Lanius/Vermetten Book Chapter

"The Hidden Epidemic: The Impact of Early Life Trauma on Health and Disease" Cambridge University Press, 2008

Chapter title:

The Relationship of Adverse Childhood Experiences to Adult Health, Well-being, Social Function, and Healthcare 6-01-2007 v1.1

Vincent J. Felitti, MD (VJFMDSDCA@mac.com) Kaiser Permanente Medical Care Program, San Diego, CA

Robert F. Anda, MD, MS (RFA1@cdc.gov)
US Centers for Disease Control and Prevention, Atlanta, GA

The Relationship of Adverse Childhood Experiences to Adult Health, Well-being, Social Function, and Healthcare

"In my beginning is my end."

T.S. Eliot, Four Quartets

Introduction:

Medicine increasingly recognizes that early life events, including oftenunrecognized emotional traumas, have both dramatic and long-lasting effects on the neural and biological systems involved in well-being, biomedical disease, social function, and psychopathology. Our current understanding is relatively new, developing in the twentieth century. The turning point in modern understanding is commonly attributed to the great French neurologist, Jean-Martin Charcot, and his studies of hysteria in the closing years of the nineteenth century. Janet, Freud, and Breuer soon were to explore the role of emotional trauma in human development.² Subsequently, psychoanalysts and poets provided the early general insights into many of these developmental phenomena. These descriptions represented an enormous shift away from Biblical and Renaissance views that mental illness and biomedical disease were god-given, or the result of satanic possession, or of moral failure. Judith Herman provides a chronology of these transformations in her book, *Trauma and Recovery*. Harry Harlow provided in his primate work experimental demonstration of the need in infancy for more than environmental warmth, physical safety, and food if a functional adult rhesus monkey is to emerge. John Bowlby⁵ described the importance of the missing ingredient in his work on human attachment. Neurobiologists are currently discovering and describing the intermediary biomedical mechanisms by which these processes manifest themselves.

The focus of this chapter will be an examination of the relations between early life traumas and the effect they have on nearly all systems of the body. To do this, we will draw on our experience with the Adverse Childhood Experiences (ACE) Study, a major American medical study providing retrospective and prospective analysis in over 17,000 individuals of the effect of traumatic life experiences during the first eighteen years of life on later well-being, social function, health risks, disease burden, healthcare costs, and life expectancy.⁶

The ACE Study is an outgrowth of repeated counterintuitive observations made while operating a weight loss program that uses the technique of supplemented fasting, which allows non-surgical weight reduction of approximately three hundred pounds (135 Kg) per year. Unexpectedly, our Weight Program had a high drop-out rate, limited almost exclusively to patients successfully losing weight. Exploring the reasons underlying the high prevalence of patients inexplicably fleeing their own success in the Weight Program ultimately led us to recognize that certain of the more intractable public health problems like obesity were *also* unconscious, or occasionally conscious, solutions to problems dating back to the earliest years, but hidden by time, by shame, by secrecy, and by social taboos against exploring certain areas of life experience. It became evident that traumatic life experiences during childhood and adolescence were far more common

than generally recognized, were complexly interrelated, and were associated in a strong and proportionate manner to outcomes important to medical practice, public health, and the social fabric of the nation. In the context of everyday medical practice, we came to recognize that the earliest years of infancy and childhood are not lost but, like a child's footprints in wet cement, are often life-long.

The findings from the ACE Study provide a remarkable insight into how we become what we are as individuals and as a nation. They are important medically, socially, and economically. Indeed, they have given us reason to reconsider the very structure of primary care medical practice in America.

Outline of the ACE Study and its setting:

The Adverse Childhood Experiences (ACE) Study was carried out in Kaiser Permanente's Department of Preventive Medicine in San Diego, in collaboration with the US Centers for Disease Control and Prevention (CDC). This particular Department of Preventive Medicine provided an ideal setting for such collaboration because for many years we have carried out at one site detailed biomedical, psychological, and social (biopsychosocial) evaluations of over 50,000 adult Kaiser Health Plan members a year. The CDC contributed the essential skill sets for study design and massive data management required for meaningful interpretation of clinical observations.

