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NONSELF HELP: How Immunology Might Reframe the Enlightenment

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Tat tvam asi
[That art Thou]

—Chandogya Upanishad 6.8.7

I. ILLOGICAL "SELF"

In 1960, Peter Medawar and F. MacFarlane Burnet were awarded the Nobel Prize for Medicine for their groundbreaking work on immunity and the role of "self" and "nonself" in maintaining and destroying organic integrity. Medawar's ideas on acquired immunologic tolerance developed from his demonstration that mice "learned" to accept foreign tissue if injected with allogenic bone marrow at or before birth—that is, that one could tolerate difference until defensive intolerance was acquired.

Burnet's contribution was that the body produced the protective antibody ("defender" of "Self"—that is *anti*—foreign *body*) for recognizing a foreign invading antigen ("Nonself"—that is, *anti*body *generator*); and that in so doing it failed to recognize "self." To put it simply, one's defense mechanism worked by not recognizing the self it protected.

By the early-1970s the idea that there was something called an "immune system" was just beginning to take root in general practice, having only first appeared in the scientific literature in the mid-1960s (Moulin 1989:221–222; 1991), which

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is why, I would argue, its recent history is so important. This change was subtle but significant; for the science of immunology moved away from an earlier, reactive definition of itself that went far back into the 19th century to one now characterized by protection through expulsion—an earlier belief that dramatic, hypersensitive reactions (such as in anaphylactic shock) were "the unavoidable preliminary step in the production of immunity" (Moulin 1989:232), now gave way to the idea that immunity once acquired stood principally as a mechanism of defense and boundary maintenance.

Immunology courses at medical schools have, since then, borne such titles as "Identity: Microbes and Defense" As one leading introductory textbook puts it:

While normally acquired immunity is carefully regulated so that it is not induced against components of "self," for various reasons, when this regulation is defective, an immune response against "self" is mounted. This type of immune response is termed *autoimmunity*. . . .

In many cases, exposure to foreign substances results from clinical situations in which tissue is transplanted or blood is transfused from one person to another.... Rejection of the transplant or transfusion is not a manifestation of some force of nature designed to frustrate the physician and the patient. Rather, such rejection occurs because of the central tenet of acquired immunity recognition and elimination of "not-self." [Benjamini and Leskowitz 1988:10]

For several decades, the work of the early pioneers of this model defined the rapidly growing field of immunology, while inspiring our present understanding of transplantation biology. In the absence of a fully articulated and wellunderstood immune system (which, to our surprise when we now look back, first appeared in the literature only in the 1960s) no one could foresee that the human "self" was now defended by a mechanism that could only function by not "knowing," or otherwise "recognizing," the very "self" it was designed to look after. Indeed, this may be immunology's first and ultimate paradox of identity namely, how can a mechanism defend by not identifying the thing, the person, defended?

In successive decades (following Medawar and Burnet), several complex theoretical models emerged: the systemic network theory of Niels Jerne, the associative recognition theory of Melvin Cohn, and the diversified cell theories of Talmage and Burnet, to name some important examples. Through these and others, the systemic nature of immune function became solidified, explaining the idea of immunologic "memory" and the logarithmic rate of antibody formation by showing

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how molecules could quickly replicate on binding to an injected antigen. Through evidencing such an orchestrated defense against what was viewed as a pathogenic assault, the concept of the "immune system" became an acknowledged fact, even though yet today its very existence remains based, to quote Melvin Cohn, the Salk Institute's dean of theoretical immunology and head of its Conceptual Immunology Group, "on experimental systems of such great complexity that many interpretations are possible and reproducibility becomes a luxury" (Cohn et al. 1980; see also Cohn 1992, 1997a, 1997b, 1998a, 1998b, 1998c]).

Almost overnight the killing and consuming concepts of pathogenic microbial invasion took root in immunology even though at the time the nature of viruses (which I will get to in a moment) was yet very poorly understood.

