

# Stochastic Process in Evolutionary Genetics ; Fokker Plank Diffusion Model-

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## Abstract

In Evolution, the genetic make up of species and population may change over time, some traits may be lost , other new one arises, while some persist unchanged, Ewens Sampling when the Sample Size is small, monocious population of varying size that reproduces in each generation according Wright-Fisher Model, if Natural selection, Hardy Weinber Law and a Markov Chain application in Genetics. then Weiner process for speciation is discussed.

Key Words: Wright Fisher Model, Hardy Weinberg Equilibrium, Ewens Sampling Branching Process, Natural Selection, Fokker Plank Diffusion Model.

## (1) Introduction

### 1.1 Stochastic Treatment: Discrete Process: Fisher Wright Model

Consider a diploid population having in each generation exactly  $N$  individuals, so there are  $2N$  genes altogether at the locus in question. Suppose that, in some ways to be chosen at out discussion, the population reproduce itself to form a daughter generation. Suppose there also the onece the daughter generation is formed , no further reproduction is possible for the parent generation. If there is no mutation, Selection gene frequencies tend to remain steady. It is reasonable to the number of  $A_1$  genes is  $X(t)$  then the number  $X(t+1)$  is a binomial variate with index  $2N$  and parameter  $X(t)/2N$ , explicitly given that ,  $X(t) = i$ , the probability  $p_{ij}$  that  $X(t+1) = j$  is given  $P_{ij} = \binom{2N}{j} (i/2N)^j (1-i/2N)^{2N-j}$  ,  $X(i)$  is a Markovian Variable with transition Matrix  $j$

$$P = \{p_{ij}\}$$

Wright Fisher (1931) introduced the concept of effective population number ,  $(N_e)$  for a population composed of equal number of males and females which would result in equivalent inbreeding or variances due to genetic drift, the object of this paper is population is is not panmix, Ewens(1972) Sampling procedure consequently and Watterson(1976) diffusion model Approximation ,.

### 1.2 Hardy Weinberg Law and a Markov Chain in genetics

- (1) According G. H. Hardy, Mendelian Population in a mixed population, Letter to the Editor, Science , N,S, Vol.28, (1908) pp:49-50, For a better under standing of the inequilibrium population , let us consider Hardy Weinberg Equilibrium (I,e) let us fix the gene frequencies,  $P$  and  $q$  ( $P+q=1$ ) , gene frequencies of  $A$  and  $a$  , In each of the two selection an  $A$  genes is selected with probability of an offspring being  $AA$  is  $p^2$ , the genotype  $Aa$  can occur in two ways is  $2pq$ ,  $aa$  with  $q^2$  probabilities.

Suppose the three genotypes AA, Aa, aa occur among males and females in the same ratio,  $u:2v:w$ ; we shall suppose  $u+2v+w=1$  say  $u:2v:w$  the genotype frequencies;  $p=u+v$ ,  $q=v+w$ . The genotype AA, Aa, aa with probabilities  $u^2=p^2$ ,  $2v^2=2pq$ ,  $w^2=q^2$

Hardy-Weinberg Equilibrium explics, Whatever the composition of the parent population produce an approximately stationary genotype distribution with unchanged gene frequencies. However equilibrium deviates.

### 1.3 :Wright Fisher and Moran Model

Evolution of the frequency of an allele A in a population which has the size N in each generation.

$Y_t$ ; Number of A alleles in generation 't'

State Space  $E = \{0, 1, 2, \dots, 2N\}$

Since the possible number of alleles lies between 0 to 2N.

The frequencies of the three genotypes in the nth generation are three random variables whose expected values are not in the ratio  $u^2:2v^2:w^2$  i.e the actual values will vary generation to generation lead for Stochastic Model in g.

