

Increased Prevalence of Neuroleptic Malignant Syndrome, Dementia and Fragmented Germ cells with Reduced Endogenous Estrogen

Proof of Concept Study-Retrospective Analysis

Dr. Elizabeth JeyaVardhini Samuel¹, Dr. Sanjoy Joseph George²

¹M.B.B.S., M.D. (Internal Medicine) Assistant Professor,

²M.B.B.S., Post Graduate Resident in Medicine, Department of Medicine, Pondicherry Institute of Medical Sciences, No.20, KanagaChettikulam, Kalapet, Pondicherry. South India. 605014

Abstract: *Background:* Era of contraception, abortions, [20th, 21st centuries] implemented as family welfare schemes witnessed, increase in prevalence of dementia, psychosis, stroke, Parkinson's disease, cerebral tumors, myoclonus, seizures and Neuroleptic malignant syndrome. *Objectives:* altruistic association of contraception [if any], with increasing dementia, stroke, Neuroleptic malignant syndrome, psychosis, cerebral tumors, Parkinson's disease in younger age was sought after. *Methods:* In 2012, retrospective analysis of, prevalence of stroke, in 350 patients of 20-35 years, 35-50 years, >50 years age groups, from data collected by convenient, stratified random sampling, from different geographical locations, between 2003-2012 and its association with presence, absence of contraception, abortion was undertaken; simultaneously, serum estrogen levels obtained from 105 patients, was also analyzed. Retrospective analysis of data of 279 patients who had presented with stroke, cerebral tumors, tuberculous meningitis, Parkinson's disease, myoclonus, seizures over 1989-2012 was undertaken. Data of another group of 318 patients presenting with psychosis including suicides, dementia, behavioral disorders spread over 1989 to 2012, of whom ~100 were college students with behavioral disorders, was also analyzed retrospectively for significant association of contraception status, cholesterol deprived diet and prevalence of neuropsychiatric disabilities. Steep rise in incidence of Neuroleptic malignant syndrome e.g. 15 patients within 6 months in 2007 was identified in young women after tubectomy [$p < 0.0005$]. *Results:* With no difference attributable to geographical location the results revealed. [a] Cholesterol deprived diet was associated with behavior disorders, including suicides as >100 fold increase in <20 years college students [$p < 0.0005$]; chronic cholesterol deprived diet without contraception was associated with 10 fold increase in psychosis, dementia among 35->50 years with a p value of <0.0005[b]Contraception with cholesterol deprived diet showed 30- 40 fold increase in psychosis starting from younger age itself 20->50years with a p value of <0.0005 and 30 fold increase in dementia in >50 years with a p value of <0.0005. [c]Neuroleptic malignant syndrome was seen with absolute increase in patients with tubectomy [$p < 0.0005$]. [d]stroke, tumors, encephalitis, Parkinson's disease, myoclonus, were seen with ~15 fold increase in contraceptive users with a p value of <0.0005. [f] Contraception reversal with essential cholesterol, fatty acids rich diet cured demyelination, behavior disturbances. *Conclusion:* The concept is Contraception, abortion preventing normal path traversed by germ cells, with smashed fragmentation of germ cells, consequent reduced endogenous estrogen: androgen, resultant defaulted genomic repertoire, deranged cell cycle, cell metabolism, degeneration of neuronal cells, neurotransmitters and cholesterol deprived diet with reduced myelin synthesis, hormone synthesis, neurotransmitter synthesis leading to ~40 fold increase of neuronal diseases with a p value of <0.0005 in young age itself. Since dopamine is inherently reduced by deranged cell metabolism after contraception, Neuroleptic: anti psychiatric drugs by further dopaminergic blockade precipitate the fatal Neuroleptic malignant syndrome; contraception reversal, with cholesterol rich diet enables therapies effective and prevents further deterioration of neurological pathologies, can prevent increasing prevalence of Neuroleptic malignant syndrome.

Keywords: Autologous germ cell replants; contraception reversal; defaulted genomic repertoire, deranged cell metabolism

1. Introduction

Era of contraception [20th, 21st centuries as family welfare schemes were progressively, stealthily implemented,] witnessed increased incidence of Neuroleptic malignant syndrome, increased prevalence of dementia, psychosis, behavior disorders including suicides, Parkinson's disease and other degenerative neurological disorders occurring in young age, cerebral tumors, and tuberculous meningitis in spite of advancing research and technologies. Cholesterol deprived diet was recommended along with contraception. Retrospective analysis was planned to detect [If any] altruistic association existed between contraception, abortion cholesterol deprived diet and increasing prevalence of neuro psychiatric disabilities.

2. Methods

Minimum of 30 samples was planned for, in each of 3 age groups, namely 20-35 years, 36-50 years, >50years; though people from the community are visiting the hospital, analysis of hospital patients alone can create a bias, hence data from the community, hospital, health screening camps, of different geographical locations were included; data collection was solely done by the corresponding author; data from each person included prevalent diseases, status of contraception, hysterectomy, type of oil ingested, life style, level of nutrition, presence of anemia; the data was tabulated as prevalent diseases, matched against the variables in each age group; retrospective bioinformatics analysis was done, by plotting histograms for the 3 age groups and cumulative graphs for

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each disease in 2012; an example of tabulation of the data is provided in the supplementary file.