Indeed, despite early optimism about the explanatory usefulness of an immune system structured to defend against invaders, the fundamental paradigm of immunology—the recognition and elimination of "nonself"—has yet to resolve the field's major concerns of autoimmunity, transplantation, and tumor immunology (Carosella and Pradeu 2006, 2010; Carosella et al. 2006; Pradeu 2009; Pradeu and Carosella 2006). Indeed, "the self-nonself" model has created so many intractable problems (e.g., Tauber 1991, 1994, 1999, 2000, 2003; Tauber and Chernyak 1991) that theorists and bench researchers alike frequently suggest that it ought to be put to rest (Tauber 1994), or replaced by a model in which antigen-presenting cells (macrophages and dendritic cells) allow the body to become "aware" of viral "danger." This was, and yet remains, the contested view of Polly Matzinger (e.g., Matzinger and Fuchs 1996; Schaffner 1997), one of the most popularly acknowledged (e.g., Pennisi 1996) and most controversial of U.S. immunologists.

There is still a fascinating book to be written specifically about how these models were rapidly transformed between the 1960s and the new millennium (see, e.g., Eichmann 2008)—both about how notions of what viruses were evolved out of the study of microbes, and about how immune systems models related in the 1950s and 1960s to the social information feedback studies of the Macy Foundation being carried out by the father of cybernetics, Norbert Weiner, along with Harvard sociologist, Talcott Parsons, and anthropologist, Gregory Bateson.

However, what is so often overlooked throughout these debates over models of immunity is the degree to which immunological identity hinges on culture-bound notions of a wholly autonomous "self"—a concept made manifest in two basic assumptions: (1) that organic integrity depends on the recognition and elimination of biological difference (there would be no immunology without recognition and cuan_1130

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elimination); and (2) that the notion of a prior and persistent "self" is not contestable. Overall, these premises converge on a singular assumption: that a "self" must preserve its integrity through a protective mechanism. Were a "self" not salient, persistent, and protectionist, that "self" would soon become, as Cohn himself once reminded me, a toxic dumpsite.

Whilst the "recognition-and-elimination" model worked well for autonomous biological pathogens (such as those bacteria and parasites which really do attack us), we now know that defending the body against a viral "attack" is nothing like defending it from invading organisms. Viruses need cells to achieve vitality, and cannot attack without the life that autogenous, "self-made" cells (ones made by our own bodies) bring to each and every viral encounter.

Furthermore, because immunity demands the ongoing and ever-changing acquisition of tolerance to one's own proteins, it is often argued by contemporary immunologists who are dissatisfied with the self-nonself model that if, as we now know, the self "is constantly being defined anew" through defensive antibody production and related acquired immunity, isn't that another way of saying "that it doesn't really exist at all"? (Richardson 1996:5). In other words, if the self is a prior and persistent Cartesian entity, then what we now know about immunity makes its focus on self both misconstrued and inappropriate. Either immunology is not about the self, or, if it is about selfhood, then that self is not Cartesian.

II. A BASIC CONUNDRUM

If so much in immunology depends on what a "self" is, can what we have learned from immunology, to reverse our terms, make possible a rethinking of our time-honored concept of selfhood? Does immunology, to reverse or inquiry, tell us something new regarding our cultural and historical assumptions about identity and what it is that makes for a "self"?

Although good bench science depends on building on existing assumptions, anthropologists can look toward other cultures for models of self and other; for there are many examples (Hinduism being one famous one) in which body image boundary is defined by a careful familiarity with, and a regulation of, new stimuli that may in turn be helpful or harming.

Without providing the extensive inventory of possible models that a full historical and anthropological assessment of these questions would require—a thing, by the way, very much in need of being done—we may and should ask what might be gained by such an inquiry.

Are other notions of "self" applicable in any way to theoretical immunology? To answer this we need briefly to revisit the famous question of whether viruses are living things; for we will see that the virus stands somewhere at the borders of "self" and "nonself," and is thus as much a conditioner and definer of a body's boundaries as it is a "single-minded attacker."