## (2) Sampling

### 2.1 Ewens(1972) Sampling Formula

Data on variability in samples are of a type that are might call configurational. That is to say, we can encourage that a sample of r genes is taken from among the 2N that are such that variability doesn't necessarily implies that a particular small samples from a population will contain several alleles at substantial frequencies. If a sample of such a structure is found a question arises about where the presence of some favourably frequent alleles is due to selection or is merely what is to be expected from random sampling, that were present in the zygote and that it contains K alleles, which are respectively represented  $n_1, n_2, \dots, n_k$  times in the sample collection  $\{n_1, n_2, \dots, n_k\}$  as a configuration, whenever it is to be assumed that  $n_1 > n_2 > \dots > n_k$ , probability that a random samples of size ,r, contains the configuration  $\{n_1, n_2, \dots, n_k\}$  is equal to

$$P\{r; k; n_1, n_2, \dots, n_k\} = r! / (n_1! n_2! \dots n_k! l_1! \dots l_k! Q^k / (Q(Q+1)) \dots (Q+r-1)) \dots (1)$$

where  $l_j$  is the number of frequencies  $n_1, \dots, n_k$  that are equal to j,  $Q = 4Nu$  and N is the population size. What is the proper effective population Nuber  $N_e$  to substitute for N in(1)

Following Karlin(1968) and chia and Pollak(1974), we shall assume that the possible population sizes are the finite numbers  $N(1) \dots N(s)$  and that the sequence of sizes  $\{N_t, t = 0, 1, \dots\}$  is finite irreducible Markov Chain

### 3. Analysis

The associated transition Matrix is  $C = [c_{ij}]$ , where  $c_{ij} = P[N_{t+1} = N(j) / N_t = N(i)]$

#### 3.1 Estimating the transition probabilities;

Let  $X = \{x_1, x_2, \dots\}$  be a random process in the discrete state space  $S$ . the conditional probability  $\{x_{t+1} = x_{t+1} / x_t = x_t, x_{t-1} = x_{t-1}, \dots, x_1 = x_1\} = P_{x_1 x_2, x_2 x_3 \dots x_{t-1} x_t}$  the conditional probability of a path conditioned on the first value is the product of the transition probabilities between successive states of the path

#### 3.2 Estimating the changes in gene frequency

In polymorphism, a gene with  $n$  alleles there  $n(n+1)/2$  possible genotypes, the relationship between gene frequency and genotype frequency for a single gene at the population level can be used to infer the genetic states of the genes in population, if the genes and genotypic frequencies are constant from generation to generation.

Population genetics, is the study of the distribution and change in Frequency of alleles within population is known Evolutionary Biology. the four Process of Evolution

(i) Fishers' Natural Selection (2) Genetic Drift (3) Gene flow(Diffusion/ Wiener process) and (4) Mutation, (Continuous time, continuous state Fokker Plank Diffusion Model) is widely used.

(i) Three State Random Walk;  $p\{x_i = 1\} = p$  and  $p\{x_i = -1\} = 1-p$

Markov chain with transition probabilities

$$P_{ij} = \begin{cases} p & \text{if } j=i+1 \\ 1-p & \text{if } j=i-1 \\ 0 & \text{if otherwise} \end{cases}$$

Then  $x_{t+1} = x_t + e_{i+t}$ , while  $e_{i+t}$  and  $x_t$  are independent

Consider the transition probabilities as follows

		$x_{n+1}$		
		-1	0	1
	-1	$p_{-1-1}$	$p_{-10}$	$p_{-11}$
$X_n$	0	$p_{01}$	$p_{00}$	$p_{01}$
	1	$p_{1-1}$	$p_{10}$	$p_{11}$

Assume all frequency is  $p_{n1} = [1 - m_i] p_{n0} + m_i p_i$

$$= p_{n0} + (m_i(p_i - p_{n0}))$$

Change in allele frequency is  $p_{n1} - p_{n0} = (m_i(p_i - p_{n0}))$

(ii) Selection:

Consider the following Life cycle stage, their corresponding effect

Stage	Type of effect
Zygote formation to Adult	differential survival
Gametic	differential output, survival
Mating	Non random mating
Incompatibility between mates	
And parent and progeny	

Selection includes differential Fertility, Fecundity, viability all ages and differential emigration.

$$P_{n1} = f_1 p_{n0}^2 + f_2 p_{n0}(1-p_{n0})$$

$$F_1 p_{2n0} + 2f_2 p_{n0}(1-p_{n0}) + f_3(1-p_{n0})^2$$

Where  $p_{n0}$  is the allelic frequency for A in the population before selection and coefficient for selection is the measure of the disadvantage, those have least I.Q. has least chance of Survival.

