

Thomas Jefferson University Jefferson Digital Commons

Department of Family & Community Medicine **Faculty Papers**

Department of Family & Community Medicine

February 2007

Toward a biopsychosocial understanding of the patient-physician relationship: an emerging dialogue

Herbert M. Adler Thomas Jefferson University, herbert.adler187@gmail.com

Let us know how access to this document benefits you

Follow this and additional works at: http://jdc.jefferson.edu/fmfp



Overage of Part of the Family Medicine Commons, and the Translational Medical Research Commons

Recommended Citation

Adler, Herbert M., "Toward a biopsychosocial understanding of the patient-physician relationship: an emerging dialogue" (2007). Department of Family & Community Medicine Faculty Papers. Paper 14. http://jdc.jefferson.edu/fmfp/14

This Article is brought to you for free and open access by the Jefferson Digital Commons. The Jefferson Digital Commons is a service of Thomas Jefferson University's Center for Teaching and Learning (CTL). The Commons is a showcase for Jefferson books and journals, peer-reviewed scholarly publications, unique historical collections from the University archives, and teaching tools. The Jefferson Digital Commons allows researchers and interested readers anywhere in the world to learn about and keep up to date with Jefferson scholarship. This article has been accepted for inclusion in Department of Family & Community Medicine Faculty Papers by an authorized administrator of the Jefferson Digital Commons. For more information, please contact: JeffersonDigitalCommons@jefferson.edu.

Biopsychosocial Patient-Physician Relationship

TITLE

Toward a Biopsychosocial Understanding of the Patient-Physician Relationship: An **Emerging Dialogue**

AUTHOR

Herbert M. Adler, MD, PhD

Clinical Professor Department of Family Medicine, Jefferson Medical College Clinical Professor Department of Psychiatry, Jefferson Medical College 1015 Chestnut Street; Suite 1000

Philadelphia, PA 19107

Phone/Fax: 215-925-3385 e-mail: herbert.adler@earthlink.net

Abstract: 79

References: 103

Key Words: patient-physician relationship, biopsychosocial, interpersonal neurobiology **Acknowledgments**: I acknowledge with gratitude the critical readings provided by Richard L. Epstein, MD, MPH, Fred W. Markham, Jr., MD, Robert L. Perkel, MD, and Howard K. Rabinowitz, MD, the editorial assistance of Jennifer Kearney-Strouse, ELS, and Richard M. Frankel, PhD for the emergent dialogue that contributed to this paper.

Conflict of Interest: None

Biopsychosocial Patient-Physician Relationshi
Toward a Biopsychosocial Understanding of the Patient-Physician Relationship: An Emerging Dialogue
Differging Dialogue

Abstract

Complexity theory has been used to view the patient-physician relationship as constituted by complex responsive processes of relating. It describes an emergent, psychosocial relational process through which patients and physicians continually and reciprocally influence each other's behavior and experience. Since psychosocial responses are necessarily biopsychosocial responses, patients and physicians must likewise be influencing each other's psychobiology. This mutual influence may be subjectively experienced as empathy, and may be skillfully employed by the clinician to directly improve the patient's psychobiology.

"By far the most frequently used drug in general practice was the doctor himself."(1)

"The secret of the care of the patient is in caring for the patient."(2)

Introduction

Suchman(3) has argued persuasively for viewing the patient-physician relationship as constituted by complex responsive psychosocial processes of relating. Accordingly, the interaction of physician and patient can be viewed as an emergent, self-organizing process. It is established and maintained by reciprocal, iterative psychosocial responses through which each mutually influences and co-regulates the other's interdependent behavior and personal experience. Psychosocial responses have biological concomitants(4) likewise making patient-physician interactions emergent, self-organizing feedback loops comprised of mutually regulatory biopsychosocial responses. This may be labeled a complex biopsychosocial relational process. In the interest of brevity, I will refer to this throughout the paper as the biopsychosocial relational process or just the relational process.

This approach to the patient-physician relationship shifts our focus from the relationship as a context for the delivery of medical treatment to the relationship itself as a medical treatment. It also redirects our view of the function of the physician—from a *provider* of treatment to a *co-participant* in treatment, with emergent consequences for both patient and physician. Three features are highlighted:

As co-creators of a complex self-organizing relationship, patients and physicians
are engaged in a moment-to-moment mutual regulation of each other's
biopsychosocial states.

- 2. The introduction by either patient or physician of even small changes in their interactive process can lead to large changes in their biopsychosocial outcomes.
- The emergence of empathy in the patient and physician may be viewed as a biopsychosocial relational process indicator, and may also serve as a guide toward desired outcomes.