In 2003 house to house survey in the community, spread over 3 weeks, to collect data of prevalent diseases of 100 people; the people who were present during the survey were included at random, by convenient sampling into the 3 age groups namely 20-35 years, 35-50 years, >50 years, to include a minimum of 30 people in each age group; serum estrogen estimation was done for 12 people as per their request; the reduced estrogen levels [5-8pg] found in young contraceptive users, was the eye opener, leading to further data analysis.

In 2004 data of 93 hospital patients was collected over a period of 6 months, including diseases prevalent, contraception status, life style, nutrition, type of oil ingested, level of hemoglobin and were assigned to the 3 age groups by stratified random sampling with a minimum of 30 patients in each age group; serum estrogen estimation was done for all 93 patients; the data was tabulated matching diseases against status of contraception and other variables; one patient was a foreign national.

In 2011, 96 people [43 couples] working in different states of our nation had attended a health screening camp conducted in the community, spread over 3 days and their data was analyzed after assigning into the 3 age groups at random, for association of diseases with status of contraception, hysterectomy and other variables; effect of contraception in both partners after contraception also could be analyzed; none had sedentary life style, low nutrition or anemia or had worn tight attires around the pelvis.

In 2012, data of 61 hospital patients including a foreign resident, from another geographical location, was collected over a span of 6 months, assigned to the 3 age groups at random and was pooled to the other data from 2003, 2004, 2011 and retrospective bio informatics analysis was undertaken for the 350 patients in 2012, by plotting histogram for the 3 age groups and cumulative graphs for each disease.

Every participant was informed about their data being included for study purpose and the concerned hospital authorities were also informed; an engineering college student of Karunya University did the bio informatics analysis as his project. Retrospective analysis of data of 279 patients who had presented with stroke, cerebral tumors, tuberculous meningitis, Parkinson's disease, myoclonus, seizures over 1989-2012 clinical practice of the corresponding author was undertaken; they were divided into the 3 age groups at random and the significance of association of contraception, abortion status with the prevalence of neurological illnesses was analyzed. Data of another group of 318 patients presenting with psychosis including suicides, dementia, behavioral disorders spread over 1989 to 2012, of whom ~100 were college students <20 years of age with behavioral disorders, was also analyzed retrospectively for significant association of contraception status, cholesterol deprived diet and prevalence of neuropsychiatric disabilities

3. Results

Irrespective of geographical location, results revealed the following -Medical patients on diet with essential fatty acids, without practicing contraception had not presented with psychosis, dementia and behavior disorders, so not seen on the figure-1. *Cholesterol deprived diet* was associated with >100 fold increase [figure-1] in behavior disorders including arrogance, terrorism, depression including suicides, reduced stress adaptability, addiction to drugs in <20 years age group, involving mainly unmarried college students, professionals, community with a p value of <0.0005, but psychosis was not seen. Longstanding cholesterol deprived diet consumption without contraception was associated with 10 fold increase in psychosis among 35- 50 years with a p value of <0.0005; 10 fold increase in dementia was seen among >50 years with a p value of <0.0005; 30 fold increase in behavior disorders, including suicides, depression were seen among >50 years with a p value of <0.0005.

Contraception with cholesterol deprived diet had 40 fold increase in psychosis, 40 fold increase in behavior disturbances in 20-35 years age group with a p value of <0.0005; 35-50 years had 30 fold increase in psychosis with a p value of <0.0005; >50 years had 30 fold increase in dementia, 40 fold increase in psychosis with a p value of <0.0005; *Contraception users on cholesterol deprived diet had 3-4 times higher risk of psychosis than patients on cholesterol deprived diet without contraception from 20years to >50 years of age group; dementia seen in >50 years age group occurs 3-4 times more in contraceptive users with cholesterol deprived diet than the patients without contraception. 20-35 years age group on cholesterol deprived diet had high prevalence of behavior disturbances only but no psychosis but as age advances the psychosis, dementia also manifest explaining worsening neural functions in relation to the prolonged deprivation of essential fatty acids-which constitute the sphingo lipids, phospho lipids- basic moiety of neuronal system.*

Number of patients without contraception also decrease with age, majority switching to contraception by 25 years (after marriage, family planning, small family norms, one child policy) leading to higher incidence of severe neuropsychiatric disorders.

Prevalence of Neuroleptic malignant syndrome requiring intensive care was increasing for e.g. 3 patients within 7 days in 20-35 years age group receiving anti psychotics; all had undergone tubectomy, of them one had associated systemic lupus erythematosus also.