A comparison of an excellent contemporary introduction to cellular biology

(Becker, Reece, and Poenie) and any of the early popular attempts at explaining whether viruses are living things, reveals that our assumptions about the vitality of the virus have changed little over the past 40 years; for despite the fact that viruses have no mobility, we continue to describe them as if they do. Indeed, today's texts put the question aside as intractable:

"The question is sometimes asked whether or not viruses are living. The answer depends crucially on what we mean by "living," and it is probably worth pondering only to the extent that it helps us more fully understand what viruses are—and what they are not. The most fundamental properties of living things are mobility, irritability (perception of, and response to, environmental stimuli), and the ability to reproduce. Viruses clearly do not satisfy the first two criteria. Outside their host cells, viruses are inert and inactive. They can, in fact, be isolated and crystallized almost like a chemical compound. It is only in an appropriate host cell that a virus becomes functional, undergoing a cycle of synthesis and assembly that gives rise to more viruses." [Becker et al. 1996:105]

And because it is intractable, we continue to use volitional, intentional, metaphors that encourage us to think of the virus as a living thing that can commandeer a healthy cell, reduce it to a nursery, and reproduce itself at breakneck speed. But how can a virus do such things if it is inert and without locomotion? How can viruses recognize, scout, trick, discover, alert, evade, sense, recruit, mobilize, prod, mask, defend, scavenge, attack, invade, adapt, appropriate, sacrifice, and kill (Napier 2003b:60) if they lack mobility and do not respond to environmental stimuli? If these are mere linguistic conventions, why should we continue to employ them when they are so inaccurate?

Here is a common example of the consequences of such conceptualizing made famous by Susan Sontag in AIDS and its Metaphors (1990). It will resonate easily with what most of us have been told:

The invader is tiny, about one sixteen-thousandth the size of the head of a pin. . . . Scouts of the body's immune system, large cells called macrophages, sense the presence of the diminutive foreigner and promptly alert the immune

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system. It begins to mobilize an array of cells that, among other things, produce antibodies to deal with the threat. Single-mindedly, the AIDS virus ignores many of the blood cells in its path, evades the rapidly advancing defenders and homes in on the master coordinator of the immune system, a helper T cell....

On the surface of that cell, it finds a receptor into which one of its envelope proteins fits perfectly, like a key in a lock. Docking with the cell, the virus penetrates the cell membrane and is stripped of its protective shell in the process....

The naked AIDS virus converts its RNA into DNA, the master molecule of life. The molecule then penetrates the cell nucleus, inserting itself into a chromosome and takes over part of the cellular machinery, directing it to produce more AIDS viruses. Eventually, overcome by its alien product, the cell swells and dies, releasing a flood of new viruses to attack other cells. [1990:105-107]

Here is another example:

When faced with a foreign invader, the immune system mounts either of two defenses. One, humoral immunity, involves primarily B cells. These white cells recognize a particular antigen, then make antibodies that bind to that molecule. The other depends heavily on T cells . . . that can destroy tumors and cells infected with viruses and bacteria. These assassins, including natural killer cells, become part of the cell-mediated immune response.

T helper cells are the sergeants that roust T or B cells into action. As helpers form in the thymus, each becomes sensitive to just one antigen trigger. They drift in the bloodstream or hang out in lymph nodes in a "naïve" state until they meet the antigen they were primed to recognize. At that moment, a helper cell's fate is sealed as either a TH1 or a TH2, or so some researchers think. If it becomes a TH1, the cell then readies cytolytic T cells to do battle, generating the TH1 response. As a TH2 cell, it *initiates* humoral immunity. . . .

These beneficial assassins can destroy a cell that has been tricked into harboring pathogens where antibodies and TH2 components can't get at them. [Pennisi 1994, emphasis added]

But if, on its own, a virus remains inert and without locomotion, why should we privilege it with agency? Why should we in fact refer to viruses as foreign agents, if a virus is lifeless outside a living cell? And if the floating virus is not an active "other" to be defeated, what generates a so-called "viral attack"? Why, furthermore,

should the body's bringing it to life be understood as primarily a defensive activity, if a virus on its own is inanimate? Finally, if the so-called defensive antibodies we create (and those proteins essential for identify foreign bacteria and viruses) are de facto not coded for marking "self," how can we see them at all as "self"—or a defensive mechanism of "self"?