Empirical Evidence

Overview

Empirical research has demonstrated that contingent interpersonal responses are accompanied by contingent neurobiological responses. Depending on the social context, such responses have been variously labeled *sociophysiology* between therapists and patients, (5, 6) as well as among nonhuman mammals; (7) *interpersonal neurobiology* in child development; (8) *affect attunement* between caregivers and infants; (9) and *physiological linkage* between empathic spouses. (10) More recently, neuroscientists have discovered a *mirror neuronal system* that contributes to this attuned responsiveness, with special relevance to empathy. (11-13)

Sociophysiology

Starting in the mid-1950s with investigations of the psychiatric interview, researchers described an "interpersonal physiology," which referred to a correlation of selected physiologic indicators of autonomic activity—heart rate,(5) heart lability, skin temperature,(14) and muscle tension(15)—that varied together between psychotherapists and patients. It was speculated that this similarity of patterning was a "physiological identification" between therapist and patient, and might turn out to be an objective

measurement of rapport.(14) These findings were confirmed by other researchers,(6) and it was also demonstrated that the physiologic correlation was the result of empathy rather than a common reaction to the same events.(16, 17) Aside from studies of the "physiological linkage"(18) between empathic spouses, few other clinical explorations of sociophysiology were performed.(10)

Animal ethologists applied sociophysiologic research strategies to look at how mutually-regulatory physiologic feedback loops establish and maintain nonhuman mammalian social organizations.(7, 19, 20) This was demonstrated at a number of levels of mammalian social organization: the mother-offspring bond,(21) conspecific (peer) relationships,(22) the adult pair-bond,(23) hierarchical relationships,(24, 25) sexual development,(26, 27) and sexual reproduction.(28, 29)

Subsequently, Gardner used the term *sociophysiology* to refer to the hypothesis that current psychopathology is a consequence of evolutionarily selected characteristics of brain physiology.(30) My use of the term preserves its original meaning as an interpersonal physiological engagement, occurring in real time and having continuous here-and-now physiologic consequences. As applied to the patient-physician relationship, sociophysiology encompasses two overlapping processes. First, intrapsychically, anticipated, planned, and remembered social experiences are inseparable from their concomitant physiology. Second, interpersonally, social relationships influence physiology and vice versa. For humans, these two processes may be hardly distinguishable because even when alone, people are usually in the company of imagined others,(31) and their physiology will reflect this. So, even between medical encounters, and years later, the biopsychosocial relational process can continue.

Interpersonal Neurobiology

The unfolding neural circuitry of the developing brain is configured by social interaction, variously described as interpersonal neurobiology(32) and the social construction of the human brain.(33) Throughout the lifespan, social interaction continues to modify neural structures(34, 35) and maintain the integrated functioning of neural circuits.(4) Consistent with complexity theory, reciprocal, modifiable neurobiological and neuroendocrine patterns of response affect and are affected by social attachment.(36-38) The propensity for the kind of self-organizing physiological attunement that regulates the mother-infant relationship may continue into adult interpersonal relationships, and may serve a similar physiologic regulatory function.(39, 40) The physiologic consequences of bereavement in adults are very similar to those of maternal separation in infants, and may be partially accounted for by the loss of an external physiologic regulator.(41)

Affect Attunement—The Regulation of Physiology and Attachment

Studies by infancy researchers have demonstrated the precisely calibrated feedback loop through which caregivers modulate infants' physiological responses by a nuanced combination of stimulation and soothing.(42-46) Stern has labeled this interactive regulatory process "affect attunement."(9) Through this self-organizing developmental process,(47) caregiver and infant co-operate the neurobiological responses that establish and maintain their attachment.

This "dance of attunement" (48) creates a secure, affectional bond (49) that synchronizes the level of autonomic arousal in both infant and caregiver, (48) is usually experienced by the caregiver as deeply satisfying, and tends to have a calming effect on the infant. To this end, caregivers report being guided in their attuned responses by an

empathic feeling with the infant.(50) Because much of this mutual responsiveness occurs too instantaneously to be under conscious control, it had been anticipated that innate imitative neural circuits would be discovered.(51) Neuroscience has now provided such a candidate neural mechanism.(11, 52)

The Mirror Neuronal System

Neuroscience research, first in monkeys(53, 54) and then in humans,(55, 56) has discovered a mirror neuronal system that can account for a cognitively unmediated responsive feedback loop underlying interpersonal communication. Mirror neurons discharge when a specific motor action is performed and when an individual observes another individual performing a similar motor action.(57, 58) Because the mirror neuronal system in both humans and monkeys(59, 60) is connected to parts of the brain that are critical for the recognition of facial expressions and emotional behaviors,(12, 59) the observation of emotions can influence the emotional experience of the observer. In this way, the mirror neuronal system may provide a neurobiological grounding for interpersonal empathy.(12, 13, 61-63)

Empathy as Interpersonal Neurobiology

Further support for the view that empathy is a neurobiological response has been provided by the use of positron emission tomography (PET) to demonstrate that accurate empathy of distress is correlated with the activation of specific neural networks.(64) Accordingly, empathy can be thought of as the neurobiological experience of *what* we know and *how* we know it.(17, 61, 65)

In a study of empathy in marital couples, those spouses who exhibited the most accurate empathy regarding each other's negative feelings had the most synchronous patterns of autonomic activation—described as a "physiological linkage." (17) In another

study, the degree of physiological synchrony between spouses on four measures (heart rate, pulse transmission time to finger, skin conductance level, and general somatic activity) was shown to correlate with both their emotional synchrony and marital satisfaction.(66)

Summary

Both complexity theory and empirical evidence support the proposition that the empathy in the patient-physician relationship is constituted by the reciprocal, emergent biopsychosocial responses of each party.

