonlinear analysis of sleep EEG data in schizophrenia: calculation of the principal Lyapunov exponent. *Psychiatry Research*. 1995; 56:257–269. [PubMed: 7568548]
- Rossiter, Thomas. The effectiveness of neurofeedback and stimulant drugs in treating AD/HD: Part I. Review of methodological issues. *Applied Psychophysiology & Biofeedback*. 2004; 29:95–112. [PubMed: 15208973]

- Rossler OE, Rossler R. Chaos in physiology. *Integrative Physiological & Behavioral Science*. 1994; 29:328–333. [PubMed: 7811652]
- Schwartz GE. A systems analysis of psychobiology and behavior therapy. Implications for behavioral medicine. *Psychotherapy & Psychosomatics*. 1981; 36(3–4):159–184. [PubMed: 7048388]
- Schwartz GE, Shapiro AP, Redmond DP, Ferguson DC, Ragland DR, Weiss SM. Behavioral medicine approaches to hypertension: an integrative analysis of theory and research. *Journal of Behavioral Medicine*. 1979; 2:311–363. [PubMed: 398408]
- Shannon CE. A mathematical theory of communication. *Bell Systems Technology Journal*. 1948; 27:379–423.
- Shannon, CE.; Weaver, W. *The Mathematical Theory of Communication*. Champaign, IL: Univ of Illinois; 1949.
- Shenefelt PD. Relaxation strategies for patients during dermatologic surgery. *Journal of Drugs in Dermatology: JDD*. 2010; 9:795–799. [PubMed: 20677535]
- Siebert, WMcC. *Circuits, signals, and systems*. Cambridge, MA: MIT Press; 1986.
- Siepmann M, Aykac V, Unterdorfer J, Petrowski K, Mueck-Weymann M. A pilot study on the effects of heart rate variability biofeedback in patients with depression and in healthy subjects. *Applied Psychophysiology and Biofeedback*. 2008; 33:195–201. [PubMed: 18807175]
- Sohn H, Kim I, Lee W, Peterson BS, Hong H, Chae J-H, Hong S, Jeong J. Linear and non-linear EEG analysis of adolescents with attention-deficit/hyperactivity disorder during a cognitive task. *Clinical Neurophysiology*. 2010; 121:1863–1870. [PubMed: 20659814]
- Sowder E, Gevirtz R, Shapiro W, Ebert C. Restoration of vagal tone: A possible mechanism for functional abdominal pain. *Applied Psychophysiology and Biofeedback*. 2010; 35:199–206. [PubMed: 20229150]
- Spiegler A, Knösche TR, Schwab K, Haueisen J, Atay FM. Modeling Brain Resonance Phenomena Using a Neural Mass Model. *PLOS Computational Biology*. 2011; 7
- Strine GN. Self-reports of pain reduction through paced respiration and heart rate variability biofeedback with nursing home residents. *Dissertation Abstracts International: Section B: The Sciences and Engineering*. 2004; 65(5-B):2685.