Kaiser Health Plan patients are middle-class Americans; all have high quality health insurance. In any 4-year period, 81% of adult Plan members in San Diego choose to come in for comprehensive medical evaluation. We asked 26,000 consecutive adults coming through the Department if they would help us understand how childhood events might affect adult health status. The majority agreed and, after certain exclusions for incomplete data and duplicate participation, the ACE Study cohort had over 17,000 individuals. The Study was carried out in two waves, to allow mid-point correction.

The participants were 80% white including Hispanic, 10% black, and 10% Asian; 74% had attended college; their average age was 57. Almost exactly half were men, half women. This is a solidly middle-class group from the 6th largest city in the nation; it is not a group that can be dismissed as atypical, aberrant, or 'not in my practice'. Disturbingly, it is us – not a point to be overlooked when considering the problems of translating this information into action.

Eight categories of adverse childhood experiences (ACEs) were initially studied in the first wave; two categories of neglect were added in the second wave. We empirically selected these categories because of their discovered high prevalence in the Weight Program. Their prevalence in a general, middle-class population was also unexpectedly high. We created for each individual an ACE Score, a count of the number of *categories* of adverse childhood experience that had occurred during the first eighteen years of life. ACE Score does not tally incidents within a category. The scoring system is simple: exposure during childhood or adolescence to any one category of adverse experience is scored as one point. There is no further scoring within a category; thus, an alcoholic and a drug user within a household score the same as one alcoholic; multiple

sexual molestations by multiple individuals are scored as one point. If anything, this tends to understate our findings. The ACE Score therefore can range from 0 to 8 or 10, depending on the data being from Wave 1 or Wave 2. Specifics of the questions underlying each category are detailed in our original article.⁶

Only one third of this middle-class population had an ACE Score of 0. If any one category was present, there is 87% likelihood that at least one more category will be present. One in six individuals had an ACE Score of 4 or more, and one in ten had an ACE Score of 5 or more. Thus, every physician sees several high ACE Score patients each day. Typically, they are the most difficult patients of the day. Women were 50% more likely than men to have experienced five or more categories. Here is a key to what, in mainstream epidemiology, appears as women's natural proneness to undefined health problems like fibromyalgia, chronic fatigue syndrome, irritable bowel syndrome, and chronic non-malignant pain syndromes. We now see these as medical constructs, artifacts resulting from medical blindness to social realities and ignorance of the impact of gender.

Somewhat surprisingly, the ACE categories turned out to be approximately equal in impact; an ACE Score of 4 consists of *any* four of the categories. The categories do not occur randomly; the number of individuals with high ACE Scores is distinctly higher than if the categories exist independently of each other. The ten reference categories experienced during childhood or adolescence are as below, with their prevalences in parentheses:

- Abuse
- 1. emotional recurrent humiliation (11%)
- 2. physical beating, not spanking (28%)
- 3. contact sexual abuse (28% women, 16% men; 22% overall)
- Household dysfunction
- 1. mother treated violently (13%)
- 2. household member was alcoholic or drug user (27%)
- 3. household member was imprisoned (6%)
- 4. household member was chronically depressed, suicidal, mentally ill, in psychiatric hospital (17%)
- 5. not raised by both biological parents (23%)
- Neglect
- 1. physical (10%)
- 2. emotional (15%)

The essence of the ACE Study has been to match retrospectively, approximately a half century after the fact, an individual's current state of health and well-being against adverse events in childhood (the ACE Score), and then to follow the cohort forward to match ACE Score prospectively against doctor office visits, ER visits, hospitalization,

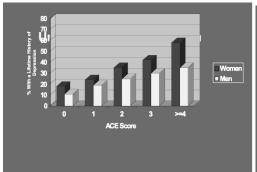
pharmacy costs, and death. We recently have passed the ten-year mark in the prospective arm of the Study.

Findings:

We will illustrate with a sampling from our findings in the ACE Study the longlasting, strongly proportionate, and often profound relationship between adverse childhood experiences and important categories of human well-being, health risks, disease burden, social function, and healthcare costs - decades later.

Well-being:

The relationship between ACE Score and self-acknowledged chronic depression is illustrated in FIG 1A.⁸ Should one not be confident of the reliability of self-acknowledged chronic depression, there is a similar but stronger relationship between ACE Score and later suicide attempts as shown in FIG 1B.⁹ The *p* value of all graphic depictions herein is .001 or better.



