

## **RECEPTOL**

### **RECEPTOL PRODUCT BRIEF**

The Receptol® oral spray consisting of Radha 108 Nanopeptides, stimulates body's own immune system as a broadspectrum immunomodulator & antiviral to fight against several diseases & prevents all communicable infections. It is a natural product manufactured by Nanobiotechnology patented proprietary processes from bovine colostrum (1st Milk after Birth).

**Keywords:** Colostrum, Receptol, Nanopeptide, Radha108, immune-modulator

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Our health is directly influenced by our immune system. A balanced and healthy immune system is centre to the body's ability to defend against infections. "It is our ability to create a healthy immune system that represents the greatest potential for gains in human health".

Today, however, many factors contribute to the general weakening of the body's defences. Antibiotics have begun to fail as the resistance of many infectious strains multiplies. Due to the failure of government control of health codes, deterioration of water quality, and frequent international travel diseases now spread more easily than ever before. Fortunately, recent research has uncovered a natural agent, which can increase our ability to fight disease and improve the quality of life for many people.

RECEPTOL® is the name given to this relatively new agent. It is found in colostrum and other sources and is a natural way of strengthening our immune systems against disease. The application of RECEPTOL® is in the medical field of Immuno-therapy which is a quiet revolution taking place in medicine. It is a form of treatment that uses the different aspects of your immune system, it cells and molecules and its various stratagems to tip the balance in your favor as your body battles to maintain health.

RECEPTOL® essentially is a complex cocktail of compounds derived from bovine lacteal sources. These compounds include Proline-Rich Polypeptides (PRPs), glycoprotein, growth factors, neurotransmitters, cytokines (IFN-alpha, IFN-beta) produced from PRPs, and enzymes. Other immune factors present include Trypsin Inhibitors, Glycoconjugates, Orotic Acid, Alpha 1-Antitrypsin, Alpha 1-Fetoprotein, Alpha 2-Macroglobulin, Alpha 2- AP Glycoprotein, C3 &

C4, Orosomucoids, Lyozyme, Thiocyanate, Peroxidase, Xanthine Oxidase, Vitamins A, B12, E, and Sulfur. Each of these compounds has specific functions in the human body.

Anyone-healthy or diseased, with a few exceptions-is benefitted from regular RECEPTOL® supplementation. The use of RECEPTOL® has resulted in no reports of serious adverse reactions, even when clinically administered in doses in excess of normal for prolonged periods. Those with specific ailments also benefit.

Numerous studies have shown the effectiveness of RECEPTOL® in eliminating or alleviating symptoms of herpes, chronic fatigue syndrome, Epstein Barr, hepatitis, secondary infection due to AIDS, Candida, cancer and many other disorders. Studies have also shown continual use provides the greatest benefit, with maximum immune activity occurring 24 to 48 hours after initial dosing.

The need for RECEPTOL® as an adjunct to better health stems from the growing awareness that prevention is the best source of treatment. With the increasing risks of antibiotic resistance and significant health threats, such as SARS, the medical community increasingly turns to the inherent concept of vaccines prevention.

RECEPTOLs are a kin to vaccines. But, rather than expose the patient's immune system to the actual disease or a deactivated version of the same, RECEPTOLs expose the patient's immune system to the memory of a health threat-whether foreign or native-and the knowledge of how to best respond to protect itself.

#### **What is Radha108:**

Radha108 is a natural, immunity enhancing component derived from bovine colostrum. Radha108 is essentially a class of peptides called Nano-peptides with molecular weight of 3500–10000 and Proline Rich Polypeptides (PRPs). Other naturally derived compounds present in the product include: Vitamin A, E, B1, B2, B5, B6, B12, B13, C, Folic acid, Sulphur and Betacarotene.

Though some of the immunity enhancing properties of Proline-Rich Polypeptides were reported earlier, the molecular mechanism and its synergistic action with low molecular weight Nanopeptides were not known until Dr. Pawan Saharan, Chief Scientific Officer and Founder Director, Chaitanya Healthcare, India identified a series of Nanopeptides, also referred to as Nano- Information-Peptides from Bovine Colostrum.

Further study and confirmation of its therapeutic efficacy in a wide range of diseases/disorders was made possible only after these therapeutically effective 'Nanopeptides' were isolated in the pure and concentrated form by using sophisticated nanotechnology and filtration techniques by Biomix Network Ltd.

One of the exciting finding was the excellent efficacy and safety of these Nanopeptides (Radha108) in the treatment of HIV/AIDS. Clinical trials conducted in a major hospital in Mumbai have clearly shown that:

A. Receptol was immensely beneficial in:

- a) Mitigating the symptoms like fever, diarrhea, nausea associated with AIDS
- b) Increasing weight of the patients
- c) Reducing the viral load
- d) Increasing the CD4 count
- e) Minimizing/eliminating associated opportunistic (viral, bacterial & fungal) infections

The observations on Receptol include:

- B. The positive effects could be observed within a span of 12 weeks
- C. No significant adverse effects observed even on prolonged usage..
- D. Can be used even as a monotherapy

#### **Radha108: Mechanism of Action:**

#### **MODE OF ACTION**

- The informational proteins (RADHA108) in RECEPTOL® are active in mitigating cell fusion
- RADHA108 series docks on glycoprotein receptor on the cell surface and thus closing doors and windows for viral entry into the immune cells.
- RADHA108 series get absorbed in the blood stream through buccal mucosa and crosses the Blood Brain Barrier (BBB).
- The levels of Interleukins & Cytokines are increased substantially.
- Supports regulation of thymus via producing functionally active NK cells
- RADHA108 augments cell-mediated immunity & activates T-cell precursors to produce helper & suppresser T-cells increasing CD4/8 counts.
- RADHA108 promotes growth & differentiation of stem cells in response to any disease

Fusion of HIV particles with human white blood cells, particularly CD4 cells occurs with the aid of glycoprotein epitopes on the viral wall. The informational proteins in Radha108 are active in mitigating cell fusion. Radha108 docks on HIV glycoprotein gp120 mimicking receptor on the cell surface and thus closing doors and windows for virus/antigen entry into the immune cells.