- Thayer JF, Lane RD. A model of neurovisceral integration in emotion regulation and dysregulation. *Journal of Affective Disorders*. 2000; 61:201–216. [PubMed: 11163422]
- Turner S, Mittermaier C, Ehrenberger K. Change of complexity patterns in human posture during aging. *Audiology & Neuro-Otology*. 2002; 7:240–248. [PubMed: 12097723]
- Timofeev I, Chauvette S. Thalamocortical oscillations: local control of EEG slow waves. *Current Topics in Medicinal Chemistry*. 2011; 11:2457–2471. [PubMed: 21906018]
- Tognoli E, Kelso JA. Brain coordination dynamics: true and false faces of phase synchrony and metastability. *Progress in Neurobiology*. 2009; 87:31–40. [PubMed: 18938209]
- Tolic I, Mosekilde E, Sturis J. Modeling the insulin-glucose feedback system: the significance of pulsatile insulin secretion. *Journal of theoretical biology*. 2000; 207:361–375. [PubMed: 11082306]
- Tsai TY-C, Choi YS, Ma W, Pomerening JR, Tang C, Ferrell JE Jr. Robust, tunable biological oscillations from interlinked positive and negative feedback loops. *Science*. 2008; 321(5885): 126–129. [PubMed: 18599789]
- Tschacher W, Scheier C, Hashimoto Y. Dynamical analysis of schizophrenia courses. *Biological Psychiatry*. 1997; 41:428–437. [PubMed: 9034537]
- Vaschillo EG, Lehrer PM, Rische N, Konstantinov M. Heart rate variability biofeedback as a method for assessing baroreflex function: A preliminary study of resonance in the cardiovascular system. *Applied Psychophysiology and Biofeedback*. 2002; 27:1–27. [PubMed: 12001882]
- Vaschillo EG, Vaschillo B, Buckman JF, Pandina RJ, Bates ME. Measurement of vascular tone and stroke volume baroreflex gain. *Psychophysiology*. 2012; 49:193–197. [PubMed: 22092290]
- Vaschillo EG, Vaschillo B, Lehrer PM. Characteristics of resonance in heart rate variability stimulated by biofeedback. *Applied Psychophysiology and Biofeedback*. 2006; 31:129–142. [PubMed: 16838124]

- Vaschillo EG, Vaschillo B, Pandina Robert J, Bates ME. Resonances in the cardiovascular system caused by rhythmical muscle tension. *Psychophysiology*. 2011; 48:927–936. [PubMed: 21143610]
- Ward LM, MacLean SE, Kirschner A. Stochastic resonance modulates neural synchronization within and between cortical sources. *PLoS ONE*. 2010; 5:e14371. [PubMed: 21179552]
- Weaver RL, Lobkis OI. On the linewidth of the ultrasonic Larsen effect in a reverberant body. *The Journal of the Acoustical Society of America*. 2006; 120:102–109.
- Weick, KE. *Making sense of the organization: the impermanent organization*. Chichester, Sussex, UK: John Wiley and Sons; 2009.
- Weinert D, Waterhouse J. The circadian rhythm of core temperature: effects of physical activity and aging. *Physiology & Behavior*. 2007; 90(2–3):246–256. [PubMed: 17069866]
- Weiss JN, Garfinkel A, Speno ML, Ditto WL. Chaos and chaos. *Journal of Clinical Investigation*. 1994; 93:1355–1360. [PubMed: 8163640]
- Wiener, N. *Cybernetics; or control and communication in the animal and the machine*. Oxford: John Wiley; 1948.
- Wiener N. *Cybernetics* (2nd ed). 1961; 212
- Wiesenfeld K, Moss F. Stochastic resonance and the benefits of noise: from ice ages to crayfish and squids. *Nature*. 1995; 373(6509):33–36. [PubMed: 7800036]
- Winterer G, Ziller M, Dorn H, Frick K, Mulert C, Dahhan N, Herrmann WM, Coppola R. Cortical activation, signal-to-noise ratio and stochastic resonance during information processing in man. *Clinical Neurophysiology*. 1999; 110:1193–1203. [PubMed: 10423185]
- Yerkes RM, Dodson JD. The relation of strength of stimulus to rapidity of habit-formation. *Journal of Comparative Neurology and Psychology*. 1908; 18:459–482.
- Yetwin, Alexis Kant. Heart rate variability biofeedback therapy for children and adolescents with chronic pain. *Dissertation Abstracts International: Section B: The Sciences and Engineering*. 2012; 72(12-B):7704.
- Yockey, Hubert P. *Information Theory, Evolution, and The Origin of Life*. New York: Cambridge University Press; 2005.
- Zarjam, P.; Epps, J.; Chen, F. Spectral EEG features for evaluating cognitive load; Paper presented at the 334d Annual International Conference of the IEEE EMBS; August 30-September 3, 2011; Boston, MA. 2011.