Childhood Experiences
Underlie Suicide Attempts

4+

99 19 15

99 19 15

10

ACE Score

Fig 1A Fig 1B

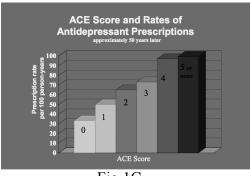


Fig 1C

One continues to see a proportionate relationship between ACE Score and depression by analysis of prescription rates for antidepressant medications after a ten year prospective follow-up, now approximately fifty to sixty years after the fact. (FIG 1C). 10 It would appear that depression, often unrecognized in medical practice, is in fact common and has deep roots, commonly going back to the developmental years of life.

An analysis of population attributable risk (that portion of a problem in the overall population whose prevalence can be attributed to specific risk factors) shows that 54% of current depression and 58% of suicide attempts in women can be attributed to adverse childhood experiences. Whatever later factors might trigger suicide, childhood experiences can not be left out of the equation. McEwen and Singer¹¹ have described this general concept of background burden as allostatic load.

A similar relationship exists between ACE Score and later hallucinations, shown in FIG 1D. Lest one reasonably suspect that, at ACE Score 7 or higher, people will likely be using street drugs or alcohol to modulate their feelings, and that *these* might be the cause of hallucinations, we have corrected for alcohol and drug use and find the same relationship exists.¹²

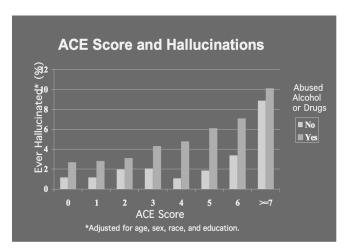
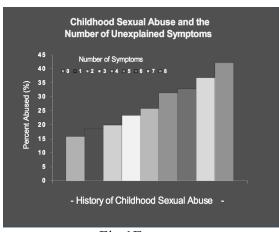


Fig 1D

Physicians dealing with somatization and psychophysiologic disorders, as well as those physicians dreading such patients, will find Figure 1E of special interest. Indeed, this figure exemplifies our observation in the Weight Program that what one sees, the presenting problem, is often only the marker for the real problem, which lies buried in time, concealed by patient shame, secrecy, and physician discomfort – and sometimes amnesia. Amnesia, usually considered a theatrical device of Hollywood movies of the 1940s is in fact alive and well, though unrecognized, in everyday medical practice. In our Weight Program, we found 12% of the participants were partially or totally amnesic for a period of their lives, typically the few years before weight gain began. In the ACE Study, we found that there was a distinct relationship of ACE Score to impaired memory of childhood, and understand this phenomenon to be reflective of dissociative responses. (FIG 1F)



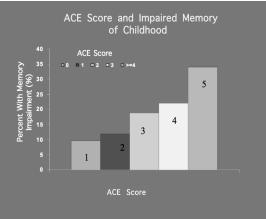


Fig 1E Fig 1F

All told, it is clear that adverse childhood experiences have a profound, proportionate, and long-lasting effect on well-being, whether measured by depression or suicide attempts, by protective unconscious devices like somatization disorders and dissociation, or by self-help attempts that are misguidedly addressed solely as long-term health risks -- perhaps because we physicians are less than comfortable acknowledging the manifest short-term benefits these "health risks" offer to the patient dealing with hidden trauma.

Health risks:

The most common contemporary health risks are smoking, alcoholism, illicit drug use, obesity, and high level promiscuity. Though widely understood to be harmful to health, each is notably difficult to give up. Conventional logic is not particularly useful in understanding this apparent paradox. As though opposing forces are not known to exist commonly in biological systems, little consideration is given to the possibility that many health risks might *also* be personally beneficial. For instance, the psychoactive benefits of nicotine were understood decades before its risks were recognized. We repeatedly hear from patients of the benefits of these "health risks." Indeed, relevant insights are even built into our language: "Have a smoke, relax." "Sit down and have something to eat. You'll feel better." Or, needing 'a fix', referring to iv drug use. Conversely, the popular term "drug abuse" serves to conceal the functionality of such behavior.

