DIFFUSION APPROXIMATION OF NON-MARKOVIAN PROCESSES

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General diffusion approximation theorems are established for sequences of non-Markovian processes. These theorems cover certain genetic models previously considered by Watterson. It follows that Watterson's conclusions concerning these models are correct, even though there is a gap in his proof.

1. Introduction. Let $I=[d_0,d_1]$ be a closed bounded interval. For each $N\geq 1$, let $\{X_n^N,\,n\geq 0\}$ be a stochastic process in I, adapted to an increasing sequence $\{\mathscr{F}_n^N,\,n\geq 0\}$ of σ -fields. The processes need not be Markovian. The conditional moments of $\Delta X_n^N=X_{n+1}^N-X_n^N$ are supposed to satisfy conditions of the form

(1)
$$\begin{split} E(\Delta X_n^N | \mathscr{F}_n^N) &= \tau_N a(X_n^N) + e_{1,n}^N ,\\ E((\Delta X_n^N)^2 | \mathscr{F}_n^N) &= \tau_N b(X_n^N) + e_{2,n}^N ,\\ E(|\Delta X_n^N|^3 | \mathscr{F}_n^N) &= e_{3,n}^N , \end{split}$$

where $\tau_N > 0$ and $\tau_N \to 0$ as $N \to \infty$, and the error terms $e_{i,n}^N$ are $o(\tau_N)$ in the sense that, for any $t < \infty$,

(2)
$$\sum_{n<[t/\tau_N]} E(|e_{i,n}^N|) \to 0 \quad \text{as } N \to \infty.$$

Let $X^N(t) = X^N_{[t/\tau_N]}$, let $0 \le t_1 < t_2 < \cdots < t_k$, and let \Rightarrow denote convergence in distribution. Our main result, Theorem 1, gives conditions on a and b that insure that $(X^N(t_1), \cdots, X^N(t_k)) \Rightarrow (X(t_1), \cdots, X(t_k))$ as $N \to \infty$ and $X^N(0) \Rightarrow X(0)$, where X(t) is a diffusion whose transition kernel, $P(t; x, A) = P(X(t) \in A \mid X(0) = x)$, satisfies the following conditions, which are analogous to (1):

(3)
$$\int_{I} (y - x)P(\tau; x, dy) = \tau a(x) + e_{1}(\tau, x), \\
\int_{I} (y - x)^{2}P(\tau; x, dy) = \tau b(x) + e_{2}(\tau, x), \\
\int_{I} |y - x|^{3}P(\tau; x, dy) = e_{3}(\tau, x).$$

Here the error terms $e_i(\tau, x)$ are $o(\tau)$ in the uniform sense:

(4)
$$\sup_{x \in I} |e_i(\tau, x)|/\tau \to 0 \qquad \text{as } \tau \to 0.$$

Let C^j be the set of functions with j continuous derivatives throughout I. At the boundaries d_i these derivatives are one-sided.

THEOREM 1. Suppose that $a \in C^3$, $a(d_0) \ge 0$, and $a(d_1) \le 0$; and that b admits a factorization $b(x) = \sigma_0(x)\sigma_1(x)$, where σ_i satisfies the following conditions: $\sigma_i \in C^3$,

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 $\sigma_i(d_i) = 0, \ \sigma_i(x) > 0 \ for \ d_0 < x < d_1, \ p(x) = \sigma_0(x)/(\sigma_0(x) + \sigma_1(x)) \ is \ non-decreasing for \ d_0 < x < d_1, \ and, \ letting \ p(d_i) = \lim_{x \to d_i} p(x), \ p \in C^3. \ Then \ there \ is \ a \ unique transition kernel P \ satisfying (3) \ and (4). \ If \ X^N(0) \Rightarrow X(0) \ as \ N \to \infty, \ then (X^N(t_1), \dots, X^N(t_k)) \Rightarrow (X(t_1), \dots, X(t_k)) \ as \ N \to \infty.$

Though the condition on b in Theorem 1 is undesirably complicated, it is not very restrictive in applications. The condition implies that $b \in C^3$, $b(d_i) = 0$, and b(x) > 0 for $d_0 < x < d_1$. If, conversely, b is analytic throughout I with $b(d_i) = 0$ and b(x) > 0 on the interior of I, the condition of Theorem 1 is satisfied. For let j_i be the order of the zero at d_i . Then $h(x) = b(x)(x - d_0)^{-j_0}(d_1 - x)^{-j_1}$ is analytic and positive throughout I, so $h(x)^{\frac{1}{2}} \in C^3$. Thus we can take $\sigma_0(x) = (x - d_0)^{j_0}h(x)^{\frac{1}{2}}$ and $\sigma_1(x) = (d_1 - x)^{j_1}h(x)^{\frac{1}{2}}$. In the genetic applications considered in Section 4, I = [0, 1], a is a polynomial, and b(x) = cx(1 - x), where c > 0.