Stroke was 4-8 fold increased among contraception users aged 20->50 years [p<0.0005] figure2. Contraception status was associated with 10 fold increase in neural disorders like glioblastoma multiforme, with a p value of <0.0005; cerebral infarcts, cerebral hemorrhages, Parkinson's disease[figure3]- was seen with 40-60 fold increase with a p value of <0.0005; myoclonus, postural tremors, seizures were seen with 20 fold

increase in contraception users with a p value of <0.0005. Schwannoma was the tumor (figure-3) seen in non contraception user of 20 years, but glioblastoma was seen in two patients above 35-50 years without contraception also; prevalence of glioblastoma multiforme [biopsy proven] was 10 fold higher among contraception users with a p value of <0.0005.

>70 years patients were non contraception users because contraception users (in our country its permanent sterilization mostly with constant fragmentation of germ cells) had embraced early demise before that age.

Seizures, myoclonus in non contraception users was due to neuro cysticercosis, herpes encephalitic sequelae, consuming fish without scales and gills (prawns) post trauma; improved, cured, controlled with treating their cause; whereas the 20 fold increase in seizures seen after contraception was with normal computerized axial tomography scan (probably degenerative etiology), rheumatic chorea (altered immunity) all stemming from smashed destruction of germ cells in contraception. Demyelination, degenerative diseases had 30-50 fold increased among 35-50 years, 50-70 years of age in contraception users including neuromyelitis optica with a p value of <0.0005; Parkinson's disease is seen in >70 years in non contraception users. (when physiological decrease of endogenous estrogen, androgen occurs to 5pg results in degenerative diseases) *whereas Parkinson's disease had occurred with 30-50 fold increase among 30-50 years among contraception users because endogenous estrogen had decreased to 5-8pg-[figure 4] after contraception in 61-75% of people [p<0.0005] in 20-50years itself leading to degenerative diseases, metabolic syndrome.*

Two subsets of population one with, the other without contraception affecting both partners is reflected in pattern of diseases` incidence, severity and prognosis; similar results can be reproduced in any part of the globe if contraception etiology [stealthily implemented, with presumption of no side effect], cholesterol deprived diet are considered.

Contraception reversal-[figure-supplementary file]-Cause and effect phenomena could be appreciated only on one patient on contraceptive pills presenting with neuromyelitis optica, dystonia, myoclonic epilepsy, urosepsis, quadriparesis recovered completely on discontinuing oral estradiol-contraceptive and other treatment including intravenous immunoglobulin, cholesterol containing diet; *other patients` tubal recanalisation could not be implemented [nonexistent protocols; unwilling patients, lack of general awareness of effective therapy with contraception reversal] though recommended; cholesterol rich diet, supplementation of essential fatty acids [consumption of oil with cholesterol, non refined oil, virgin olive oil, gingili oil, coconut oil, palmolein oil, and avoiding sunflower oil with 6 times less vitamin E] reverted behavior disturbances in youth remarkably.*

Contraception is associated with 275% increase in diseases including neuro psychiatric illnesses in addition to cholesterol

deprivation-[figure-supplementary file]. *Consumption of Fish without scales and gills consumption also precipitates myoclonus, transverse myelopathies, radiculopathies in both contraceptive and non contraceptive users but higher proportion is seen in association with contraception.*

4. Discussion

Psychosis refers to disordered functioning in the areas of emotion, perception, thinking, memory; in general population 1-2% has psychosis; 10% of patients attending psychiatrists out patients` clinics have psychosis; genetic contribution is polygenic; brain structure is generally normal but function of brain is altered due to imbalance in neurotransmitters like dopamine, nor adrenaline, serotonin, acetylcholine;¹ Dementia affects 5% in >65years, 20% in >85 years; global impairment of cognitive function, memory is most affected; language disability, defect in concentration, attention gradually become apparent associated with atrophy of brain tissue.

Alzheimer`s dementia² shows diffuse vascular disease with loss of previously acquired intellectual function when arousal is not impaired; 15% is familial, early onset is autosomal dominant, late onset is polygenic; mutations in several genes cause the disease; cerebral cortex is atrophic with histo pathological examination characterized by senile plaques, neurofibrillary tangles in cortex with significant quantity of amyloid in the plaques; stain for ubiquitin protein noted positive i.e. damaged proteins; neurotransmitter abnormalities, impaired cholinergic transmission, so donepezil helps with familiar environment.

Homovanillic acid-major catabolite of dopamine, 5 hydroxyl indole acetic acid –major catabolite of serotonin³ are normally present in cerebrospinal fluid and are altered in patients with idiopathic, drug induced parkinsonism.

First, second generation of antipsychotics act by blocking D2 dopamine receptors⁴ in the brain; extra pyramidal effects of dopamine blockade are akathisia, Parkinson`s disease, dystonia, tar dive dyskinesia; cholinergic blockade are the dry mouth, blurred vision, urinary retention, impotence.

Neuroleptic malignant syndrome is a rare manifestation secondary to idiosyncratic reaction to antipsychotics, presenting with fever, rigidity, tremor, confusion, autonomic instability; associated with 20% mortality untreated; with intensive care, mortality is 5%; antipsychotics have to be discontinued with hydration, bromocriptine, dantrolene sodium, temperature reduction, monitoring.