Receptol, a pure natural product having informational nano peptides and Proline Rich Polypeptides (PRP), is an effective and safe in increasing weight, general well being of patients, decrease in viral load and increase in CD4 cell count.

RECEPTOL® proteins directly support the Natural Killer (NK) cells of the immune system. Natural Killer Cells provide the front line of defense specially equipped to locate and kill disease cells. NK cells attach to the surfaces of foreign substances or their outer cell wall, and inject a chemical “grenade” (granule) into the interior. Once inside, the granules explode and destroy the foreign invader within five minutes. The NK cell itself remains intact and moves on to destroy the next immune attacker. Many doctors and clinics are finding RECEPTOL® helpful in promoting NK function and activity as well as supporting a healthy immune system for all patients. The immune system plays a great role in the quality of our health. Strong, active and optimally functioning NK cells promote optimal health and deter foreign substances from affecting immune function.

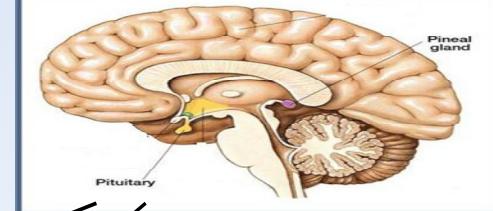
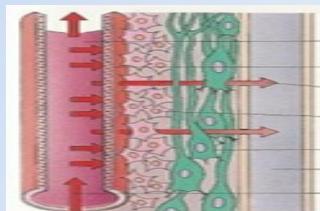
- Radha108 (PRP) promotes differentiation of B cells, differentiation and maturation of macrophages and monocytes.
- Activates natural killer (NK) cells, cytotoxic cells of the innate immune system
- Mitigates cell fusion and docks on HIV glycoprotein like Gp120, 180, 160 and 41 mimicking receptor on the cell surface closing entry of viruses.
- Stimulates production of cytokines IL-1 to IL-11, TNF- $\alpha$ , INF- $\gamma$ .
- Stimulates the maturation of immature thymocytes into either helper or suppressor T cells
- Radha108 also functions as a molecular signaling device which works through receptors on target cell surfaces.
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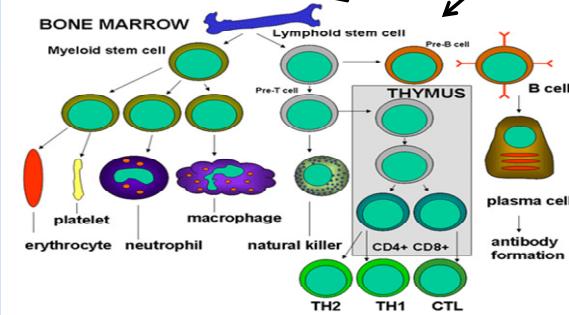
## Pharmacodynamics ( Mode of action ) of PRP (Radha 108)



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### **Phase III Radha108 clinical trials:**

A three months Phase III trial of Radha108 in patients with HIV seropositive status was conducted. After positive results of safety and efficacy Phase I and Phase II trials of Radha108 in the US & Africa, Ministry of Health and Family Welfare, Government of India under the Pharmaceuticals Research and Development Support Fund, funded Phase III trial of its efficacy in HIV/AIDS patients coordinated by National AIDS Control Organization (NACO)/ Indian Council of Medical Research (ICMR)/ Director General Health services (DGHS)/ Drugs Controller General of India (DCGI) in consultation with Dr. Pawan Saharan, Chairman, Biomix Network Limited, India at Lokmanya Tilak Municipal Medical College, Sion, Mumbai during October, 2005 and October, 2006.

An open label accelerated prospective Phase III efficacy study for Radha108 was conducted for 101 HIV sero-positive patients between the ages of 18 and 60 in India 50 patients were enrolled in Radha108 Trial, October 2005 and successively 51 patients were enrolled in Radha108 Trial, October 2006. All the patients were evaluated for a medical history, physical examinations, blood and urine tests including CD4 cell count and HIV viral load as per protocol.

Patients were taught self-administration of Radha108 liquid in pump spray form 6 times a day at 4 hour intervals. Each administration consisted of 4 sprays directly on the buccal mucosa (inner cheek) and was to circulate in the mouth for 60 seconds before swallowing. Each pump of the spray delivered 0.75ml of Radha108 liquid.

Participants were evaluated in the clinic once a week through the first 4 weeks of therapy and then once every 2 week for the remaining 8 weeks of the study. They were evaluated for clinical and physical symptoms via symptom assessment form. Physical examinations, side effects of treatment and compliance to the drug on follow up visit, blood tests to measure CD4 cell counts, HIV viral load, hemoglobin, white blood cell count, liver function test and renal function test were done at baseline and at end of study.

CD4 and CCDS cell count were analyzed via Flow Cytometry and viral load via Polymerase Chain Reaction at National Accreditation Board for Testing and Calibration Laboratory (NABL) and the College of American Pathologists Accredited Metropolis Health Services (I) Pvt. Ltd. Laboratory, Mumbai at the beginning and at the end of 12 weeks of Phase III of Radha108 Trial, October 2005 and at the end of 12 weeks, patients were evaluated at Indian Institute of Hematology (IIH), an ICMR Institute, KEM Hospital, Mumbai for Radha108 Trial, October 2006.

Primary efficacy end points of study were the effect of this therapy on the markers of HIV disease including clinical symptoms and physical findings, weight gain/loss along with the general well being of the patients and HIV viral load and CD4 cell count which showed consistently positive results in both the trials.

## Results of phase III Radha108 Trial - October 2005

There was excellent compliance for Radha108 liquid in pump spray form. The investigational product Radha108 was well tolerated and there were no serious adverse events.

**Weight gain:** Consistent weight gain was observed in all patients on follow up visits, which was one of the primary end points (Table 1). The average weight gain after 12 weeks of treatment was 4.73kg per patients which is statistically highly significant ( $p<0.001$ ).