Might it be that our persistent characterizations of viruses as active agents, arises partly from the cultural belief that harboring otherness within us is principally dangerous, a belief whereby a persistent "self" must in turn always be protected against things "foreign"? Although we all have cancer cells within our bodies, for example, we never say we have cancer until those cells become problematic. In fact, this ambivalence about, and concern over, things foreign within us is why, until the arrival of stem cell research, studying the maternal—fetal was so often unsatisfying for immunologists (Charlesworth et al. 1989).

To put things in perspective in an era of research now dominated by attention to "uncommitted" embryonic stem cells, it is worth listening to a well-known Australian immunologist's view of the maternal—fetal only a few decades ago. In 1975—76, the Nuffiled Foundation had supported the distinguished Australian immunologist, Graham Mitchell, in what was hoped to be new, groundbreaking work in this area. But this kind of immunology was unsatisfying for most immunologists whose training emerged from the understanding of the invasive agents of bacteriology and parasitology.

As Mitchell put it, commenting on this period of his research in the 1970s:

The maternal/foetal was a good idea, but we just didn't have the techniques, just didn't have the way of approaching the question. It's something which a lot of people have actually got into for a short time and then got out of—the immunological aspects of nature's most successful foreign graft [i.e., pregnancy]—how the foetus actually survives inside the immunologically hostile mother. We know she is responding to antigens from the foetus. So we did a little bit on that and then got a couple of publications. But that was a bad year. We had the idea and we thought, wouldn't it be nice to have got some money for it, and then we were under pressure to deliver on the money we got from the Nuffield Foundation. . . . We weren't into the field enough and I must say we really didn't get into the field because I got absolutely seduced by host/parasite relationships. [Charlesworth et al.:214]

As a group of social scientists then studying Mitchell's work pointed out, it is clear that

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Mitchell's own background, experience and contacts fitted him better for research into the immunology of host/parasite relationships. He had no special background in reproductive physiology [most immunologists didn't prior to the era of stem cell research] and was unfamiliar with key techniques needed in the foetal/maternal studies—as he [Mitchell] put it, "we were not particularly good at tissue culture." But he was *au fait* with the basics of parasitology: "I didn't [Mitchell claimed] have the hangup, which a lot of students do have, of learning the list of fifty parasites and remembering the life cycles and remembering what host they parasitize and so on. . . . That was behind me as an undergraduate. A little bit of boning up and that's all I needed for that." [Charlesworth et al.:214—215]

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But Mitchell was far from alone at that time in being unsettled by the areas of embryonic cell research that now dominate the science news media; embryology had since World War II faded in importance as microbiology grew dramatically, leaving immunology and virology to evolve alongside and out of the studies of invasive agents and infectious diseases. Why *do* autoimmune symptoms often subside in pregnancy? Why do woman have much higher rates of autoimmune infection (as much a seven or eight times) than men? Might even childbirth itself, as immunologists often think, be understood in terms of immunological rejection? These are today's question more than yesterday's.

23 24 But given them (perhaps precisely because of them), we should, I would argue, persist in revisiting our earlier question: how can viral antigens be considered *foreign* invaders if our own cells animate viruses? In Darwinian evolution there is no reason for a healthy organism to seek out something that might harm itself, except when in retrospect we call that encounter conditioning. And, because organisms do not spontaneously endanger themselves in a "naturally selective" world, viruses must at some level, it is assumed, be aggressive killers, even if they are actually inert until brought to life by a cell.

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There is, however, another possible view—namely, that antibodies are a key to exploring the dividing line between self and other, a mechanism by which one's identity, as a living thing, becomes contested and eventually defined. Identity, in this sense remains—to the considerable relief of immunologists—a prior and persistent condition, but one that is now defined at the peripheries of selfhood where it is contested and challenged. If we can accept such a view of identity (one widely evidenced in the anthropological literature), there is only one logically consistent and satisfying conclusion to be drawn: namely, if viruses need cells to

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come alive, the so-called immune "self" (our antibodies) must attract, or otherwise absorb those potentially pathological viral antigens. We know, that is, who we are by the risky, dynamic, and sometimes painful process of discovering what we can and cannot accommodate. "If you can't stand the heat," as Richard Nixon was so fond of repeating ad nauseam, "get out of the kitchen"—survival, in this view, being predicated on stepping forward, "taking the heat," and hopefully doing something with it besides caving in.