Assuming random mating among AA and Aa and an appear in the total population be  $u, 2v, w$ . The corresponding frequencies for parents are the

$$U^* = u/1-w, \quad 2v^* = 2v/1-w, \quad w^* = u \quad \dots(1)$$

$$P = u+v/1-w, \quad q = 2v/1-w, \quad w^* = 0 \quad \dots(2)$$

The probabilities of the three genotypes in first filial generation are  $p_1 = p^2, 2v_1 = 2pq, w_1 = q^2$

$$\text{In general } p_n = u + v_n/1-w_n, \quad q_n = 1-w_1/1-w_n \quad \dots(3)$$

$$\text{And } u_{n+1} = p^2 n \quad ; \quad 2v_{n+2} = 2p_n q_n; \quad w_{n+1} = q_n^2 \quad \dots(4)$$

$$\text{From (3) and (4). } P_{n+1} = u_{n+1} + u_{n+1}/1-w_{n+1} = p_n/1-q_n^2 = 1/1+q_n \quad \dots(5)$$

$$Q_{n+1} = v_{n+1}/1-w_{n+1} = q_n/1+q_n \quad \dots(6)$$

From (6) we can calculate  $q_n$  explicitly taking reciprocal we get

$$q_{n+1}^{-1} = 1 + q_n^{-1} \quad \dots(7)$$

$$\text{hence substituting. } Q_1^{-1} = 1 + q^{-1}, \quad q_2^{-1} = 2 + q^{-1} \quad \dots(8)$$

$$q_n = q/1+nq, \quad w_{n+1} = ((q/1+q_n)^2) \quad \dots(9)$$

unproductive (or undesirable genes) genotype gradually drops out . the selection may be

(i) systematic effect, in which both the size and direction of the change are in principle

determinants

- (ii) Dispensive effect, for which the size is determinant, in principle but the direction of the change is not
- (iii) Non –recurrent Event, for which neither, size nor direction of change is determinate.
- (iv) Genetic Drift

Genetic drift refers to the chance changes in frequency of alleles from one generation to the next

### 3.3 Derivation of the two fundamental equations of Ewens Sampling

We shall first derive a recurrence equation that relates the probabilities of configurations in generations  $t$  and  $t+1$ . We assume that the Wright Fisher Model holds. Thus if  $N(t) = N(i)$  and  $N_{t+1} = N(v)$  the  $2N(i)$  genes among the zygote of the offspring generation are obtained by repeated sampling with replacement of the  $2N(i)$  genes of the parent generation. We also assume that as each gene is passed from the parent to the offspring it has a probability  $V$  of being is mutant to a type that didn't previously exist in the population. Then if a sample of  $r+1$  genes is taken from among the offspring and we denote by  $q(r+1, m/N(t) = N(i), N(t+1) = N(v))$ , the probability that this sample was transmitted from exactly  $m$  distinct parent genes,

$$\begin{aligned}
 & q(r+1, r+1/N_{t=N(i)}, N_{t+1} = N(v)), \dots(2) \\
 & = 2N(i)(2N(i)-1) \dots (2N(i)-r) / \{2N(i)\}^{r+1} \\
 & = 1-r(r+1)/4N(i) + O(N(i)^{-2}) \dots(3)
 \end{aligned}$$

$$\begin{aligned}
 & q(r+1, r/N_{t=c=N(i)}, N_{t+1} = N(v)) \\
 & = r+1 C_{2N(i)}(2N(i)-1) \dots (2N(i)-R+1) / (2N(i))^{R+1} \\
 & = R(R+1)/4n(I) + O[9n(I)^{-2}] \dots(4)
 \end{aligned}$$

as  $N(i) \rightarrow \infty$ , we obtain expression (4) because  $r+1 C_{2N(i)}$  is the number of ways to choose 2 genes from among  $r+1$  that were derived from the same parent and  $2N(i)$  that were derived from the same parent and  $2N(i)(2N(i)-1) \dots (2N(i)-r+1)$  is the number of ways to select  $r$  distinct parental genes and a specified repeated parent from among  $2N(i)$ ,

Now let  $P(t+1(r+1; k; n_1, \dots, n_k) / m, N(i), N(v))$  and  $P_{t+1}(r+1; k; n_1, \dots, n_k / N(i), N(v))$  respectively denote conditional probabilities of the offspring configuration  $\{n_1, \dots, n_k\}$  given  $m, N(i) \dots N(v)$  and  $N(i), N(v)$ .