In the ACE Study, we found strong, proportionate relationships between the number of categories of adverse childhood experience (ACE Score) and the use of various psychoactive materials or behaviors. The saying, "It's hard to get enough of something that *almost* works." provides insight. Three common categories of what are usually termed addictions (the unconscious compulsive use of psychoactive agents) are illustrated in this section. Self-acknowledged current smoking (FIG 2A)^{13, 14}, self-defined alcoholism (FIG 2B)^{6, 15}, and self-acknowledged intravenous drug use (FIG 2C)¹⁶ are strongly related in a proportionate manner to our several specific categories of adverse experiences during childhood.



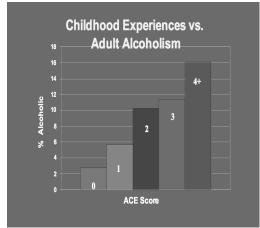


Fig 2A Fig 2B

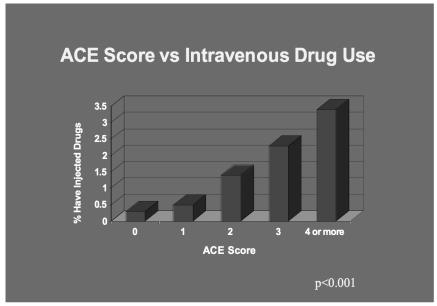


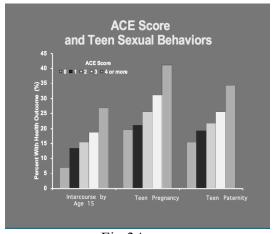
Fig 2C

The relationship of ACE Score to iv drug use is particularly striking, given that male children with ACE Score 6 or more have a 4,600% increased likelihood of later becoming an iv drug user, as contrasted with an ACE Score 0 male; this moves the probability from an arithmetic to an exponential progression. Relationships of this magnitude are rare in Epidemiology. This, coupled with related information, suggests that the basic cause of addiction is predominantly experience-dependent during childhood and not substance-dependent. This challenge to the usual concept of the cause of addictions has significant implications for medical practice and for treatment programs.¹⁷

Social function

Using teen pregnancy and promiscuity as measures of social function, we found that ACE Score has a proportionate relationship to these outcomes. (FIG 3A, 3B.) So

too does miscarriage of pregnancy, indicating the complexity of the relationship of early life psychosocial experience to what are usually considered purely biomedical outcomes. Additionally, we found that self-rated job performance correlated inversely with ACE Score. (FIG 3C). This of course is in addition to previously discussed problems like depression, alcoholism, and iv drug use that often are used as markers of impaired work performance by those studying social function.



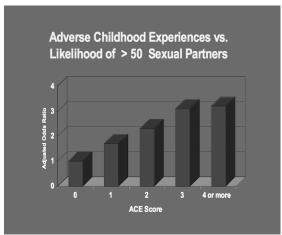


Fig 3A Fig 3B

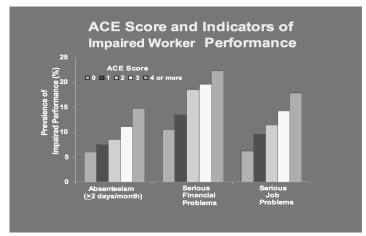
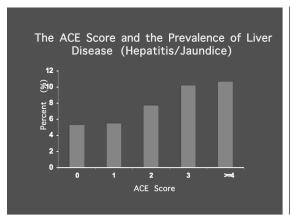


Fig 3C

Biomedical disease

We found in the ACE Study that biomedical disease in adults has a significant relationship to adverse life experiences in childhood. The implication of this observation that life experience can transmute into organic disease over time is a profound change from an earlier era when infectious diseases like rheumatic fever or polio, or nutritional deficiency like pellagra, would come to mind as the main medical link between childhood events and adult disease. In spite of this change in the etiology of biomedical outcomes, we find no evidence that there has been a change in the frequency of overall adverse childhood experiences in various age cohorts spanning the twentieth century.²⁰

Three examples of the links between childhood experience and adult biomedical health are the relationship of ACE Score to liver disease²¹ (FIG 4A), chronic obstructive pulmonary disease or COPD²² (FIG 4B), and coronary artery disease or CAD²³ (FIG 4C). The data for CAD show the effect of ACE Score after correcting for, or in the absence of, the conventional risk factors for coronary disease like hyperlipidemia, smoking, etc.