Central nervous system has 100 billion neurons, 10-50 times this number of glial cells, all in constant metabolism; transport speed 400mm/d, ~as that of electricity; made up of sphingolipids, phospholipids,⁵ lipid complex myelin is produced by Schwann cells wrapping its membrane 100 times around axon; oligodendrocytes send off multiple processes to form myelin on neighboring axons

Concept is contraception (without therapeutic indication as against the principles of practice of medicine, without evidence for need, safety,) results in smashed destruction of germ cells⁶ i.e. a centric fragments, chromatid breaks, ring chromosomes, resultant reduced endogenous estrogen-figure-4, androgen mediated by endorphins to hypothalamic pituitary axis leads to degenerative pathologies of every cell, with its mediators, transmitters, mutations including neurons, because the cell differentiation, controlled multiplication⁷ metabolism is under this endogenous estrogen androgen surveillance; hence contraception is associated with 40 fold increase in psychosis, dementia in 35->50 years; ~15 fold increase in tumors, 40-60 fold increase in parkinsonism, demyelination, degenerative pathologies, infarcts secondary to increased thrombogenicity associated with low estrogen.

Contraception has decreased, degenerated levels of dopamine, acetyl choline, other neuro transmitters leading to increase in psychosis; antipsychotics acting by dopaminergic blockade, precipitate Neuroleptic malignant syndrome with absolute increase 100% after permanent methods of sterilization i.e. tubectomy [that's why its clearly seen in India, than can be seen in intermittent temporary contraceptive measures abroad] pro dopamine drugs e.g. bromocriptine can help reduce the incidence; for people on contraception pro dopamine antipsychotics may help than anti dopaminergic, contraception reversal, prohibition of abortions will revert, prevent this dramatic neuroleptic malignant syndrome associated with significant mortality.

Cholesterol of oil, derived from seeds harboring seedlings with phyto estrogen, vitamin E, lipoproteins is highly essential for neuronal metabolism, neuronal synthesis⁸ and hormonal synthesis; hence cholesterol deprivation [consumption of sunflower oil (-it has 6 times less vitamin E than ground nut seed), refined oil, bleached oil] relating to its duration manifests with 100 fold behavior disorders in 20-35 years, 10 fold increase in psychosis, dementia in 35 years to >50 years. Consumption of fish without scales, gills (prawns, crabs) are associated with myoclonus, demyelination secondary to probably toxin mediated altered immunity so need to be avoided. Contraception reversal, cholesterol rich diet, decreases diseases as a cause and effect phenomenon.

5. Conclusion

Contraception implemented, permitted by Life Sciences as guillotine protocol (without therapeutic indication, without evidence for need or safety as against principles of practice of Medicine) with its unaware, overlooked smashed destruction of Germ Cells, resultant reduced endogenous estrogen, androgen, resultant increased degeneration of every cell, transmitters, cytokines, autoimmune destruction, demyelination, neoplastic transformation, increased thrombogenicity has led to 40 fold increase in psychosis, dementia, 40-60 fold increase in cerebral infarcts, parkinsonism, demyelination, 10 fold increase in malignant neoplasm, 30 fold increase in myoclonus, seizures with a p value of <0.0005.

Essential fatty acids, cholesterol deprived diet due to decreased neuronal, myelin elemental synthesis since the basic fatty acids for nerve synthesis are deprived, results in 10 fold increase in psychosis, dementia in 35-50 years, 50-70 years; whereas in 20-35 years age group 100 fold increase in behavior disorders, correlating with the longevity of cholesterol deprivation more severe degradative impairment in functional, structural pathologies.

Neuroleptic Malignant syndrome though rare is increasingly seen with absolute significance in people after tubectomy, on antipsychotics; clinical subset of population-contraception users have impaired production, function of neurotransmitters including dopamine; hence Neuroleptic with dopaminergic blockade produce Neuroleptic malignant syndrome in contraceptive users with high mortality.

Prodopamine antipsychotics may reduce Neuroleptic malignant syndrome in contraception users; halt, reversal of contraception by life sciences will definitely reduce these disorders as in the era before contraception making 21st centuries advanced therapies very effective.

Consumption of fish without scales, gills [prawns, crabs] also precipitate myoclonus, transverse myelopathies, radiculopathies, demyelination, angioneurotic edema, pancreatic cancers, diverticulitis, urticaria, nephritis, gastroenteritis probably due to toxin, hap ten mediated autoimmunity.

As a cause and effect phenomena, *contraception reversal – autologous germ cell replant effect*, with *cholesterol rich diet restores endogenous hormones*, cures demyelination, dystonia, behavior disorders without relapse. Similar results can be obtained in any part of the globe, since basic cell function is uniform irrespective of geographical variation, if only contraception, cholesterol deprivation is included in the analysis towards elucidation of etiology; biochemical attempts to elucidate the toxin in fish without scales, gills which induces autoimmunity, demyelination can be considered.