Although a completely different paradigm, the notion that antibodies absorb potentially harmful bits of inert matter to transform (for better or worse) their packaging of information actually makes good sense and withstands all logical trials we might set against immunology's major conundrums; for an antibody is the tool that enables a living thing to explore the boundaries of life—to engage that danger that is the precondition of real change—whether that change be life giving or life taking. Although many may find the concept of inviting danger intolerable, in some non-Western notions of selfhood it is considered both logical and coherent. There are multiple anthropological examples of how self is made up and defined by potentially dangerous encounters at one's boundaries.²

III. NONSELF HELP: COMPATIBILITY RECONSIDERED

I have suggested how an alternative view of viral activation—one arising from within cotemporary immunology—can reframe immunology's former "paradox of identity"; but what of its other more vexing paradoxes? Can this assimilative view of viral life also reframe them? Let us briefly apply such an interpretation to two other outstanding and famous paradoxes of theoretical immunology.

First, the Evolutionary Paradox (Silverstein 1989): How can the human body continue to create a broad specificity repertoire—a wide range of "B"- and "T"-cells—when natural selection would otherwise demand that apparently superfluous "deformations" be eliminated?

The answer is that, in an anthropologically and historically informed model, natural selection is no longer a problem because the body is creating possible versions of "otherness" rather than mutations of "self"—that is, the bone marrow ("B") and thymus ("T") cells explore "otherness" rather than defend "self"; for the human body appears to produce an extraordinary number of antigen-inducing antibodies (some threatening, some harmless). Indeed, estimates of the human antibody repertoire run from 10 to the 5th to 10 to the 16th. And if these figures weren't already mind-boggling, "because most antigens have many epitopes [i.e., amino acids or sugar residues that are antigenic—'outside'—determinants] and a NONSELF HELP

given epitope can be recognized by more than one lymphocyte [response cells in the lymph nodes], the number of lymphocytes that can respond to a given antigen is much larger than the number of cells possessing a certain antigen receptor" (Becker et al. 1996:788).

According to natural selection, these numbers should be naturally limited. Superfluous creations should be eliminated. Bodies don't, for instance, spontaneously generate organic variations in anticipation of unanticipated events. In evolution the body responds and adjust to stimuli; it does not create spontaneous mutants, or at least does not do so as a function of health-enhancing activity. Saltationsudden and unexpected change—troubled Darwin; it was thought unnatural.

As, however, cellular diversity by definition proliferates in immunology, this so-called "immunological repertoire" appears "paradoxical" to immunologists when it creates mutants that have no apparent target. A body should not produce spontaneous, experimental deformities in anticipation of a single cellular mutation out of millions on millions that might be useful for some future, unknown threat. Furthermore, for binding to take place, an antibody must be created with a locking mechanism that works for an unknown antigen. This makes no sense either because variation, by definition, is infinite—unless immunologists, as have some evolutionary biologists, come to accept the possibility that antigens develop from the nucleic acids of normal cell genes (i.e., out of the attenuated nucleic acids of [as it were] "former selves," or at least something very much like them).

If this connection between pathology and prior normality is posited, what of antibodies—those mutations we generate in anticipation? Might they not be seen, then, as creative attempts to engage risk at the borders of self? In numerous non-Western models identity depends on just such a dynamic engagement. However, I refrain from making the complex digressions that such an explication would necessitate, but refer to the many examples of spiritual engagement, ritual warfare, and marital customs described in the anthropological literature wherein the assimilation—the vaccination—of "otherness" proliferates. Although elsewhere I discuss alternate concepts of self at some length (e.g., Napier 1986, 1992:190-99; 2003b), suffice it here to say that the anthropological literature is replete with examples not only of how body-image boundaries may become transformed or otherwise malleable but also where they position themselves to assimilate other selves or, as it were, evolutionary "former selves," as a means of establishing and negotiating identity and body boundaries.