The Clinical Application of Empathy in the Patient-Physician Relationship

Empathy includes both the subjective *perception* of attuned interpersonal neurobiology and the moment-to-moment *process* of this attunement—the more accurate the reciprocal responses, the more synchronous the attunement. Importantly, even small changes introduced by either patient or physician can cascade into large neurobiological changes. Herein lies the therapeutic potential of *clinical empathy*, defined as the physician's use of the empathic process to directly affect the patient's psychobiology. Whether clinical empathy is conceptualized as a primarily cognitive process(67) that makes patients feel understood or as a primarily affective process(68) that makes them "feel felt,"(32) it is an emergent neurobiological process.

Clinical Empathy as a Clinical Procedure

While biopsychosocial responsiveness between patients and physicians is reciprocal and mutual, it is not symmetrical because patients and society grant clinicians the responsibility to focus attention and treat biological and psychological aspects of a patient's disease. Treatment includes: prescribing medication, providing information,

and performing clinical procedures. The physician's use of empathy, warrants consideration as a clinical procedure because it uses "emotional resonance" (69) to achieve skilled "communicative attunement" (70) that produces a neurobiological intervention.

Three additional features of clinical empathy support its consideration as a clinical procedure: 1) It has a medical indication; 2) It is a skilled, interpersonal performance requiring "emotional labor";(71) and 3) It attempts to achieve a specific outcome—an improvement in the patient's psychobiology.

Beginning with the indication, the distress of sickness can result in both an activation of the hypothalamic-pituitary-adrenal axis and an ensuing need to seek psychobiological relief through the formation of a secure attachment bond with a caregiver. (72) At this vulnerable point in the patient's life, the effect of clinical empathy on the patient's psychobiology is likely to be enhanced. (73, 74) With regard to performance, clinical empathy is a skilled interpersonal intervention that uses an asymmetrical affect attunement to modify the patient's psychobiology. This attunement may be facilitated(71) by inserting a collaborative comment or question at the right moment during the history—"Let me see if I have this right" (75)—or by making a permissive request at the seeming conclusion—"Was there anything else?"—that can bridge the synapses between and within patients and physicians. With regard to outcome, the process of self-organizing attunement is also its product. (76) Clinicians' active coparticipation in their patients' state of autonomic arousal may shift it toward homeostasis(77, 78) and decrease their allostatic load—the physiologic burden of adjusting to stressors. (79) Such an interactive physiologic regulation may even reestablish the patient's positive psychobiological state. (80)

Clinicians can learn much about the process and therapeutic potential of interpersonal neurobiology from studies of caregiver-infant interaction. Almost immediately postpartum, both caregiver and infant engage in a feedback loop of contingent, responsive, matching behaviors, primarily those conveying emotions.(51, 81) Through such communications, caregiver and infant mutually regulate each other's psychobiology, but not to an equal degree, because the caregiver's self-regulatory capacity acts as an external organizer of the infant's biobehavior.(82, 83) When successful, they self-organize a unique relationship(80) that is both their process of attunement and its product—a more stable infant neurobiology on its way to resilience and self-regulation.(84)

I am proposing that the empathic clinician may similarly use the relational process to effect a direct biological treatment. This clinical procedure is guided by the subjective experience of empathy and is operationalized by saying the right words in the right way at the right time. The intended clinical outcome is an improvement in the patient's psychobiology, perhaps just for the duration of the medical encounter, perhaps for much longer.

Two Illustrative Clinical Examples

Effective Empathy

Matthews and colleagues have described the feelings attendant to "connexional moments" in the medical encounter as "the culmination of effective empathy."(85) In essence, they argue that *effective empathy* is the subjective experience of a salutary attunement in a biopsychosocial relational process.

One of the authors (Suchman),(85) describes a patient under his care, an oft-hospitalized asthmatic woman with a "borderline personality," who was, once again, admitted for extreme dyspnea. Her respiratory distress could not be accounted for by her physical findings. Suchman encouraged the patient to talk about what she was experiencing during her latest episode. As he listened earnestly to her story, he found himself palpably experiencing the void she must have been experiencing. He conveyed this by saying, "I'm beginning to understand how hard it is to be you." Then, he recounted, "Her eyes welled up, and she nodded slowly. Seeing how much it meant to her to have someone grasp even momentarily the private hell she had to endure, I found my eyes welling up, too, and I felt a chill in my neck and spine. For a moment, it felt like we were joined, both parts of some larger whole; it was very peaceful and reassuring, even loving. A feeling of calm and joy was with me for the rest of the day. R seemed peaceful, too. She went home the next day, and although she is certainly not 'cured' of her personality disorder, she has not been admitted again in the 5 years since." (85)

This vignette illustrates how clinical empathy was used as a clinical procedure.