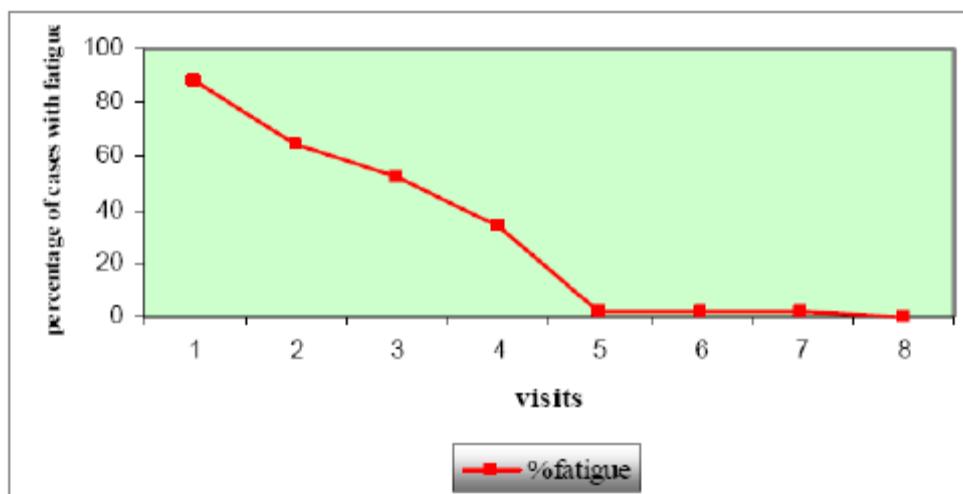
Table 1 : changes in mean weight after treatment

duration in weeks	mean weight (N = 50)
basal	$50.48 \pm 10.97$
2	$50.77 \pm 11.26$
3	$51.38 \pm 10.94$
4	$52.33 \pm 10.73$
6	$53.89 \pm 11.17$
8	$54.59 \pm 10.89$
10	$55.44 \pm 11.07$
12	$55.21 \pm 9.42$

**Clinical parameters:** All the clinical symptoms disappeared during the 12 weeks of treatment. These reductions are statistically significant ( $p<0.05$ ).

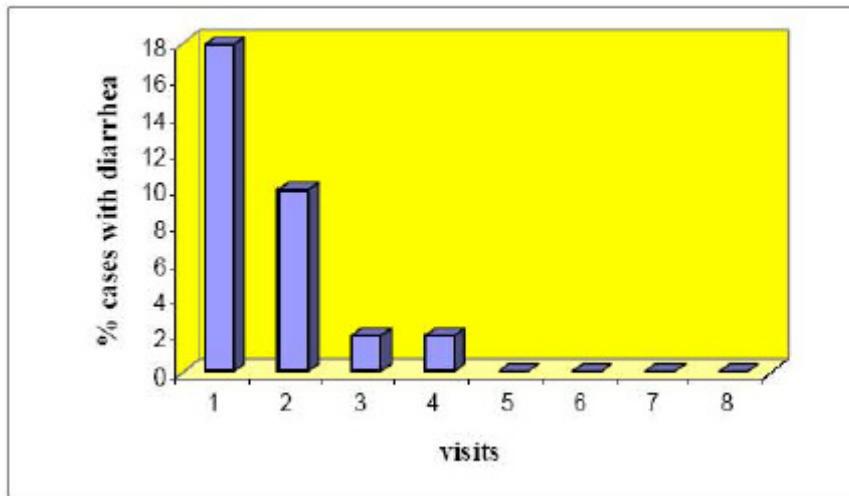
**Fatigue/ Malaise:** 88% of the total study cases had symptoms of fatigue at basal. After treatment at the end of 2<sup>nd</sup> week proportion of symptoms of fatigue had a significant fall from basal. After 6<sup>th</sup> week onwards only one or two patients had fatigue (Figure 1).

Figure 1 : percentage of cases with fatigue / malaise after the treatment



**Diarrhea:** 18% of total study cases had diarrhea at basal and after treatment from 5<sup>th</sup> week onwards all the patients had relief from diarrhea (Figure 2).

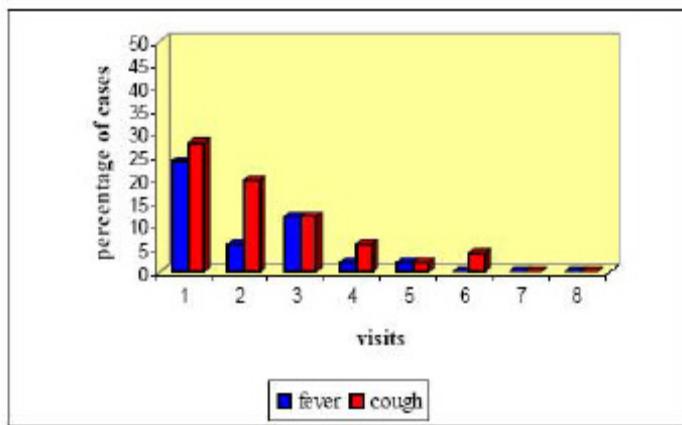
Figure 2 : percentage of cases with diarrhea after the treatment



**Fever:** Fever was reported by 24% of total study cases at basal and from 7<sup>th</sup> week onwards not a single patient had fever with significant fall which started 4<sup>th</sup> week onwards.

**Cough:** 28% of the total study cases had a significant fall from 3<sup>rd</sup> week onwards and all improved 10<sup>th</sup> week onwards (Figure 3).

Figure 3 : percentage of cases with fever / cough after therapy.