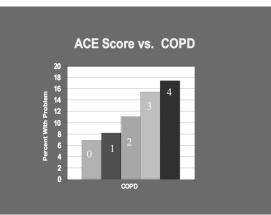


Fig 4A Fig 4B

Physica Sexual:	l abuse 1.5	X	
Cornel			
Sexual	abuse 1.4		
Domest	ic violence 1.4		
Mental	illness 1.4	x	
Substar	ice abuse 1.3	x	
Househ	old criminal 1.7	\mathbf{x}	
Emotio	nal neglect 1.3	x	
Physica	l neglect 1.4	x	

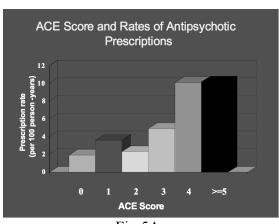
Fig 4C

Certain of these relationships of childhood experience to later biomedical disease might initially be thought to be straight-forward, for instance assuming that COPD or CAD are merely the obvious outcomes of cigarette smoking. In this case, one might reasonably assume that the total relationship of adverse childhood experience to later biomedical disease lies in the observation that stressful early life experience leads to a coping behavior like smoking, which becomes the mechanism of biomedical damage. While this hypothesis is true, it is incomplete; the actual situation is more complex. For instance, in an analysis published in *Circulation*²³, we found that there was a strong relationship of ACE Score to coronary disease, *after* correcting for all the conventional risk factors like smoking, cholesterol, etc. This illustrates the likelihood that adverse experiences in childhood are related to adult disease by two broad etiologic mechanisms:

- conventional risk factors that actually are attempts at self-help through the use of agents like nicotine with its broad psychoactive benefits, in addition to its now well-recognized cardiovascular risks, and
- the effects of chronic stress as mediated through the mechanisms of chronic hypercortisolemia, inflammatory cytokines, and other stress responses on the developing brain and body systems, and sometimes by the total dysregulation of the stress response.

Healthcare costs

Thus far, at the ten-year point in the prospective arm of the Study, we have only begun to analyze pharmacy data. Given the average age of our cohort, we are looking at prescription drug use fifty to sixty years after the fact. Prescription costs are now an increasingly significant portion of rapidly rising national healthcare expenditures in the United States. The relationship of ACE Score to antidepressant prescription rates has already been shown in FIG 1C. Below, in FIG 5A and FIG 5B, we show the relationship of adverse childhood experiences to the decades-later use of anti-psychotic and anxiolytic medications¹⁰. Analyses of the relationships of ACE Score to doctor office visits, Emergency Department visits, hospitalization, and death are in progress. The economic effect of FIG 1E will be intuitively obvious to practicing physicians who have observed that multi-volume patients typically do not have a unifying diagnosis underlying all the medical attention. Rather, they have a multiplicity of symptoms: illness, but not disease. Kirkengen has more fully discussed the nature, origins, and often unwitting medical creation of this complex phenomenon in her book, *Inscribed Bodies*.²⁴



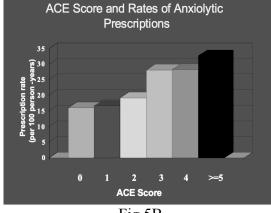


Fig 5A

Fig 5B

Life expectancy

Although we have not yet begun our prospective analysis of adult death rates as they may be related to adverse childhood experiences, a suggestive insight can be provided by use of the Null Hypothesis. Using the Null Hypothesis, we might propose that if there is *no* relationship of ACE Score to ultimate mortality, then we ought to be able to predict certain expected findings and consequently test for them. Thus, if there is no relationship of ACE Score to adult mortality, the age distribution of Kaiser Health

Plan members choosing to come in for comprehensive medical evaluation ought to be independent of ACE Score. In FIG 6A, we see that the age distribution for ACE Score 0 individuals is what one would expect: old people are more likely to come in for a complete medical evaluation than are young people, and intermediate age quartiles have the expected relative proportionality. However, at ACE Score 2, what had been the most common age quartile has become the least common, and what had been the least common has become the most common. At ACE Score 4, the initially most common age quartile has almost disappeared. We anticipate that, when our prospective analysis of death rates is completed, it may illustrate convincingly that there is an increasing death rate as the ACE Score increases. Certainly, this would be the expected continuation of our findings that ACE Score is strongly related first to health risks, then to disease, then to one outcome of disease: death.