The clinical indication was a problem with R's psychobiology that had not responded very well to her previous medical care. The clinical procedure began with Suchman's recognition that his dysphoric feelings about R were empathic indicators of what she was probably feeling. His statement, "I'm beginning to understand how hard it is to be you," could only have been convincing because it was accompanied by emotional expressions that R perceived as authentic and attuned to her predicament.(71) This attuned empathic communication joined Suchman and R as co-participants in an emergent interpersonal neurobiology. Their co-participation was evidenced by the responsive welling up of R's

eyes leading to a reciprocal autonomic response in Suchman. Their emergent relationship was evidenced by their apparently shared feelings of joy and calmness.

The clinical outcome of this encounter may be gauged by the ensuing feelings of mutual satisfaction, understood as subjective indicators of at least two overlapping psychobiological effects. The experience of a secure attachment has a non-specific stress-buffering effect.(86) In addition, this clinical procedure may have had a transformative effect that changed how R and Suchman felt about themselves as well as how they felt about each other. Other clinical outcomes were a decrease in hospital admissions for R and a decrease in the risk of burnout(87, 88) for Suchman.

The mutually salutary effects in the case of Suchman and R apparently continued long after the medical encounter. The effectiveness of that clinical procedure is very likely renewed at relevant moments by the recollections each party has given to the other. Effective Attitude

Clinicians may employ their affect attunement to improve patients' attitudes toward their personhood, now threatened by a medical problem. The successful conveyance of a salutary attitude can change the meaning of the experience along with its psychobiologic consequences.

One of my patients reported the lifelong consequences of a change in attitude that occurred during a medical encounter. She will always remember an off-hand response that rescued her from self-defeating despair. When she was 20 and single, her gynecologist diagnosed genital herpes during a pelvic examination. The patient felt like a pariah. "No one will ever want me," she remembers sobbing. "Can I ever have an honest sex life?" The gynecologist matter-of-factly replied, "I don't know why not." He then

followed up with information about herpes, recommended a helpful book, and informed her of an Internet dating service for people with herpes. He also pointed out that now she had another good reason to establish trusting relationships before sexual relations. In the telling of this story, now 10 years later, the patient triumphantly reenacted the casual hand gesture, shrug, and bemused expression that accompanied the physician's words. That attitude with its concomitant psychobiology was no longer just his; it had become hers. The patient recalled that she had immediately felt herself transformed from a disdained miscreant to a person with a manageable problem. She also recalled that what mainly repaired her self-image was seeing herself reflected by her physician's expression. This deftly performed interpersonal clinical procedure, which entailed one phrase, a few expressive gestures, and medical information, revitalized her psychobiology—then and now. While I do not know this physician, the biopsychosocial relational process suggests that his psychobiology likewise benefited from this attuned self-organizing process because the feedback loop was now infused with the patient's appreciation.(89)

The Issue of Clinical Significance

An empathic patient-physician relationship has been found to improve patients' adherence to and satisfaction with their treatment.(90) Patients' satisfaction can be considered an indicator of a salutary psychobiology.(4, 91) Since adherence and satisfaction contribute both indirectly and directly to health outcomes,(92) the clinical significance of clinical empathy is strongly supported. Even if the criterion for clinical

significance is more narrowly defined as the kind of direct biological effects attributable to a pharmacologic agent, there is still strong supporting evidence from three levels of psychosocial research. At the macro level, epidemiologic studies have long demonstrated that social support, a major component of which is emotional support, influences biological variables that affect the development and course of a wide range of biomedical diseases.(93, 94) At the micro level, psychosocial influences have been demonstrated to exert similar effects on relevant biological variables with similar biomedical consequences.(95, 96) Less work has been done at the dyadic level of relationships, but studies have demonstrated that marital conflict can result in deleterious alterations in cellular immune regulation and endocrine function, while harmonious relationships can enhance these physiological systems.(97, 98)

One caution about the biomedical consequences of psychosocial interventions is that while the changes in relevant biological variables are statistically significant and in the right direction they may be too small to be clinically significant. Nevertheless, biological changes reported as lacking clinical significance in short-term studies may later be found by long-term studies to be biomedical risk factors. Many years separate sun exposure and melanoma, head injury and Alzheimer's disease, influenza and Parkinsonism.

According to complexity theory, even small statistically significant changes in relevant biological variables like glycohemoglobin, blood pressure, and cholesterol levels may eventually have clinical consequences. By analogy, even though the psychobiological effects of a change of attitude may be too small to be clinically significant during the medical encounter, they may have large biomedical effects over time.

Future Directions

Empirical studies of clinical empathy might proceed in three steps. The first would be to establish the presence of interpersonal neurobiological and empathic responses during the medical encounter. This could be done during the medical encounter by performing neuroimaging and physiologic studies that have been used to monitor the process of psychotherapy,(99, 100) and immediately afterward by administering an empathy scale, such as the Relationship Inventory,(101) to both patients and physicians. Second, researchers could demonstrate subsequent changes in biological variables that are plausibly relevant to disease, employing the methodology used to study the psychoneuroendocrine effects of conflict and resolution in spouses.(97, 102) The third and final step would be to explore the strategies and techniques(71) that an empathic physician can use with the patient(103) in a way that maximizes the therapeutic potential of the biopsychosocial relational process.