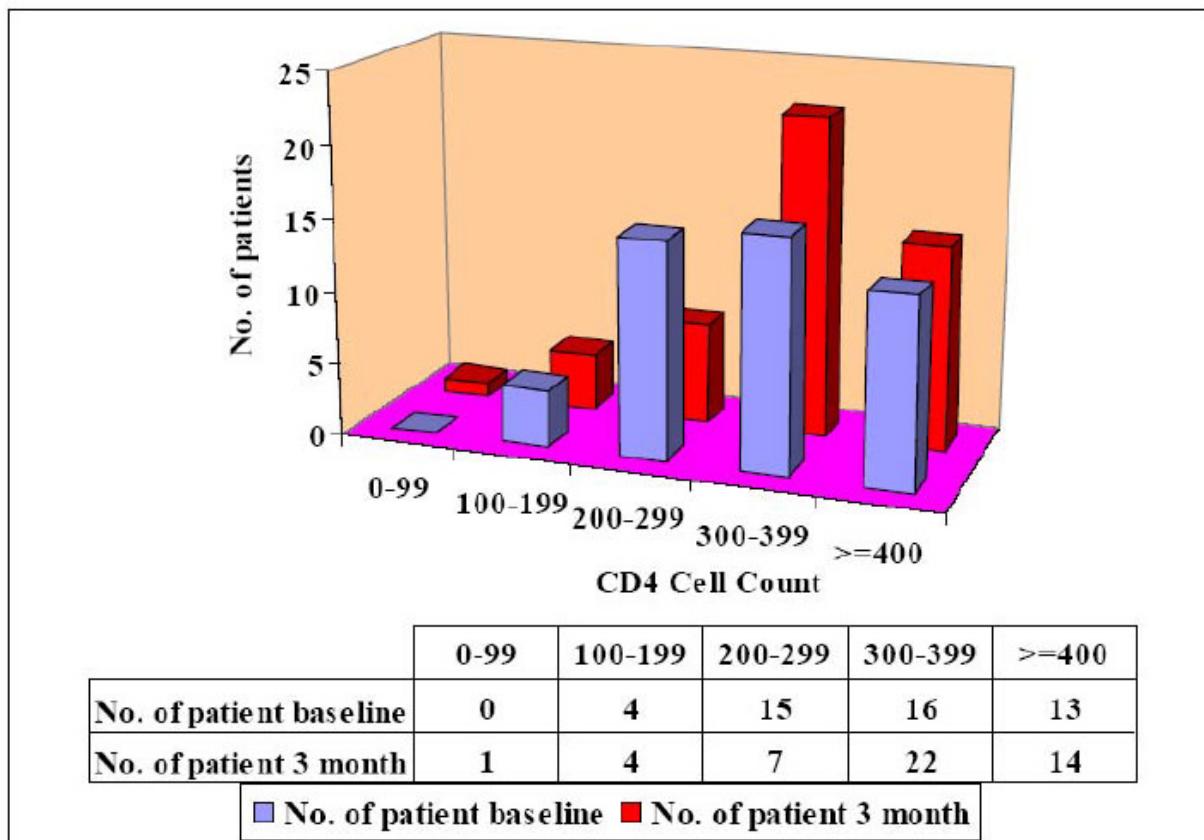


**Skin rash and Herpes zoster:** 14% and 12% of total study cases had skin rash and herpes zoster at basal respectively and from 4<sup>th</sup> week onwards not even a single patient had suffered from skin rash.

Similarly, nausea, vomiting and disturbed sleep were improved significantly during therapy with Radha108.

**Absolute CD4 cell count and HIV viral load:** CD4 cell count was available for 48 patients with pre and post treatment values. There was increase in CD4 count on the average by 51 (median CD4 cell counts from 312 to 363). This is of borderline statistical significance ( $p<0.06$ ). 30 out of 48 (62.5%) patients showed rise in CD4 count and 18 patients (37.5%) showed decrease in CD4 cell count. CD4 cell count range with number of patient at baseline and at the end of study is shown in Figure 4.

Figure 4 : No. of patients at baseline and at end of treatment with CD4 cell count range.



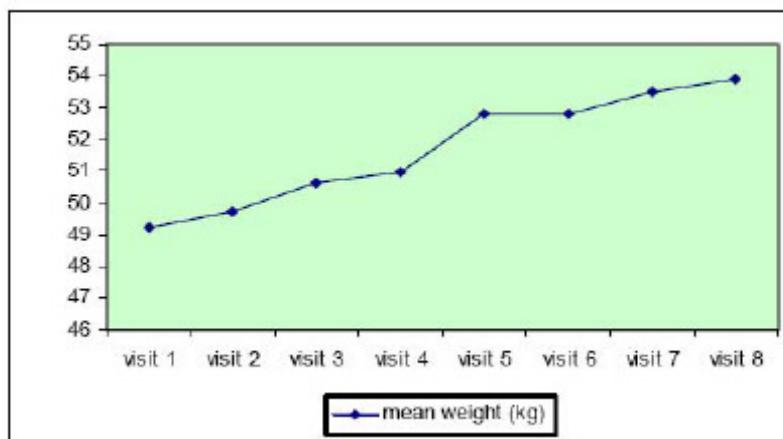
The mean HIV log viral load has statistically significantly dropped from 4.63 to 4.18 after 12 weeks of treatment ( $p=0.03$ ). Similar trend is also seen in the pre treatment and post treatment median viral loads (from 92458 vs. 25332,  $p<0.001$ ).

### **Result of phase III Radha108 trial – October 2006:**

Replica of positive result of Phase III Radha108 trial October 2005 was observed during Phase III Radha108 trial October 2006. During the study 51 patients were enrolled without any dropouts.

**Weight gain:** Weight monitored on every visit showed significant gain in all 51 HIV patients with mean weight gain of 4.68 # 1.9 kg after 12 weeks of Radha108 therapy ( $p<0.001$ ).(Figure 5).

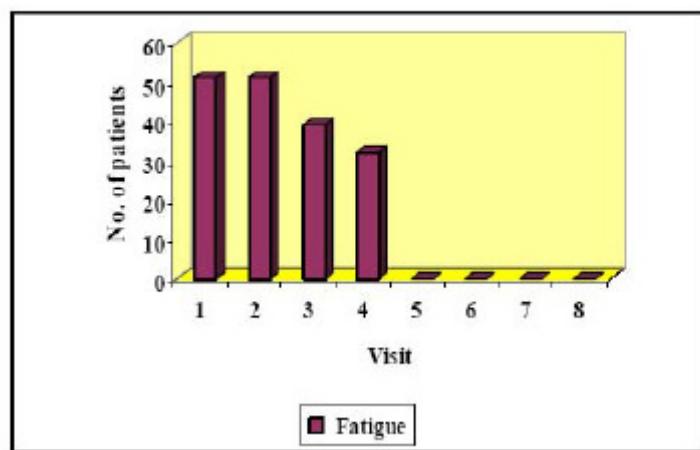
Figure 5 : mean weight in kg during receptol therapy



**Clinical Parameters:** All the clinical symptoms had disappeared during the 12 weeks of treatment. These reductions are statistically significant ( $p<0.005$ ).