6. Key Points

- Neuroleptic malignant syndrome occurs with absolute increase after tubectomy
- Contraception users have 40 fold increases in dementia, psychosis, cerebral infarcts, Parkinsonism, demyelination; 10 fold increases in [glioblastoma] malignant neoplasm; 30 fold increases in myoclonus, seizures with a p value of <0.0005.
- Cholesterol deprived diet produces 100 fold increase in behavior disorders in 20-35 years age group, 10 fold increases in psychosis, dementia in 35->50 years with a p value of <0.0005.
- Consumption of fish (water animals) without scales, gills are associated with myoclonus, demyelination, transverse myelopathies, and radiculopathies.

- Autologous Germ cell replant effect- contraception reversal with essential cholesterol rich diet cures demyelination, degenerative diseases like Parkinson’s syndrome, behavior disorders.

7. Conflicts of Interests

None Declared

8. Acknowledgements

Mr. Jobin John - an Engineering College student of Karunya University planned the bioinformatics figures to enable retrospective analysis. His present contact address: Jobin John, B.Tech (Bioinformatics), ZIFO Technologies Pvt. Ltd, No. 21A, Anna Salai, Little Mount, Saidapet, Chennai, India

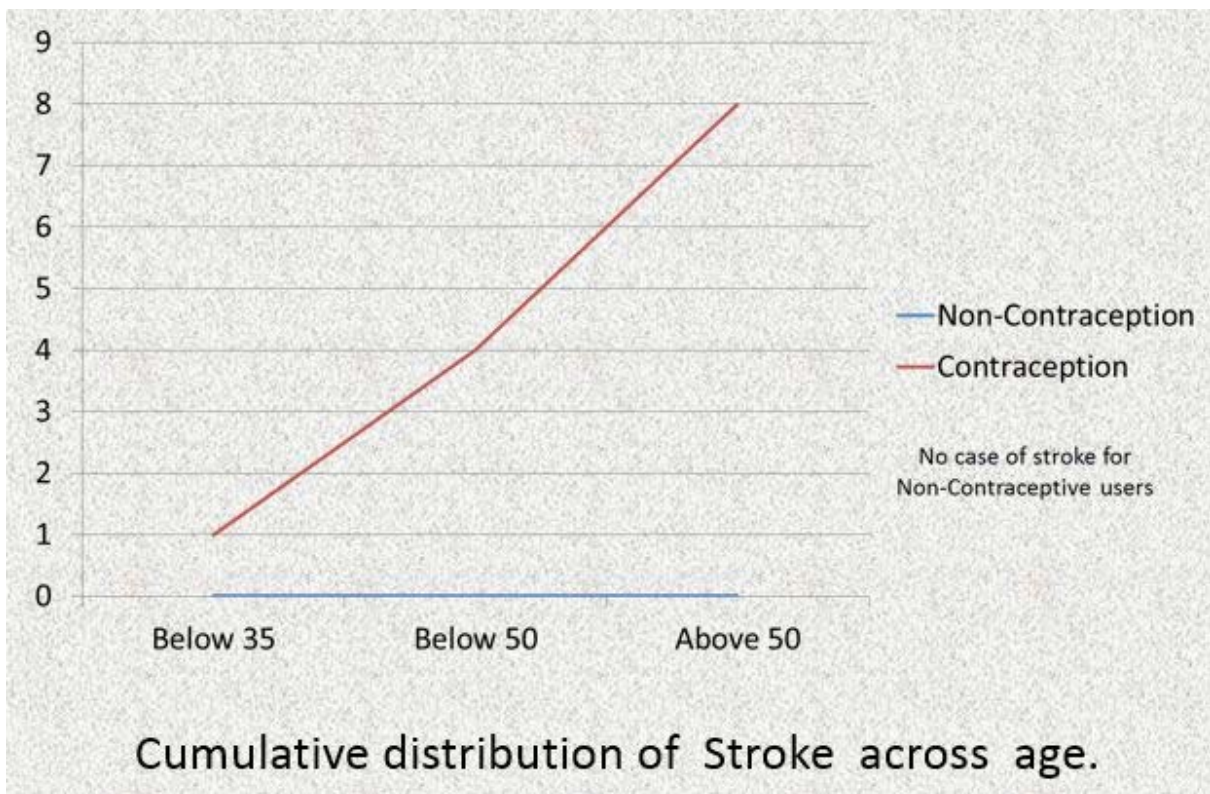
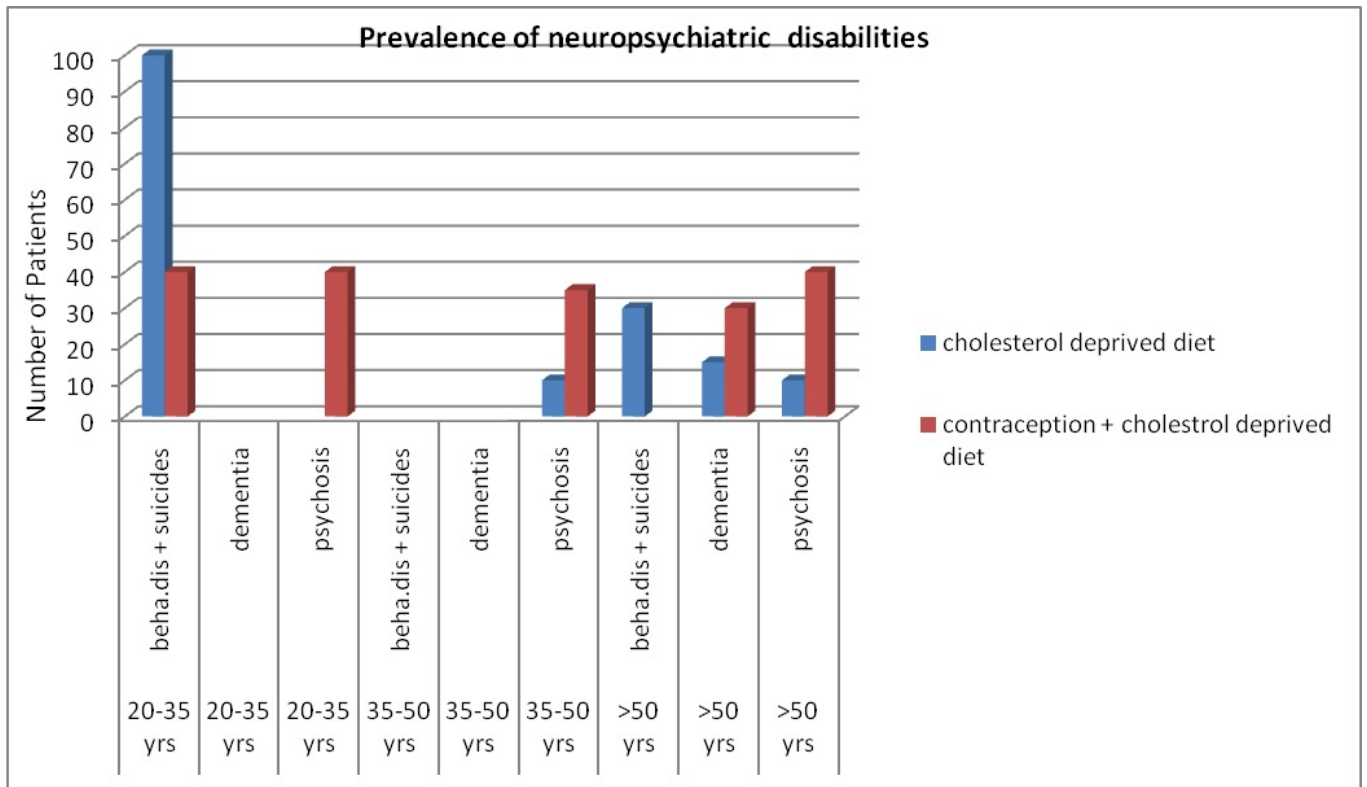
References