References

- 1. **Balint M.** *The Doctor, His Patient, And The Illness* New York: International Universities Press; 1957.
- 2. **Peabody F.** The care of the patient. *JAMA*. 1927;88:877-82.
- 3. **Suchman AL.** A new theoretical foundation for relationship-centered care. Complex responsive processes of relating. *J Gen Intern Med.* 2006;21(Suppl 1):S40-4.
- 4. **Kandel ER, Squire LR.** Neuroscience: Breaking down scientific barriers to the study of brain and mind. *Science*. 2000;290(5494):1113-120.
- 5. **DiMascio A, Boyd R, Greenblatt M, Solomon H.** The psychiatric interview: A sociophysiologic study. *Dis Nerv Syst.* 1955;16:4-9.
- 6. **Waid WM.** Origins of sociophysiology. In: Waid WM, editor. *Sociophysiology*. New York,: Springer-Verlag; 1984:3-20.
- 7. **Mendoza SP.** Sociophysiology of well-being in nonhuman primates. *Lab Anim Sci.* 1991;41(4):344-9.
- 8. **Siegel D.** Toward an interpersonal neurobiology of the developing mind:

 Attachment relationships, "mindsight," and neural integration. *Infant Ment Health J.* 2001;22(1-2):67-94.
- 9. **Stern D.** The Interpersonal World of the Infant: A View from Psychoanalysis and Developmental Psychology New York: Basic Books; 1985.

- Levenson R, Ruef A. Physiological aspects of emotional knowledge and rapport.
 In: Ickes W, editor. *Empathic Accuracy*. New York: The Guilford Press; 1997:44-72.
- 11. **Rizzolatti G, Craighero L.** The mirror-neuron system. *Ann Rev Neurosci*. 2004;27:169-92.
- 12. **Carr L, Iacoboni M, Dubeau MC, Mazziotta JC, Lenzi GL.** Neural mechanisms of empathy in humans: A relay from neural systems for imitation to limbic areas. *Proc Natl Acad Sci U S A*. 2003;100(9):5497-502.
- 13. **Gallese V.** The roots of empathy: The shared manifold hypothesis and the neural basis of intersubjectivity. *Psychopathology*. 2003;36(4):171-80.
- 14. DiMascio A, Boyd R, Greenblatt M. Physiological correlates of tension and antagonism during psychotherapy: A study of "interpersonal physiology."
 Psychosom Med. 1957;19:99-104.
- 15. **Malmo RB, Boag TJ, Smith AA.** The physiological study of personal interaction. *Psychosom Med.* 1957;19:105-19.
- 16. **Robinson J, Herman A, Kaplan B.** Autonomic responses correlate with counselor-client empathy. *J Couns Psychol*. 1982;29:195-8.
- 17. **Levenson RW, Ruef AM.** Empathy: a physiological substrate. *J Pers Soc Psychol*. 1992;63(2):234-46.
- 18. **Levenson RW, Gottman JM.** Marital interaction: physiological linkage and affective exchange. *J Pers Soc Psychol*. 1983;45(3):587-97.
- 19. **Barchas P, Jose II W, Payne B, Harris W.** An attention-regulating function of social hierarchies: high status, attention, and the CNV brain wave. In: Barchas P,

- editors. *Social Hierarchies: Essays Toward a Sociophysiological Perspective*. Westport, CT: Greenwood Press; 1984:133-146.
- Mendoza S. The psychobiology of social relationships. In: Barchas P, Mendoza S, eds. Social Cohesion: Essays Toward a Sociophysiological Perspective.
 Westport, Connecticut: Greenwood Press; 1984:3-30.
- 21. **Gust DA, Gordon TP, Brodie AR, McClure HM.** Behavioral and physiological response of juvenile sooty mangabeys to reunion with their mothers following a year's absence. *Dev Psychobiol.* 1992;25(8):613-22.
- 22. McKinney WT. Animal research and its relevance to psychiatry 1. In: Kaplan HI, Saddock BJ, editors. *Comprehensive Textbook of Psychiatry*. Vol. 1. Baltimore: Williams & Wilkins; 1995:397-411.
- 23. **Insel TR, Hulihan TJ.** A gender-specific mechanism for pair bonding: oxytocin and partner preference formation in monogamous voles. *Behav Neurosci*. 1995;109:782–9.
- 24. **Haber S, Barchas P.** The regulatory effect of social rank on behavior after amphetamine administration. In: Barchas P, ed. *Social Hierarchies, Essays Toward a Sociphysiological Perspective*. Wesport, Connecticut: Greenwood Press; 1984:119-132.
- 25. Gust DA, Gordon TP, Wilson ME, Ahmed-Ansari A, Brodie AR, McClure HM. Formation of a new social group of unfamiliar female rhesus monkeys affects the immune and pituitary adrenocortical systems. *Brain Behav Immun*. 1991;5(3):296-307.