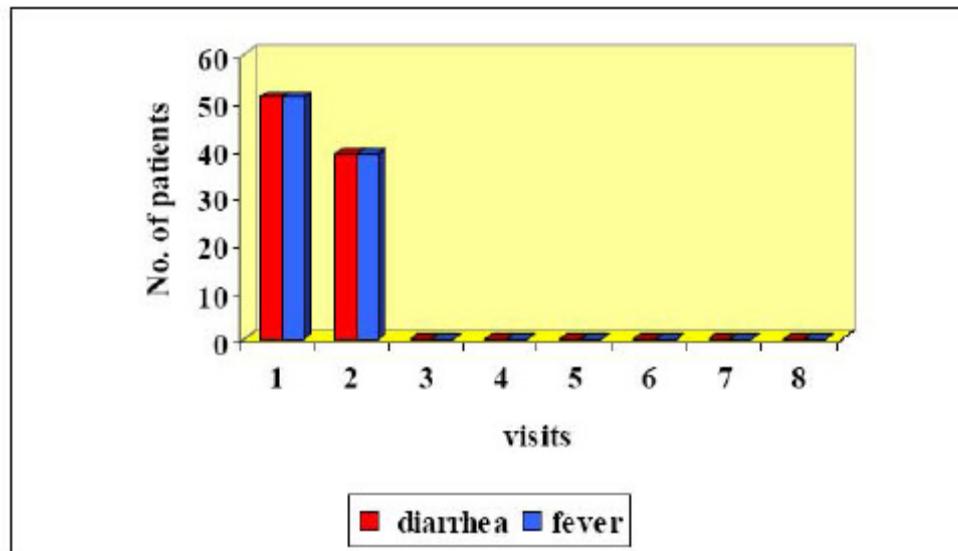
**Fatigue/Malaise:** 100% of the total study cases had symptoms of fatigue at basal. After treatment at the end of 2<sup>nd</sup> week proportion of symptoms of fatigue had a significant fall from basal. After 6<sup>th</sup> week onwards all patients had relief from fatigue (Figure 6).

Figure 6 : No. of patients with fatigue / malaise after therapy



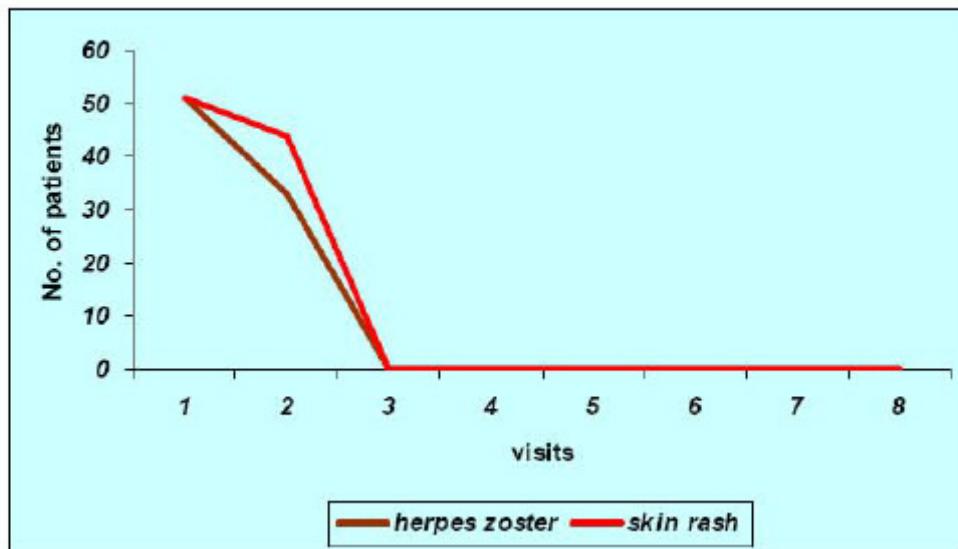
**Fever and diarrhea:** 100% of the total study cases had fever and diarrhea at basal and after treatment from 3<sup>rd</sup> week onwards all the patients had relief from diarrhea and fever (Figure 7).

Figure 7 : No. of patients with fever / diarrhea with receptol therapy.



**Skin rash and Herpes Zoster:** All the patients had HIV associated skin rash and herpes zoster at basal and became asymptomatic after 3<sup>rd</sup> week onwards with treatment (Figure 8).

Figure 8 : No. of patients having HIV associated skin rash and herpes zoster after therapy