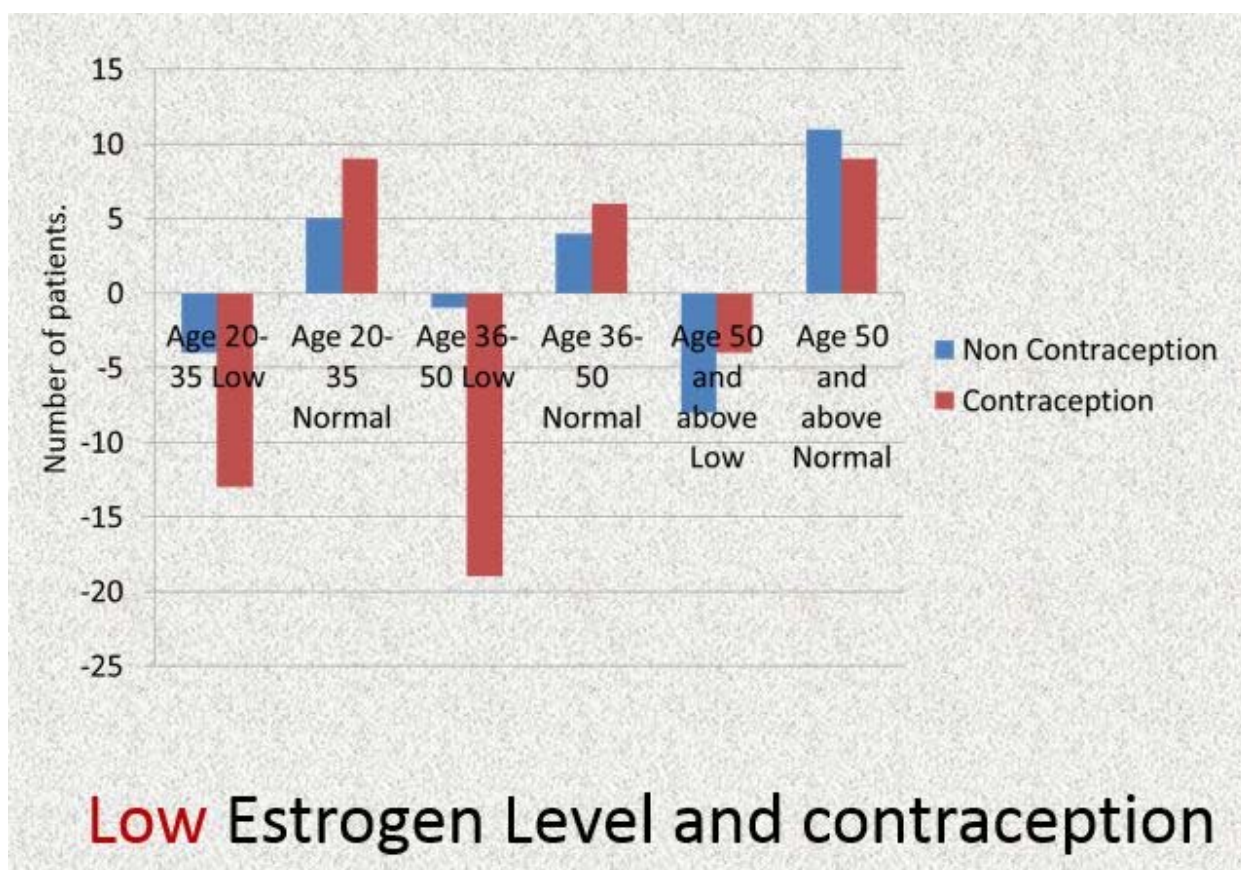
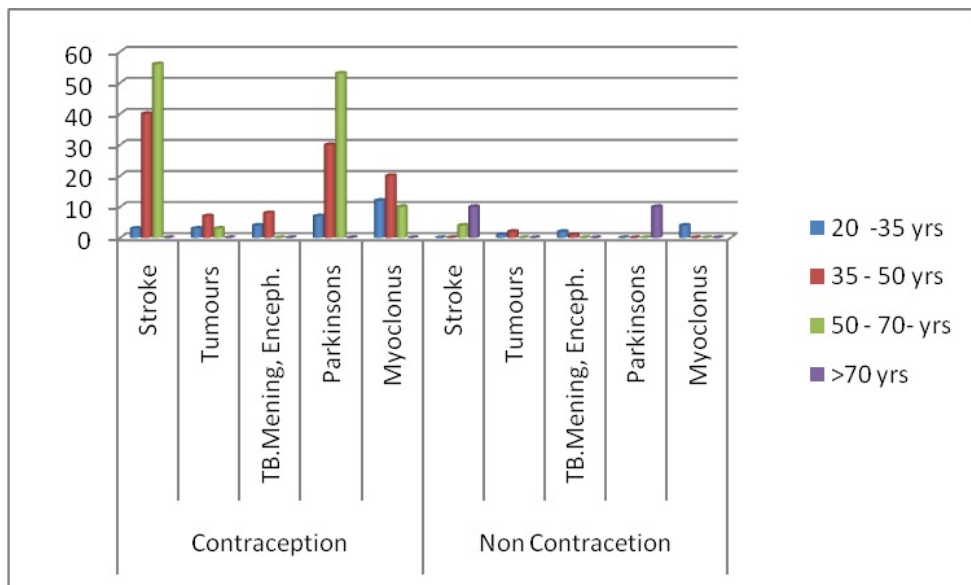
[1] M.C.Sharpe, S.M.Lawrie: Epidemiology of Psychiatric Disorders, Davidson’s Principles and Practice of Medicine 21st edition,2010 pages-230-231;248-249