- 26. Darney KJ Jr Goldman J, Vandenbergh J. Neuroendocrine responses to social regulation of puberty in the female house mouse. *Neuroendocrinology*. 1992;55(4):434-43.
- 27. **Dunlap JL, Zadina JE, Gougis G.** Prenatal stress interacts with prepuberal social isolation to reduce male copulatory behavior. *Physiol & Behav*. 1978;21:(873-5).
- 28. **Virgin CJ Jr, Sapolsky RM.** Styles of male social behavior and their endocrine correlates among low-ranking baboons. *Am J Primatol*. 1997;42(1):25-39.
- 29. **Sachs BD, Akasofu K, Citron JH, Daniels SB, Natoli JH.** Noncontact stimulation from estrous females evokes penile erection in rats. *Physiol Behav*. 1994;55(6):1073-9.
- 30. **Gardner R Jr.** Sociophysiology as the basic science of psychiatry. *Theor Med*. 1997;18(4):335-56.
- 31. **Stern D.** The Present Moment in Psychotherapy and Everyday Life New York: WW Norton & Co; 2004.
- 32. **Siegel D.** The Developing Mind: Toward a Neurobiology of Interpersonal Experience New York: Guilford Press; 1999.
- 33. **Eisenberg L.** The social construction of the human brain. *Am J Psychiatry*. 1995;152:1563-75.
- 34. **Kandel ER.** Psychotherapy and the single synapse: The impact of psychiatric thought on neurobiologic research. *N Eng J Med.* 1979;301(19):1028-37.
- 35. **Greenough W, Black J.** Induction of brain structure by experience: substrates for cognitive development. In: Gunnar M, Nelson C, editors. *Minnesota Symposia on*

- *Child Psychology: Developmental Neuroscience*. Vol. 24. Hillsdale, NJ: Lawrence Erlbaum; 1992.
- 36. **Insel T.** A neurological basis of social attachment. *Am J Psychiatry*. 1997;154:726-35.
- 37. **Nelson E, Panksepp J.** Brain substrates of infant-mother attachment: contributions of opioids, oxytocin, and norepinephrine. *Neurosci Biobehav Rev*. 1998;22(3):437-52.
- 38. **Young LJ, Wang Z, Insel TR.** Neuroendocrine bases of monogamy. *Trends Neurosci.* 1998;21(2):71-5.
- 39. **Hofer MA.** Relationships as regulators: a psychobiologic perspective on bereavement. *Psychosom Med.* 1984;46:183-97.
- 40. **Hofer MA.** The Roots of Human Behavior, An Introduction to the Psychobiology of Early Development 2 San Francisco: W. N. Freeman; 1981.
- 41. **Hofer MA.** On the nature and consequences of early loss. *Psychosom Med.* 1996;58(6):570-81.
- 42. **Beebe B, Alson D, Jaffe J, Feldstein S, Crown C.** Vocal congruence in mother-infant play. *J Psycholinguist Res.* 1988;17(3):245-59.
- 43. **Tronick ED, Als H, Brazelton, TB.** Mutuality in mother-infant interaction. *J Commun.* 1977;27:74-79.
- 44. **Cohn J, Tronick E.** Mother-infant face-to-face interaction: Influence is bidirectional and unrelated to periodic cycles in either partner's behavior. *Dev Psychol.* 1988;24:386-392.

- 45. **Jaffe J, Beebe B, Feldstein S, Crown CL, Jasnow MD.** Rhythms of dialogue in infancy: Coordinated timing in development. *Monogr Soc Res Child Dev*. 2001;66(2):i-viii, 1-132.
- 46. Jonsson CO, Clinton DN, Fahrman M, Mazzaglia G, Novak S, Sorhus K.
 How do mothers signal shared feeling-states to their infants? An investigation of affect attunement and imitation during the first year of life. *Scand J Psychol*.
 2001;42(4):377-81.
- 47. **Schore AN.** Early organization of the nonlinear right brain and development of a predisposition to psychiatric disorders. *Dev Psychopathol*. 1997;9(4):595-631.
- 48. **Faude J, Jones C, Robins M.** The affective life of infants: Empirical and theoretical foundations. In: Nathanson D, editor. *Knowing Feeling, Affect, Script, and Psychotherapy*. New York: WW Norton & Co; 1996:219-56.
- 49. **Bowlby J.** A Secure Base New York: Basic Books; 1988.
- 50. **Murray AD.** Infant crying as an elicitor of parental behavior: An examination of two models. *Psychol Bull.* 1979;86:191-215.
- 51. **Meltzoff AN, Moore MK.** Newborn infants imitate adult facial gestures. *Child Dev.* 1983;54(3):702-9.
- 52. **Rizzolatti G, Fadiga L, Fogassi L, Gallese V.** From mirror neurons to imitation: Facts and speculations. In: Meltzoff A, Prinz W, editors. *The Imitative Mind:*Development, Evolution, and Brain Bases. Cambridge Studies in Cognitive

 Perceptual Development. New York, NY: Cambridge University Press; 2002:247-66.