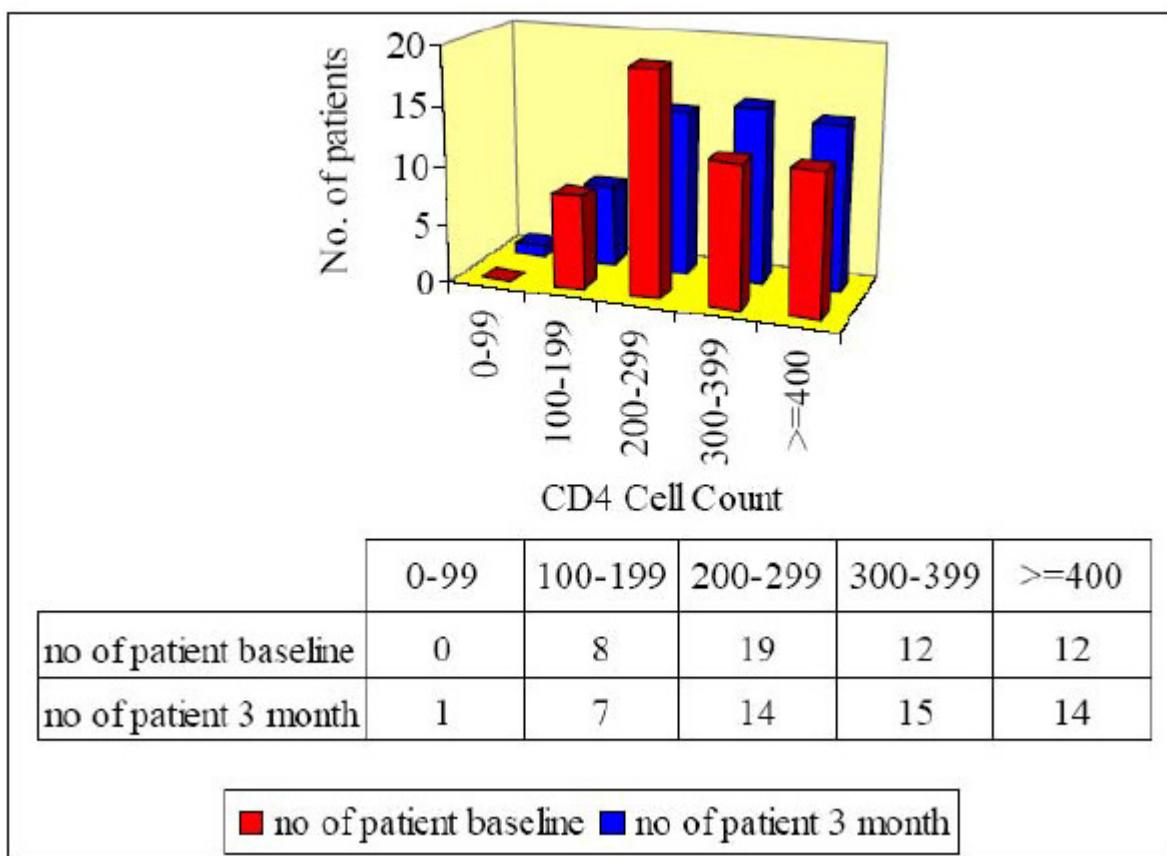


Similar improvements were seen in nausea, vomiting, cough and disturbed sleep and all patients became asymptomatic from 3<sup>rd</sup> week onwards.

### Absolute CD4 cell count:

CD4 cell count was available for all 51 patients with pre and post treatment values. There was increase in CD4 cell counts from 276 to 305). This value is statistically significant ( $p<0.05$ ). CD4 cell count range with number of patients at baseline and at the end of study is shown in Figure 9.

Figure 9 : No. of patients and CD4 cell count range



### HIV vital load:

After Radha108 12 weeks treatment at the end of study mean viral load showed a significant fall ( $p<0.01$ ) from the baseline conducted at Institute of Immuno Hematology (IIH) and ICMR Institute KEM Hospital Mumbai.

### Side Effects:

All patients tolerated Radha108 well with no side effects. Milk allergies are caused by the large milk proteins primarily casein and to a lesser extent the immunoglobulins. These proteins are completely removed from the Radha108. As Radha108 is a food substance derived from colostrum, it is entirely safe for human consumption.

## **Conclusion**

These results show that Radha108 is an effective and safe drug in increasing weight and general well being of patients, decrease in viral load and increase in CD4 cell count. Thus, Radha108, a pure natural drug having informational nanopeptides and PRPs as active pharmaceutical ingredient holds good promise for the treatment of HIV patient across all age groups.

“Radha108 preparations have been used to effectively treat a wide range of diseases this include bacterial, mycobacterial, fungal, parasitic, and viral and cancer.”

Radha108 is also useful as immunomodulator therapy in recurrent fungal, viral, bacterial infections, allergic and autoimmune disorders like asthma, SLE, rheumatoid arthritis etc, chronic fatigue syndrome, chronic dermatologic disorders, hepatocellular carcinoma, Alzheimer disease, Parkinson’s disease and other neurological disorder.

## References

1. Fierlbeck G et al [ Intralesional therapy of melanoma metastases with recombinant interferon-beta] Hautarzt. 43: 16-21 (1992);
2. Stuart-Harris RC et al The clinical application of the interferons: a review. NSW Therapeutic Assessment Group Med. Journal of Aust. 156: 869-72 (1992);
3. Gifford GE and Duckworth DH Introduction to TNF and related lymphokines. Biotherapy 3: 103-11 (1991);
4. Sato N et al Actions of TNF and IFN-gamma on angiogenesis in vitro. Journal of Investigative Dermatology 95: 85S-9S (1990);
- 5.. Granitov, VM et al...Usage of RECEPTOL in treatment of HIV – Infected patients. Russian Journal of HIV AIDS and Related Problems 2002, 1, 79-80
- 6.. World Health Organization. Progress on global access to HIV antiretroviral therapy: a report on "3 by 5" and beyond. 2006.
7. Brahmhatt H, Kigozi G, Wabwire-Mangen F, Serwadda D, Lutalo T, Nalugoda F, Sewankambo N, Kiduggavu M, Wawer M, Gray R. Mortality in HIV-infected and 11 uninfected children of HIV-infected and uninfected mothers in rural Uganda. J Acquire Immune Defic Snyder 2006;41(4):504-8.
8. Newell ML, Coovadia H, Cortina-Borja M, Rollins N, Gaillard P, Dabis F. Mortality of infected and uninfected infants born to HIV-infected mothers in Africa: a pooled analysis. Lancet 2004; 364(9441):1236-43.
9. Taha TE, Dallabetta GA, Canner JK, Chipangwi JD, Liomba G, Hoover DR, Miotti PG. The effect of human immunodeficiency virus infection on birth weight and infant and child mortality in urban Malawi. Int J Epidemiol 1995; 24(5):102
10. Barnes, D. Debate over potential AIDS drug. Science, 237(4811), pages 128-130, July 10, 1987.
11. Brenneman, D. and others. Neuronal cell killing by the envelope protein of HIV and its prevention by vocation intestinal peptide. Nature, 335(6191), pages 639-642, October 13, 1988.
12. Brenneman, D. and others. Peptide T prevents gp120 induced neuronal cell death in vitro: relevance to AIDS dementia. Drug Development Research, volume 15, pages 361-369, 1988.
13. Saharan P., Singh T., Safety and efficacy Clinical Trial of RECEPTOL\*: New Nanobiotechnology based immunomodulator in HIV therapy. Unpublished Patented data (2005-2006).