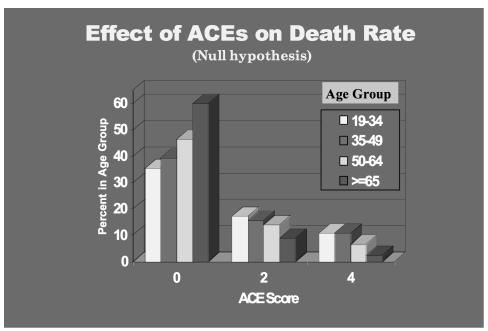


Fig 6A

Reasonably, one might challenge this interpretation of selective attrition by hypothesizing that our patients are progressively so humiliated by exposure of their increasing ACE Scores that they are subsequently avoiding necessary medical care. Such an hypothesis is not supported by our findings. Some years ago we had on site for six months a fully trained psychiatrist-psychoanalyst who saw selected high ACE Score patients immediately after their comprehensive medical evaluation, rather than after referral to Psychiatry. An anonymous questionnaire, returned by 81% of the patients he saw, showed that his hour-long interview was interpreted by patients as highly desirable and appreciated. Talking about the worst secret of one's life with an experienced person, being understood, and coming away feeling still accepted as a human being, seems to be remarkably important and beneficial, perhaps not unlike Confession in the Roman Catholic Church. Indeed, seeking understanding and acceptance is a deep and human need, harkening back at least to the biblical enjoinder, "With all thy getting, get thee understanding."

Integration into clinical practice:

We have made a limited but instructive attempt to integrate the ACE Study findings into clinical practice. At Kaiser Permanente's high-volume Department of Preventive Medicine in San Diego, we have used what we learned to expand radically the nature of our Review of Systems (ROS) questionnaire. We have now asked routinely of over 440,000 adult individuals undergoing our comprehensive medical evaluation a number of questions of newly discovered relevance, the following of which are a sample:

- Have you ever been a combat soldier?
- Have you ever lived in a war zone?
- Have you been physically abused as a child?
- Have you been sexually molested as a child or adolescent?
- Have you ever been raped?
- Who in your family has been murdered?
- Who in your family has had a nervous breakdown?
- Who in your family has been a suicide?
- Who in your family has been alcoholic or a drug user?

Such questions have been accepted by patients in the context of a well-devised ROS questionnaire that is filled out at home. Examiners have learned that the most productive response to a Yes answer is, "I see that you have - - - . Tell me how that has affected you later in your life." While not a simple transition for staff, and one requiring an organized training effort, it has been an effective transition with measured benefits. An independent organization carried out for us a neural net analysis of the data from 120,000 patient evaluations (2 years' work) using this new approach, a biopsychosocial approach to comprehensive medical evaluation. Surprisingly, a 35% reduction in doctor office visits (DOVs) was found in the year subsequent to evaluation, compared to the year before. Additionally, analysis showed an 11% reduction in Emergency Department (ED) visits and a 3% reduction in hospitalizations. This change was dramatically and unexpectedly different from a much smaller, 700-patient evaluation carried out 20 years earlier, when we worked in a more usual biomedical mode. That earlier approach provided a net 11% reduction in DOVs compared to the antecedent year, in spite of a 14% referral rate. No evaluation was made then of ED visits or hospitalization. Finally, we found that the recent notable reductions in DOVs and ED visits totally disappeared in the second year after comprehensive evaluation, when there was a reversion to prior baseline. While the underlying information was present in charts with laser-printed clarity, it was almost never integrated into subsequent medical visits.

If these first year results are replicable, and we believe they should be, the implications for primary medical care are those of a paradigm shift. While offering tremendous opportunity, paradigm shifts are resisted. The philosopher, Eric Hoffer, has discussed this problem in his book, *The Ordeal of Change*²⁵. Jeffrey Masson, in <u>Assault on Truth</u>²⁶ describes the enormous social pressures on Freud to recant his interpretation of his findings of traumatic sexual experiences in childhood as being valid. Louise DeSalvo points out in *Virginia Woolf*²⁷ how literary commentators almost uniformly

avoid the thematic content (incest) of Woolf's work in favor of erudite discussions of her style and literary techniques.