- 53. **Gallese V, Fadiga L, Fogassi L, Rizzolatti G.** Action recognition in the premotor cortex. *Brain*. 1996;119(Pt 2):593-609.
- 54. **Rizzolatti G, Fadiga L, Gallese V, Fogassi L.** Premotor cortex and the recognition of motor actions. *Brain Res Cogn Brain Res.* 1996;3(2):131-41.
- 55. **Fadiga L, Fogassi L, Pavesi G, Rizzolatti G.** Motor facilitation during action observation: A magnetic stimulation study. *J Neurophysiol*. 1995;73(6):2608-11.
- 56. **Iacoboni M, Woods RP, Brass M, Bekkering H, Mazziotta JC, Rizzolatti G.**Cortical mechanisms of human imitation. *Science*. 1999;286(5449):2526-8.
- 57. **Fogassi L.** Evolution of language from action understanding. The International Conference on Audio-Visual Speech Processing. St. Jorioz, France; 2003.
- 58. **Buccino G, Binkofski F, Fink G, Fadiga L, Fogassi L, Gallese V, et al.** Action observation activates premotor and parietal areas in a somatotopic manner: An fMRI study. *Eur J Neurosci*. 2001;13(2):400-4.
- 59. **Bremmer F, Schlack A, Shah N, Zafiris O, Kubischik M, Hoffmann K, et al.**Polymodal motion processing in posterior parietal and premotor cortex: a human fMRI study strongly implies equivalencies between humans and monkeys.

 Neuron. 2001;29:287-96.
- 60. **Fecteau S, Carmant L, Tremblay C, Robert M, Bouthillier A, Theoret H.** A motor resonance mechanism in children? Evidence from subdural electrodes in a 36-month-old child. *Neuroreport*. 2004;15(17):2625-7.
- 61. **Iacoboni M.** Understanding others: Imitation, language, empathy. In: Hurley S, Chater N, editors. *Perspective on Imitation: From Neurons to Memes*. Vol. 1:

- Mechanisms of Imitation and Imitation in Animals. Cambridge, MA: MIT Press; 2003.
- 62. **Rizzolatti G, Fadiga L, Fogassi L, Gallese V.** Resonance behaviors and mirror neurons. *Arch Ital Biol.* 1999;137(2-3):85-100.
- 63. **Leslie K, Johnson-Frey S, Grafton S.** Functional imaging of face and hand imitation: Towards a motor theory of empathy. *Neuroimage*. 2004;21(2):601-7.
- 64. Shamay-Tsoorya SG, Lester H, Chisin R, Israel O, Bar-Shalom R, Peretz A, et al. The neural correlates of understanding the other's distress: A positron emission tomography investigation of accurate empathy. *NeuroImage*. 2005;27(2):468-72.
- 65. **Vanderpool JP, Barratt ES.** Empathy: towards a psychophysiological definition. *Dis Nerv Syst.* 1970;31(7):464-7.
- 66. **Gottman JM, Levenson RW.** A valid procedure for obtaining self-report of affect in marital interaction. *J Consult Clin Psychol*. 1985;53(2):151-60.
- 67. **Hojat M, Gonnella JS, Nasca TJ, Mangione S, Vergare M, Magee M.**Physician empathy: Definition, components, measurement, and relationship to gender and specialty. *Am J Psychiatry*. 2002;159:1563-9.
- 68. **Eisenberg N, Fabes R.** Empathy: Conceptualization, measurement, and relation to prosocial behavior. *Motivation and Emotion*. 1990;14(2):131-50.
- 69. Halpern J. Empathy: Using resonance emotions in the service of curiosity. In: Spiro H, Cunen M, Peschel E, St James D, editors. *Empathy and the Practice of Medicine: Beyond Pills and Scalpel*. New Haven, CT: Yale University Press; 1993:160-73.

- 70. **Orlinksy D, Grawe K, Parks B.** Process and outcome in psychotherapy—noch einmal. In: Bergn A, Garfield S, editors. *Handbook of Psychotherapy and Behavior*. 4th ed. New York: Wiley; 1994:270-378.
- 71. **Larson EB, Yao X.** Clinical empathy as emotional labor in the patient-physician relationship. *JAMA*. 2005;249(9):1100-06.
- 72. **Simpson J, Rholes W.** Stress and secure base relationships in adulthood. *Adv Pers Rel.* 1994;5:181-204.
- 73. **Spiegel D, Hunt T, Dondershine HE.** Dissociation and hypnotizability in posttraumatic stress disorder. *Am J Psychiatry*. 1988;145(3):301-05.
- 74. **Miller WR.** Rediscovering fire: Small interventions, large effects. *Psychol Addictive Behav.* 2000;14(1):6-18.
- 75. **Coulehan JL, Platt FW, Egener B, Frankel R, Lin CT, Lown B, et al.** Let me see if I have this right . . . : words that help build empathy. *Ann Intern Med*. 2001;135(3):221-7.
- 76. **Stacey R.** Organizational identity: The paradox of continuity and potential transformation at the same time. *Group Analysis*. 2005;38(4):477-94.
- 77. **Porges SW.** Love: An emergent property of the mammalian autonomic nervous system. *Psychoneuroendocrinology*. 1998;23(8):837-61.
- 78. **Carter CS.** Neuroendocrine perspectives on social attachment and love. *Psychoneuroendocrinology*. 1998;23(8):779-818.
- 79. **McEwen BS.** Stress, adaptation, and disease. Allostasis and allostatic load. *Ann N Y Acad Sci.* 1998;840:33-44.