14. Janusz M., Staroscik K., Zimeckt M., Wieczorek Z., Lisowski J., Aproline-rich polypeptide (PRP) with immunoregulatory properties isolated from ovine colostrums. *Archivum immunologiac therapies experimentalis* (Warszawa) 34(4): 427-436 (1986).
- 15..Wieczorec Z., Zimecki M., Spiegel K., Lisowski J., Janusz M., Differentiation of T-Cells from immature precursors: identification of a target cell for a proline-rich polypeptide (PRP) *Archivum immunological therapies experimentalists* (Warszawa) 37(3-4):313-322(1989).
- 16..Kubis A., Marcinkowska E., Janusz M., Lisowski J. Studies on mechanism of action of a proline-rich polypeptide complex (PRP): Effect on stage of cell differentiation peptides 26(11) : 2188-2192 (2005).
17. Modulation of 4HNE-mediated signaling by proline-rich polypeptides from ovine colostrums. *Journal of Molecular Neuroscience* 20(2):125-134 (2003).
18. Immunology, Immunopathology and Immunity, Sell S, Appleton and Lange: Stamford CT 1996.
19. Bishop GA., Haxhinasto SA., Slunz LL., Hostager BS. Antigen specific B-lymphocyte activation. *Critical reviews in immunology* 23(3): 159-197 (2003).
20. Zibiocka A., Janusz M., Rybka K., Wirkus – Romanowska I. Kupryszeński G., Lisowski J. Cytoline inducing activity of a proline-rich polypeptide (PRP) from ovine colostrum and its active nanopeptide fragment analogs. *European Cytokine Network* 12(3) :462-467 (2001).
- 21.Fernandez- ortega C. Dubed M. Ruibal O. Vilarruba OL. Menendez de San Pedro JC. Navea L. Ojeda M. Arana MJ. Inhibition of in vitro HIV infection by dialyzable leucocyte extracts, *Biotherapy* 9(1-3)33-40 (1996).
22. Claes – Henrik Floren S., Chinene L., Elfstrand C., Hagman L. Thse. Coloplus a new product based on bovine colostrum alleviates HIV- associated diarrhoea. *Scandinavian journal of Gastroenterology* 2006: 41-68 2-686.
23. Saharan P. Mammalian Colostrum derived nanopeptides for broad spectrum viral and recurrent infection with a method of isolation thereof. *Patent submitted to Indian Patent office*, 2007.
24. Zimecki M,Staroscik K, Janusz M, Lisowski J, Wieczorek Z. The inhibitory activity of proline-rich polypeptide on the immune response to polyvinyl pyrrolidone (PVP). *Arch Immunol Ther Exp (Warsz)*1983;31(6):895-903.
25. Julius MH, Janusz M, Lisowski J. A colostral protein that induces the growth and differentiation of resting B lymphocytes. *J Immunol*, 1988; 140(5):1366-371.

26. Effects of oral dietary supplementation with Ai/E10® upon Natural Killer (NK) cell activity in a healthy human population. *Quantum Research, Inc.*, Scottsdale, Arizona, 2001.
27. An examination of Immune Response Modulation in Humans by Ai/E10® utilizing a double blind study. *Immune Consultants, Inc.*, Tucson, Arizona, 2001.
28. Boldogh I, Liebenthal D, Hughes TK, Juelich TL, Georgiades JA, Kruzel ML, Stanton GJ. Modulation of 4HNE-mediated signaling by a proline-rich polypeptides from ovine colostrum. *J Mol Neurosci*. 2003;20(2):125-134.
29. Pizza G, Chiodo F, Colangeli V, Gritti F, Rais E, Fudenberg HH. Preliminary observations using HIV-specific transfer factor in AIDS. *Biotherapy*. 1996;9(1-3):4-47.
30. Eggena MP, Barugahare B, Jones N, Okello M, Mutalya S, Kityo C, Mugyenyi P, Cao H. Depletion of regulatory T cells in HIV infection is associated with immune activation. *J Immunol*. 2005; 174(7):4407-4414.
31. Macroft A, Youle M, Moore A, et al. Reasons for modification and discontinuation of antiretrovirals : results from a single center. *AIDS*, 2001; 15:185-94.
32. Fatenkeheur G, Romer K, Cramer P, et al. High rates of changes of first antiretroviral combination regimen in an unselected cohort of HIV 1 infected patients. *8th ECCAT Greece*, 2001; abstract no. 50.
33. Rais E, Guerra L, Viza D, Pizza G, De Vinci C, Schiattone ML, Rocaccio L, Cicognani M, Gritti F. Preliminary results in HIV I infected patients with transfer factor (TF) and zidovudine (ZDV). *Biotherapy*. 1996;9(1-3):49-54.
34. Shi M, Hao S, Chan T, Xiang J. CD4+ T cells stimulate memory CD8+ T cell expansion via acquired pMHC I complexes and costimulatory molecules, and IL-2 secretion. *J Leuco Biol*, 2006; 80(6):1354-1363.
35. Khan A. Non-Specificity of transfer factor. *Annals of Allergy*, 1977. 38(5): 320-322.
72. Razonable, Raymund, R., et.al. Division of Infectious Diseases and Internal Medicine, Mayo Clinic and Foundation, Selective Reactivation of Human Herpesvirus 6 variant A Occurs in Critically Ill Immunocompetent Hosts, *The Journal of Infectious Diseases*, January , 2002.
73. Dwyer JM. The use of antigen-specific RECEPTOL in the management of infections with herpes viruses. In: Kirkpatrick CH, Burger DR and Lawrence HS eds. *Immunobiology of RECEPTOL*. New York Academic Press 1983:233-243.