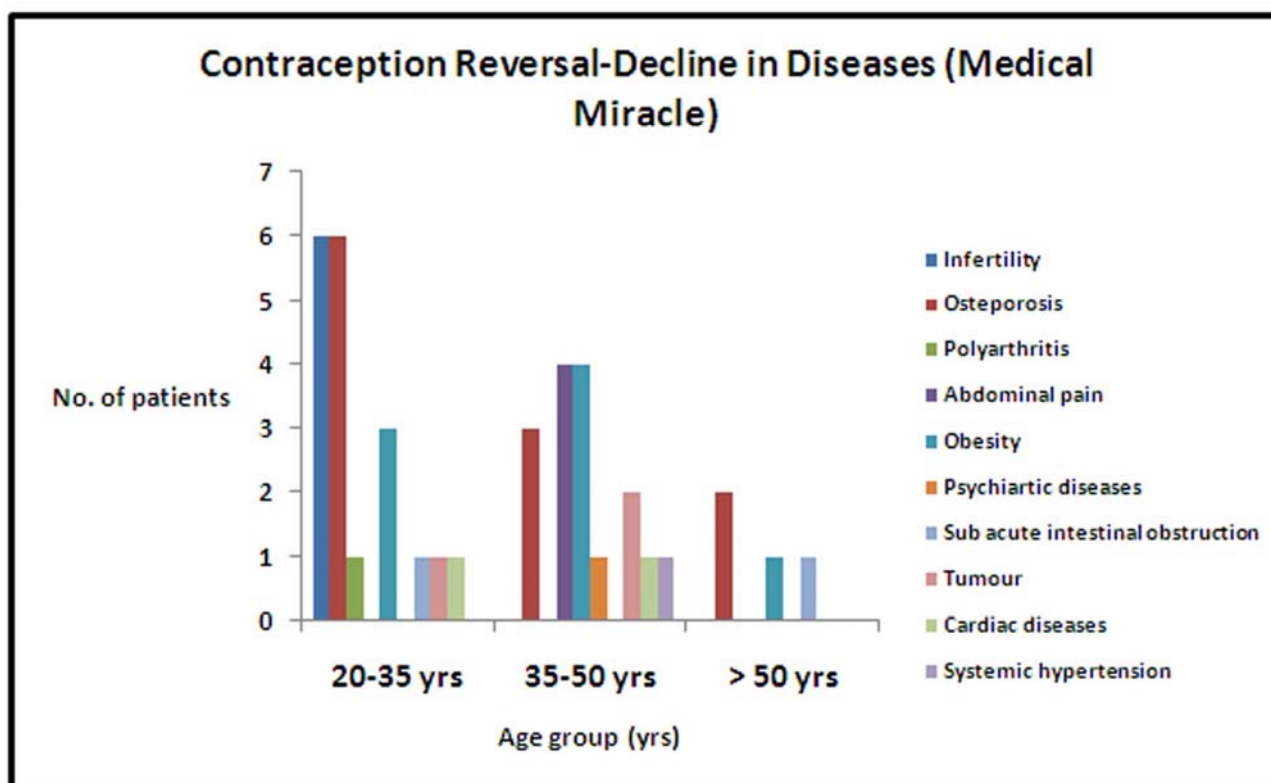
[2] C.M.C.Allen, C.J.Leuck, M.Dennis: Neurological Disease; Davidson’s Principles and Practice of Medicine, 21st edition 2010; page-1197-1198
 [3] Allan H Ropper, Robert H Brown: Adams and Victor’s Principles of Neurology 8th edition page 14-17
 [4] M.C.Sharpe, S.M.Lawrie: Medical Psychiatry, Davidson’s Principles and Practice of Medicine 21st edition 2010;page-245
 [5] William .F.Ganong: Physiology of nerve and muscle cells, Review of Medical Physiology; page 49-50
 [6] Bruce C Gilliland: Systemic Sclerosis, Harrison’s Principles of Internal Medicine, 11th edition, 1987 2nd volume; page1429
 [7] John MendelSohn: Principles of Neoplasia, Harrison’s Principles of Internal Medicine, 11th edition 1987, volume-1, page-422
 [8] Michael S Brown, Joseph L. Goldstein: Exogenous Pathway, endogenous pathway, Goodman and Gilman’s Pharmacological Basis of Therapeutics 7th edition; page-827-828