If the treatment implications of what we found in the ACE Study are far-reaching, the problems of integrating this information into clinical practice are absolutely daunting. Simply put, it is easier for all of us to deal with the presenting symptom of the moment than to attempt to understand it in the full context of the patient. Though the latter approach demonstrably would save time and money in the long run, most of us operate in the short run, and respond to valid forces that are both external and internal.

The very nature of the material in the ACE Study is such as to make most of us uncomfortable. Why would a physician want to leave the familiarity of traditional biomedical disease and enter this area of threatening uncertainty that none of us have been trained to deal with? As physicians, we typically focus our attention on tertiary consequences, far downstream, while the primary causes are well protected by time, social convention, and taboo. We physicians have often limited ourselves to the smallest part of the problem, that part in which we are erudite and comfortable as mere prescribers of medication or users of impressive technologies. Thus, although the ACE Study and its fifty-some publications have generated significant intellectual interest in North America and Europe during the past ten years, its findings have yet to be translated into significant clinical or social action. The reasons for this are important to consider if this information is to be converted into meaningful social and medical opportunity.

Conclusion:

The influence of childhood experience, including often-unrecognized traumatic events, is biomedically, socially, and psychologically as powerful as psychoanalysts originally described it to be. This influence is long-lasting, and neurobiologists are now describing its intermediary mechanisms. Unfortunately, and in spite of this major advance, psychodynamic and biological psychiatry remain at odds, rather than taking advantage of the new discoveries to reinforce each other.

Many of our most intractable public health problems are the result of attempted personal solutions to problems caused by traumatic childhood experiences, which are lost in time and concealed by shame, secrecy, and social taboo against the exploration of certain topics. The findings of the Adverse Childhood Experiences (ACE) Study provide a credible basis for a new paradigm of primary care medical practice that would start with comprehensive biopsychosocial evaluation of all patients at the outset of ongoing medical care. We have demonstrated that this approach is acceptable to patients, affordable, and beneficial in multiple ways. The potential gain is huge. So too is the likelihood of physician and institutional resistance to this change. Actualizing the benefits of this paradigm shift will depend on first identifying and resolving the various bases for resistance to it. In reality, this will require far more planning than would be needed to introduce a purely intellectual or technical advance. However, our experience suggests that it can be done.

References

- 1. Eliot, TS. Four Quartets. Harcourt, Brace, and World. New York, 1943.
- 2. Breuer J, Freud S. *Studies on Hysteria*. Standard Edition of the Complete Psychological Works of Sigmund Freud; v2. Hogarth Press. London, 1943.
- 3. Herman JL. *Trauma and Recovery*. Basic Books. New York, 1992
- 4. Harlow HF. *Learning to Love*. Jason Aronson. New York, 1974
- 5. Bowlby J. A Secure Base: Parent-child attachment and healthy human development. Basic Books. New York, 1988.
- 6. Felitti VJ, Anda RF, Nordenberg D, Williamson DF, Spitz AM, Edwards V, Koss MP, Marks JS. The relationship of adult health status to childhood abuse and household dysfunction. <u>American Journal of Preventive Medicine</u>. 1998; 14:245-258.
- 7. Dong M, Anda RF, Felitti, VJ, Dube SR, Williamson DF, Thompson TJ, Loo CM. Giles WH. The interrelatedness of multiple forms of childhood abuse, neglect, and household dysfunction. <u>Child Abuse and Neglect.</u> 2004; 28(7):771-784.
- 8. Anda RF, Whitfield CL, Felitti VJ, Chapman D, Edwards VJ, Dube SR, Williamson DF. Alcohol-impaired parents and adverse childhood experiences: the risk of depression and alcoholism during adulthood. <u>Journal of Psychiatric Services</u> 2002; 53:1001-1009.
- 9. Dube SR, Anda RF, Felitti VJ, Chapman D, Williamson DF, Giles WH. Childhood abuse, household dysfunction and the risk of attempted suicide throughout the life span: Findings from the Adverse Childhood Experiences Study. Journal of the American Medical Association. 2001; 286:3089-3096.
- 10. Anda RF, Brown DW, Felitti VJ, Bremner JD, Dube SR, Giles WH. The Relationship of Adverse Childhood Experiences to Rates of Prescribed Psychotropic Medications in Adulthood. <u>American Journal of Preventive Medicine.</u> 2007; 32(5):389-94.
- 11. Seeman T, McEwen B, Rowe J, Singer B. Allostatic load as a marker of cumulative biological risk. <u>Proceedings of the National Academy of Sciences</u>. 2001; 98: 4770-4775.
- 12. Whitfield CL, Dube SR, Felitti VJ, Anda RF. Adverse Childhood Experiences and Subsequent Hallucinations. <u>Child Abuse & Neglect.</u> 2005; 29(7) 797-810.