- 74.Viza D, Vich JM, Phillips J et al. Orally administered specific RECEPTOL for the treatment of herpesvirus infections. *Lymphok Res* 1985;4:27-30.
- 75.Jones JF, Jeter WS, Fulginiti VA et al. Treatment of childhood combined Epstein-Barr virus/cytomegalovirus infection with oral bovine RECEPTOL. *Lancet* 1981;2:122-124.
- 76.Ablashi DV, Levine PH, DeVinci C et al. Use of anti HHV-6 RECEPTOL for the treatment of two patients with chronic fatigue syndrome (CFS). Two case reports. *Biotherapy* 1996;9:81-86.
- 77.Steele RW, Myers MG and Monroe VM. RECEPTOL for the prevention of varicella-zoster infection in childhood. *N Engl J Med* 1980;303:355-359.
- 78.Lang I, Nekam H, Gergely P et al. Effect of in vivo and in vitro treatment with dialyzable leukocyte extracts on human natural killer cell activity. *Clin Immunol and Immunopathol* 1982;25:139-144.
- 79.Boucheix C, Phillips J, Pizza G et al. Activity of animal RECEPTOL in man. *Lancet* 1977;1:198-199.
- 80.Fudenberg H and Pizza G. RECEPTOL 1993: New frontiers. *Progress in Drug Res* 1994;42:309-400.
- 81.Aralla-Chaves M, Ramos MTF and Rosado RMF. Evidence for prompt and intense constitution of cell-mediated immunity by means of RECEPTOL in a case of complex immune deficiency. *Cell. Immunol.* 1974;12:160.
- 82.Ballow M and Good RA. Report of a patient with T-cell deficiency and normal B-cell function: a new immunodeficiency disease with response to RECEPTOL. *Cell. Immunol.* 1975;19:219.
- 83.Jones JF, Pizza G, DeVinci C. Infectious mononucleosis: immunotherapy with EBV-specific RECEPTOL. *J Exp Pathol* 1987;3:399-406.
- 84.Khan A, Hansen B, Hill NO et al. RECEPTOL in the treatment of herpes simplex types 1 and 2. *Dermatologica* 1981;163:177-185.
- 85.Winkelmann RK, DeRemee RA, Ritts RE Jr. Treatment of varicella-zoster pneumonia with RECEPTOL. *Cutis* 1984;34:278-281.
- 86.Rozzo SJ and Kirkpatrick CH. Purification of RECEPTOLs. *Mol Immunol* 1992;29:167-182.
- 87.Pizza G, Viza D, Roda A et al. RECEPTOL for the treatment of chronic active hepatitis. *N Engl J Med* 1979;300:1332.

- 88.Nkrumah F, Pizza G, Viza D et al. Regression of progressive lymphadenopathy in a young child with acute cytomegalovirus (CMV) infection following the administration with specific anti-CMV activity. *Lymphok Res* 1985;4:237-241.
- 89.Neequaye J, Viza D, Levine PH et al. Specific RECEPTOL with activity against Epstein-Barr virus reduces late relapse in endemic Burkittâs lymphoma. *Anticancer Res* 1990;10:1183-1187.
- 90.Viza D, Vich JM, Phillips J et al. Specific RECEPTOL protects mice against lethal challenge with herpes simplex virus. *Cell Immun* 1986;100:555-562.
- 91.Wilson GB, Poindexter C, Fort JD et al. De novo initiation of specific cell-mediated immune responsiveness in chickens by RECEPTOL (specific immunity inducer) obtained from bovine colostrum and milk. *ACTA Virol* 1988;32:6-18.
- 92.Kirkpatrick CH, Hamad AR, and Morton LC. Murine RECEPTOLs: dose-response relationships and routes of administration. *Cell Immunol* 1995;164:203-206.
- 93.Viza D, Lefesvre A, Patrasco M et al. A preliminary report on three AIDS patients treated with anti-HIV specific RECEPTOL. *J Exp Path* 1987;3:653-659.
- 94.Pizza G, DeVinci C, Palareti A et al. 25 years of clinical experience with RECEPTOLs. XI International Symposium on RECEPTOL. March 1-4, 1999. Monterey, Mexico.
- 95.Barnes, D. Debate over potential AIDS drug. *Science*, 237(4811), pages 128-130,July10,1987.
- 96.Brenneman, D. and others. Neuronal cell killing by the envelope protein of HIV and its prevention by vasoactive intestinal peptide. *Nature*, 335(6191), pages 639-642,October13,1988.
- 97.Brenneman, D. and others. Peptide T prevents gp120 induced neuronal cell death in vitro: relevance to AIDS dementia. *Drug Development Research*, volume 15,pages361-369,1988.
- 98.Bridge, P. and others. Peptide T: Improvements in phase I trial of AIDS patients.Draft of letter submitted to Lancet, July 1989.
- 99.Kowalski, M. and others. Functional regions of the envelope glycoprotein of human immunodeficiency virus type 1. *Science*, 237 (4820), pages 1351-1355, 1987.
- 100.Lasky and others. Delineation of a region of the human immunodeficiency virus type 1 gp120 glycoprotein critical for interaction with the CD4 receptor. *Cell*, volume 50 number 6, pages 975-985, 1987.

101.Nygren and others. 95- and 25-kDa fragments of the human immunodeficiency virus envelope glycoprotein gp120 bind to the CD4 receptor. Proceedings of the National Academy of Sciences U. S. A., volume 85 number 17, pages6543-6546,1988.

102.Pert, C., and others. Octapeptides deduced from the neuropeptide receptor-like pattern of antigen T4 in brain potently inhibit human immunodeficiency virus receptor binding and T-cell infectivity. Proceedings of the National Academy of Sciences U. S. A., volume 83, pages 9254-9258, December 1986.

103.Pert interview, Science Impact, pp. 6-7, June 1987.  
Ruff, M., and others. Peptide T[4-8] is core HIV envelope sequence required for CD4 receptor attachment.Lancet, 2(8561), page 751, Sept. 26, 1987.

104.Sodroski, J., and others. HIV envelope-CD4 interaction not inhibited by synthetic octapeptides. Lancet, 1(8547), pages 1428- 1429, June 20, 1987.

105.Wetterberg, L., and others. Treatment with peptide T in seven immunodepressed HIV infected patients. Draft of paper submitted to AIDS, Gower Academy Journal, London, June, 1988.

106.Wetterberg, L., and others. Peptide T in treatment of AIDS. The Lancet, 1(8525), page 159, Jan. 17, 1987.