If natural selection is the parent of traditional immunological identity, cell biology shows that there is nothing natural about selection, at least as it relates

to the benefits and drawback of encountering danger; for a self understood as a "prior and persistent identity" can never ascertain in advance whether a selective pressure is ultimately destructive or constructive. (Which is why, incidentally, natural selection is so frequently accused of being teleological.) Therefore such a "self" can only constantly attempt, as it were "selfishly," to eliminate difference. The result of this elimination may reflect the immunology we have inherited, but it will not answer the contradictions of the field now being taken on by contemporary immunology—contradictions whose resolutions shake the very foundations of Cartesian thinking about what makes a "self" prior, persistent, and autonomous.

So much for the evolutionary paradox and immunology. But there is another, equally famous, outstanding paradox that an assimilative view of viral encounters resolves. This is what is known as the *Repertoire Paradox* (Silverstein 1989). And it goes like this: *How can we argue that the proliferation of B-cells and T-cells is adaptive, when neither the size of the immunological repertoire, nor even the presence or absence of the thymus (our major generator of immunity), can be construed as an indicator of resistance? Some organisms do very well indeed with a quite limited repertoire; and it is widely accepted that repertoire scope is only relevant to the number of potentially pathogenic viruses that can influence an organism: one may not need a thymus at all if there is no danger—no alienated variation of identity—no "similar," "other-," or "former" self, as it were—being expressed.*

Furthermore, "foreign" viruses and prokaryotes (those organisms whose DNA is not contained inside a cellular nucleus), are not always harmful: some are innocuous, others are real mutation—transformation factories without a clearly defined "target," undermining the very notion that the body seeks always to rid itself of difference.

But identifying "nonself" need not be defensive in this new, immunologically, and anthropologically informed model of "self"; for immunity can now be understood more as a creative attempt to engage difference, than solely as a battle to eliminate it. Here, immunity is less a system of protection and defense than a system of information assessment, even of creation—for we survive through risking engagement with difference—as so many new students of mirror neurons, for instance, are eager to tell us.

Today, then, immunology sees itself quite differently, and I would argue is well-positioned—perhaps better positioned than any other domain of modern science—to help us rethink notions of the self that have dominated Western philosophy at least since the Enlightenment; for it is with the Enlightenment that we get the birth of scientific agnosticism and the first consistent argument for a prior and persistent person that is not predicated on divine intervention.³

disorders.

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Compare the "recognition and elimination" model of immunity with, for instance, a recent Salk Institute discovery that certain skin cells with a hereditary blood disease called *Fanconi anaemia* could be reengineered to revert to stem cells, which could then be recommitted by use of a virus. What this means is that the repaired cells—constructed by reverse engineering to a naive stem cell and then as it were, reinfecting it—can now be used to replace the defective bone marrow cells characteristic of a disease that also results in very low blood cell counts. A cure becomes possible because, rather than suppressing immune responses, we reshape them by encouraging and feeding novel viral information—information of a new

What made this possible, I would argue, is an unrecognized but giant leap that has almost subconsciously become a core immunological precept—namely that viruses do not invade us, but that we, for better or worse, bring life to the sometimes dangerous encounters that define the limits of who we are, that limit what we can be, and that (hopefully) do so without taking the very life that those viruses, once embodied, now inform—or, as we used to say, infect.

type that one day may well lead to therapies for what were once incurable genetic

Burnet's antibodies, which recognize "foreignness" ("nonself") but fail to recognize "self," are now replaced by antibodies that function as "self" search engines—search engines for the information (harmful or helpful) that sits latently in viruses like books in a library.