- 80. **Schore A.** Early relational trauma, disorganized attachment, and the development of a predisposition to violence. In: Solomon M, Siegel D, editors. *Healing Trauma*. New York: WW Norton & Company; 2003:107-167.
- 81. **Ziskind E.** The social context of nonverbal behavior. In: Philippot P, Feldman R, Coats E, eds. *Infant and Child Development*. New York, NY: Cambridge University Press; 1999.
- 82. **Stern D.** The early differentiation of self and other. In: Kaplan S, Lichtenberg J, editors. *Reflections on Self Psychology*. Hillsdale, NJ: Analytic Press; 1983.
- 83. **Sroufe L.** *Emotional Development: The Organization of Emotional Life in the Early Years* New York: Cambridge University Press; 1996.
- 84. **Feldman R, Greenbaum CW Yirmiya N.** Mother-infant affect synchrony as an antecedent of the emergence of self-control. *Dev Psychol.* 1999;35(1):223-31.
- 85. Matthews DA, Suchman AL, Branch WT Jr. Making "connexions":
 Enhancing the therapeutic potential of patient-clinician relationships. *Ann Intern Med.* 1993;118(12):973-7.
- 86. **Berkman LF, Glass T, Brissette I, Seeman TE.** From social integration to health: Durkheim in the new millennium. *Soc Sci Med.* 2000;51:843-57.
- 87. **Chopra SS, Sotile WM, Sotile MO.** Physician burnout. *JAMA*. 2004;291(5):633.
- 88. **Maslach C, Schaufeli WB, Leiter MP.** Job burnout. *Annu Rev Psychol*. 2001;52:397-422.
- 89. **Horowitz CR, Suchman AL, Branch WT Jr, Frankel RM.** What do doctors find meaningful about their work? *Ann Intern Med.* 2003;138(9):772-5.

- 90. **Roter DL, Frankel RM, Hall JA, Sluyter D.** The expression of emotion through nonverbal behavior in medical visits: Mechanisms and outcomes. *J Gen Intern Med*. 2006;21(Suppl 1):S 28-34.
- 91. **Levine S, Lyons DM, Schatzberg AF.** Psychobiological consequences of social relationships. *Ann N Y Acad Sci.* 1997;807:210-8.
- 92. **Hauck FR, Zyzanski SJ, Alemagno SA, Medalie JH.** Patient perceptions of humanism in physicians: effects on positive health behaviors. *Fam Med*. 1990;22(6):447-52.
- 93. **Seeman TE, Crimmins E.** Social environment effects on health and aging: integrating epidemiologic and demographic approaches and perspectives. *Ann NY Acad Sci.* 2001;954:88-117.
- 94. **Cobb S.** Social support as a moderator of life stress. *Psychosom Med*. 1976;38(5):300-14.
- 95. **Smith TW, Ruiz JM.** Psychosocial influences on the development and course of coronary heart disease: Current status and implications for research and practice. *J Consult Clin Psychol.* 2002;70(3):548-568.
- 96. **Vitetta L, Anton B, Cortizo F, Sali A.** Mind-body medicine: Stress and its impact on overall health and longevity. *Ann NY Acad Sci.* 2005;1057:492-505.
- 97. **Kiecolt-Glaser J, Newton T, Cacioppo JT, MacCallum RC, Glaser R, Malarkey WB.** Marital conflict and endocrine function: Are men really more physiologically affected than women? *J Consult Clin Psychol*. 1996;64(2):324-32.

- 98. **Kiecolt-Glaser JK, Malarkey W, Chee M, Newton T, Cacioppo JT, Mao HY, et al..** Negative behavior during marital conflict is associated with immunological down-regulation. *Psychosom Med.* 1993;55(5):395-409.
- 99. **Kandel E.** Cellular mechanisms of learning and the biological basis of individuality. In: Kandel E, Schwartz J, Jessell T, editors. *Principles of Neural Science*. 4th ed. New York: McGraw-Hill; 2000:1247-79.
- 100. **Marci CD, Moran EK, Orr SP.** Physiologic evidence for the interpersonal role of laughter during psychotherapy. *J Nerv Ment Dis.* 2004;192(10):689-95.
- 101. **Barrett-Leonard G.** The Relationship Inventory now: Issues and advances in theory, method, and use. In: Greenberg L, Pinsoff W, editors. *The Psychotherapeutic Process: A Research Handbook*. New York: Guilford; 1986:439-476.
- 102. Kiecolt-Glaser JK, Glaser R, Cacioppo JT, MacCallum RC, Snydersmith M, Kim C, et al. Marital conflict in older adults: endocrinological and immunological correlates. *Psychosom Med.* 1997;59(4):339-49.
- 103. **Novack DH, Suchman AL, Clark W, Epstein RM, Najberg E, Kaplan C.**Calibrating the physician: Personal awareness and effective patient care. *JAMA*. 1997;278(6):502-9.