It follows from (3) and (4) that,  $P(t+1(r+1; k; n_1, n_2 \dots n_k / N(i), N(v))$

$$\begin{aligned}
 & = \{1-r(r+1)/4N(i)\} P_{t+1}(r+1; k; n_1, n_2 \dots n_k / r+1), N(i), \dots, N(v) + r(r+1)/4N(i) \\
 & P_{t+1}(r+1; k; n_1 \dots n_k / r, N(i), N(v)) \dots(5)
 \end{aligned}$$

if the possible population sizes are all large. we shall now calculate the probabilities on the right side of (5), if  $l=0$ . In this case the  $r+1$  offspring genes must be unmated couples of the parental genes. Thus  $P_{t+1}(r+1; k; n_1, \dots, n_k / r+1, N(i), N(v))$

$$= (1-u)^{r+1} p_{t+1}(r+1; k; n_1, \dots, n_k) \dots(6)$$

The second conditional configuration, probability of  $\{A_1, \dots, A_k\}$  is the probability of an event that, can occur in several mutually exclusive ways. Each of these is associated with  $r$  parental genes one of which is used as parent twice, on way is to have the parental configuration  $\{n_1, \dots, n_{j-1}, \dots, n_{j+1}, \dots, n_k\}$ . the probability of this is  $\{l(n_j)\}^{-1}$  times as large  $P(r; k; n_1, \dots, n_{j-1}, n_{j+1}, \dots, n_k)$  if there are  $l(n_j)$  offspring genes in a sample of size of size  $r+1$  that are represented  $n_j$  times. Now, if the offspring genes represented  $n_{j-1}$  times are  $l(n_{j-1})$  in number, there are  $l(n_{j-1})+1$  if such alleles among the parents.

Hence, some parental alleles represented  $n_{j-1}$  times is chosen to produce two copies with probability  $(n_{j-1})/(l(n_{j-1})+1)/r$ , if parental genes are randomly chosen to be replicated twice. Hence

$$P_{t+1}(r+1; k; n_1, \dots, n_k / r, N(i), N(v)) = (n_{j-1})/(l(n_{j-1})+1)/r \cdot l(n_j) \cdot X_{Pt}(r; k; n_1, \dots, n_{j-1}, n_{j+1}, \dots, n_k / N(i)) \dots (7)$$

Therefore, if we combine (2), (5), (6) and (7) we obtain

$$P_{t+1}(r+1; k, n_1, \dots, n_k / N(i), N(v)) P(N_t = N(i)) C_{1v} = P_{t+1}(r+1; k, n_1, \dots, n_k / N(i), N(v)) \cdot 0((N_t = N(i)) C_{1v})$$

$$P(N_t = N(i)) C_{1v} (1 - (r+1)/4N(i) - n_{j-1} X_{l(n_{j-1})+1} / r l(n_j) X_{Pt}(r; k; n_1, \dots, n_{j-1}, n_{j+1}, \dots, n_k) / N(i))$$

it applies if  $l_i = 0$

Now we shall now derive the second if the fundamental equation, which only refer to configuration within one generation. To do this, a sample of  $r+1$  genes in generation  $t$  will be looked upon as consisting of two parts. first ‘ $v$ ’ genes are drawn and next the resulting subsample is supplement by the drawing of one more gene. with random sampling the configuration  $\{n_1, \dots, n_k\}$  among the first  $r$  genes is  $1/(r+1)$  times as probable all sets of  $r+1$  genes containing this as a subset of  $v$ . also one of the ways to have  $\{n_1, \dots, n_k\}$  among the first  $r$  genes is to have it followed by a gene represented  $n_{j+1}$  times among  $r+1$  genes and  $n_j$  times any  $r$ . thus, if  $l(n_i)$  is equal to the number of alleles represented  $n_j$  times among the  $r$  genes, each of these ways is  $1/l(n_j)$  times as probable as all configurations of the type  $\{n_1, n_2, \dots, n_{j-1}, n_{j+1}, \dots, n_k\}$

finally, there are  $(n_{j+1})[l(n_{j+1})+1](n_{j+1})/l(n_j)$  ways to pick a gene times in the sample, therefore,

$$P_t(r; k, n_1, \dots, n_k) = \{1/n_{j+1} + 1(n_{j+1})/l(n_j)(r+1) X_{Pt}(r+1; k, \dots, n_{j-1}, n_j, n_{j+1}, \dots, n_k) + 1 + 1/r + 1 P_t(r+1, k+1, n, n_k, l)$$

where  $R_i$  is the number of alleles represented once among the first ‘ $r$ ’ genes.