- 13. Anda RF, Croft JB, Felitti VJ, Nordenberg D, Giles WH, Williamson DF, Giovino GA. Adverse childhood experiences and smoking during adolescence and adulthood. <u>Journal of the American Medical Association</u>. 1999; 282:1652-1658
- 14. Edwards VJ, Anda RF, Gu D, Dube SR, Felitti VJ. Adverse childhood experiences and smoking persistence in adults with smoking-related symptoms and illness. The Permanente Journal. 2007; 11: 5-13.
- 15. Dube SR, Miller JW, Brown DW, Giles WH, Felitti VJ, Dong M, Anda RF. Adverse Childhood Experiences and the Association with Ever Using Alcohol and Initiating Alcohol Use During Adolescence. <u>Journal of Adolescent Health.</u> 2006; 38: 444.e1-10.
- 16. Dube SR, Anda RF, Felitti VJ, Chapman DP, Giles WH. Childhood Abuse, Neglect, and Household Dysfunction and the Risk of Illicit Drug Use: The Adverse Childhood Experiences Study. <u>Pediatrics</u>. 2003; 111:564-572.
- 17. Felitti VJ. Ursprünge des Suchtverhaltens Evidenzen aus einer Studie zu belastenden Kindheitserfahrungen . <u>Praxis der Kinderpsychologie und Kinderpsychiatrie</u>. 2003; 52:547-559.
- 18. Hillis SD, Anda RF, Dube SR, Felitti VJ, Marchbanks PA, Marks JS. The Association Between Adolescent Pregnancy, Long-Term Psychosocial Outcomes, and Fetal Death. Pediatrics. 2004; 113:320-7.
- 19. Anda RF, Fleisher VI, Felitti VJ, Edwards VJ, Whitfield CL, Dube SR, Williamson DF. Childhood Abuse, Household Dysfunction and Indicators of Impaired Worker Performance in Adulthood. <u>The Permanente Journal.</u> 2004; 8:30-38.
- 20. Dube SR, Felitti VJ, Dong M, Giles WH, Anda RF. The Impact of Adverse Childhood Experiences on Health Problems: Evidence from Four Birth Cohorts Dating Back to 1900. <u>Preventive Medicine</u>. 2003; 37: 268-77.
- 21. Dong M, Dube SR, Felitti VJ, Giles WH, Anda RF. Adverse Childhood Experiences and Self-reported Liver Disease: New Insights into a Causal Pathway. <u>Archives of Internal Medicine</u>, 2003; 163:1949-1956.
- 22. Anda RF, Brown DW, Dube SR, Bremner JD, Felitti VJ, Giles WH. The Relationship of Adverse Childhood Experiences to the Prevalence, Incidence of Hospitalization, and Rates of Prescription Drug Use of Obstructive Pulmonary Disease in a Cohort of Adults. (in preparation)

- 23. Dong M, Giles WH, Felitti VJ, Dube, SR, Williams JE, Chapman DP, Anda RF. Insights into causal pathways for ischemic heart disease: Adverse Childhood Experiences Study. <u>Circulation</u>. 2004; 110:1761-1766.
- 24. Kirkengen, AL. *Inscribed Bodies*. Kluwer Academic Publishers. Dordrecht. 2001.
- 25. Hoffer, Eric. The Ordeal of Change. Harper and Row. New York, 1959.
- 26. Masson JM. Assault on Truth. Farrar, Straus, and Giroux. New York, 1984.
- 27. DeSalvo, Louise. Virginia Woolf: The impact of childhood sexual abuse on her life and work. Beacon Press. Boston, 1989.