If, as evolutionary biologists now widely suppose, pathogens evolve from normal cell genes or entities much like genes (for how else can they bind with cell surface receptors?), what our antibodies must be recognizing are as much aspects of *selfhood* as aspects of *foreignness*. The difference made possible by this view is neither subtle nor semantic; for it shows us that cellular antibodies risk bringing life to alien viruses to revitalize these biologically binding—and therefore related, even if dangerous—forms of "self" so as to adjust and respond to them. In this view the "self" stands not only as something prior and persistent but also as a living thing capable of constantly recreating and reclaiming its identity through engaging viral information.

"Know your friends well, you enemies better." Thus understood, the body interacts with its environment in an attempt to create a new future. Burnet, who said "it could be no other way" than for antibodies to fail in recognizing "self," did not suspect that there was another way of resolving immunology's paradoxes—a way that might change how we view the self and its environment. Part of this not knowing was because of immunology's evolving out of cell biology—where real

organisms do attack one another. Part of this was because we needed molecular genetics and stem cell research to demonstrate the real advantages of modifying and loading cells over killing them off.

The alternative view outlined here, although not the last word on the immunological self, does provide an explanation by which immunology's paradoxes can be reconciled with what is known. If proven correct, it may or may not help reshape scientific research, as it, however, seems now at the threshold of so doing; but it will certainly force us to reconsider immunology's contribution to the metaphysics of human identity.

ABSTRACT

The classical immunological paradigm is predicated on the body's ability to recognize and eliminate "nonself." However, the "self-nonself" model has yet to facilitate any resolution of the field's major concerns, and may thus prove to be of limited use. Merely discarding it is no solution, as the juxtaposition of "self" and "nonself" persists in research, in clinical settings, and in everyday practice despite the best efforts of theoretical immunologists. Instead, the very conception of "selfhood" may prove to be key. Replacing immunology's prior and persistent "self" with less static concepts derived from non-Western contexts not only resolves immunology's famous paradoxes but also offers a new and more accurate model that allows immunology to reframe what may become an outmoded Enlightenment construct of "self." In such a new paradigm, immunology's well-known system of protection and defense is replaced with a view in which nonself becomes not only the body's enemy but also its primary mechanism for the creative assimilation of difference. This incorporative model—in which the "immune system" functions more as a search engine than as an expeller of difference—both resolves outstanding paradoxes, and complies more accurately with contemporary knowledge and research practice. [medical anthropology, immunology, identity]

NOTES

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For further discussion see The Age of Immunology: Conceiving a Future in an Alienating World (Napier 2003b). See also Cohen 2004, 2009; Martin 1990, 1994; Moulin 1989, 1990; and Napier 1996a, 1996b.

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2.	See several examples in Napier 2003a, 2003b. On the idea that identity is defined at its
	peripheries, see Fredrik Barth's landmark Ethnic Groups and Boundaries: The Social Organization
	of Cultural Difference (1969).

More specifically, I say "Enlightenment" because Descartes begins his own Meditations with the claim that he is about to embark on his famous exploration of (self-) consciousness, and that (like a modern-day "infectious" virus) he comes forth masked.

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Editors' Notes: Cultural Anthropology has published many articles on the history and anthropology of science, including, Mette N. Svendsen's "Articulating Potentiality:

Notes on the Delineation of the Blank Figure in Human Embryonic Stem Cell Research" (2011), Celia Lowe's "Viral Clouds: Becoming H5N1 in Indonesia, Michael M. J. Fischer's "Four Genealogies for a Recombinant Anthropology of Science and Technology" (2007), and Deepa S. Reddy's "Good Gifts for the

Common Good: Blood and Bioethics in the Market of Genetic Research" (2007).

Cultural Anthropology has also published articles on conceptions of self and personhood. See, for example, Arafaat A. Valiani's "Physical Training, Ethical Discipline, and Creative Violence: Zones of Self-Mastery in the Hindu Nationalist Movement" (2010), Tomas Matza's "Moscow's Echo: Technologies of the Self, Publics, and Politics on the Russian Talk Show" (2009), Nancy Ries's "Potato Ontology: Surviving Postsocialism in Russia" (2009), and Julie Livingston's "Suicide, Risk, and Investment in the Heart of the African Miracle" (2009).