Ewens sampling when the sample size is small in comparison with that of the population selection plays a negligible role, Mutation is nonrecurrent and the population is at equilibrium under mutation random drift.

Ewens Sampling Formula:  $Q = 4Na$ , where  $N$  is the population size

To determine the overall frequency  $P$  for a given generation,

$$P = \frac{1}{4}(p_{mm} + p_{mf} + p_{fm} + p_{ff})$$

Where  $p_{yz}$  represent the allele frequencies transmitted from the  $y$  sex of parents to the  $z$  sex of offspring chosen to be used as parents with  $m$ =male and  $f$ =female when the chance occurrences along these pathways are independent, we can represent by

$$V_p = \frac{1}{16}(V_{pmm} + v_{pmf} + v_{pfm} + v_{pff})$$

Where  $v_{pyz}$  is the variance of the frequency along each path. In an idealized reference situation of random sampling with replacement of alleles the variances would be binomial with

$$V_{pyz} = p_{yz}(1-p_{yz})/2N_{eyz}$$

$$\text{And } V_p = p(1-p)/2N_e$$

Where  $N_{eyz}$  is the effective population number for specific path, sex  $y$  to sex  $Z$  and  $N_e$  is the overall effective number. Using the combinatorial mathematics, what is the proper effective population number  $N_e = [1/d \sum_i n_i(N_i - 1)]$ .

If there is a noncoincidental population of varying size that reproduce in each generation according to the Wright Fisher Model, Karlin (1968) and Coir and Pollak (1974), we shall assume. The population sizes are the finite numbers  $N(1) \dots N(s)$  and that the queue of sizes  $\{N_t, t=0,1,\dots\}$  is a finite irreducible Markov chain, the associative transition matrix is  $C = \{c_{iv}\}$  where

$$C_{iv} + P\{N_{t+1} = N(v)/N(N_t) = N(1)\}$$

Which apply to theoretical population as well as Wright-Fisher Reproduction population structure.

Effective Breeding Population Size, For Structural Random Mating with Random or directional Selection pioneer work of Wright (1931) introduced the concept of effective population number to reflect the magnitude of expected random variation and fixation in gene frequency due to finite size of population known as effective population number ( $N_e$ ) for a population composed of  $N_m$  breeding males and  $N_f$  breeding female was  $1/N_e = 1/4N_m + 1/4N_f$

Four pathways allele pass between generation (i) male to male (ii) Male to female

(iii) Female to male (iv) female to female, which leads to four types of Stochastic process

### 3.4 Fisher's Natural Selection

**The essence of the theory of evolution through the selection is that in any population there will exist genetic variation between individuals and that those genotypes which are better suited to the environment than other will contribute rather more than their fair share of offspring to the following generation. Thus the genetical make-up of the following generation will differ somewhat from that of the parent generation leading to substantial changes over large number of generation. such evolution depends as the genetical variation in the population, so that it might be expected that the greater variation, the greater will be the changes which occur. Further it appears that in some sense the process leads to the improvement in the population. Selection differences among genotypes generally leads to changes in gene frequencies.**

## IV Conclusion

### Fokker Plank Diffusion Model

Continuous state continuous Parameter Diffusion Approximation of Fokker plank Diffusion Model. May be studied

Consider the random variable differing over  $(0,1)$ , in such a way that if at any time,  $t$ , the random variable assumes the value  $x$ , then the value  $x+x$  assumed at time  $t+dt$  is a random variable

$$E(dx) = m(x)dt + o(dt)^2$$

$$V(dx) = v(x)dt + o(dt)^2$$

$$E(dx) = o(dt)^i \quad i > 3$$

Here  $m(x)$  and  $v(x)$  are function of  $x$ , but not of  $t$ , and are called respectively the drift and diffusion co-efficient of the process. If  $f(x;t)$  is the probability density of the random variable at time  $t$ , the theorem of total probability shows

$$f(x; t+ t) = \int F(x-dx;t)g(dx; x-dx).d(dx)$$

where  $g(dx; x-dx)$  is the probability density of change in the value of the random variable from  $x-dx$  to  $x$  in the time interval  $(t, t+dt)$

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