Tabulation

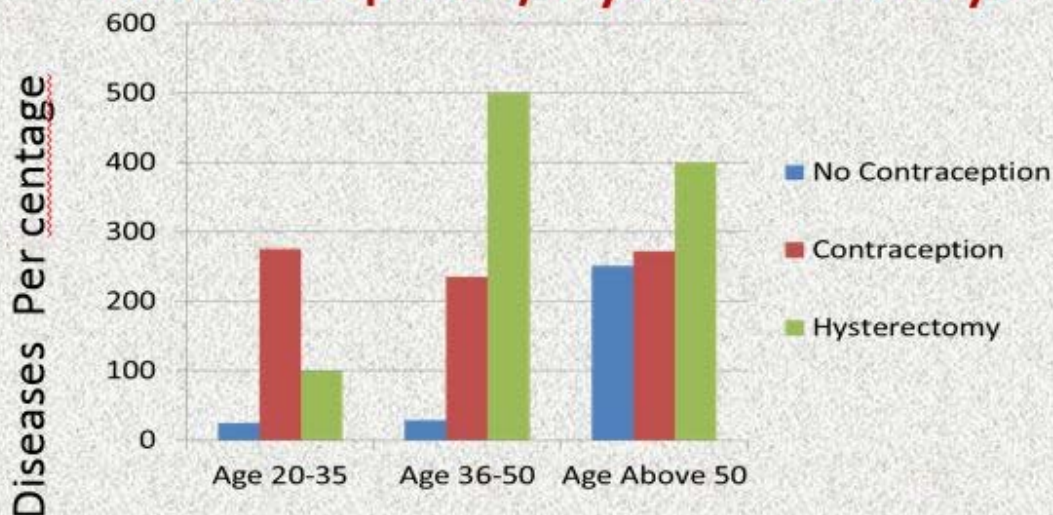
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| 36- 50yrs | contr acep | | | | | | | | | | | | | | | | |
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Diseases/morbidity- contraception/hysterectomy.



Tubal Recanalization – mandatory/essential - will return life factors – result in decreased diseases. Hysterectomy should be reserved only for PPH/Uterine Cancers. Hysterectomy to be replaced by tubal-recanalization, myomectomy, Pelvic Floor Repair.