Theorem 1 is an extension of Theorem 9.1.1 of [5] to non-Markovian processes. The latter theorem includes the existence and uniqueness of P. It is noteworthy that the uniformity condition (4) can be used in place of conventional boundary conditions to determine P uniquely. The proof of Theorem 9.1.1 shows that the equation

$$T_t f(x) = \int_I f(y) P(t; x, dy)$$

defines a strongly continuous conservative semigroup on $C=C^0$ (supremum norm). Let Γ be the generator of this semigroup, and let $\mathscr{D}(\Gamma)$ be the domain of Γ . Then T_t is the unique strongly continuous conservative semigroup on C for which $\mathscr{D}(\Gamma) \supset C^2$ and

$$\Gamma f(x) = a(x)f'(x) + 2^{-1}b(x)f''(x)$$

for $f \in C^2$ and $x \in I$ ([5] page 150). In terms of the Feller boundary theory, exit boundaries are adhesive $(\Gamma f(d_i) = 0 \text{ for } f \in \mathcal{D}(\Gamma))$ and regular boundaries are reflecting $((d/dp)f(d_i) = 0 \text{ for } f \in \mathcal{D}(\Gamma)$, where $dp(x) = e^{-B(x)} dx$ and $dB(x) = 2a(x)b(x)^{-1} dx$ ([5] page 148).

The possibility and desirability of extending Theorem 9.1.1 of [5] to non-Markovian processes was suggested by an interesting paper of Watterson [6]. Watterson's result is similar to the special case of Theorem 1 corresponding to k=1, b(x)=cx(1-x) and a(x) a third degree polynomial. Unfortunately, there is a large gap in Watterson's proof. Let $F_N(x,n)=P(X_n^N\leq x)$ and $F(x,u)=P(X(u)\leq x)$. The transition from

$$\lim_{N\to\infty} \int_0^\infty e^{-\alpha u} F_N(x, [N^m u]) du = \int_0^\infty e^{-\alpha u} F(x, u) du$$

to

(5)
$$\lim_{N\to\infty} F_N(x, [N^m u]) = F(x, u)$$

at the bottom of page 950 of [6] cannot be justified by "the uniqueness theorem for Laplace transforms." (Watterson's N^m corresponds to our τ_N^{-1} .) Standard continuity theorems ([3] page 433) yield, not (5), but the integrated version

$$\lim_{N\to\infty} \int_0^t F_N(x, [N^m u]) du = \int_0^t F(x, u) du$$
.

Watterson's paper is oriented toward applications to two genetic models, one with overlapping generations, the other with non-overlapping generations. These applications are described in detail in a later paper [7]. In Section 4 we will show that these models fall within the scope of Theorem 1. Guess [4], using Watterson's result as a lemma, proved weak convergence of the distribution of the process $\{X^N(t), t \ge 0\}$ for a class of models that includes the non-overlapping generation model. Theorem 2 of Section 3 is a weak convergence theorem that complements Theorem 1 and applies to both of the models considered by Watterson.

2. Proof of Theorem 1. We first show that, for $n \ge m$ and $f \in C$,

(6)
$$E[|E(f(X_n)|\mathscr{F}_m) - T_{(n-m)\tau}f(X_m)|] \le \sum_{i=m}^{n-1} E[|E(g_{i+1}(X_{i+1})|\mathscr{F}_i) - T_{\tau}g_{j+1}(X_j)|],$$

where $g_j = T_{(n-j)\tau}f$ and N's have been suppressed. Clearly

$$f(X_n) - T_{(n-m)\tau} f(X_m) = g_n(X_n) - g_m(X_m) = \sum_{j=m}^{n-1} (g_{j+1}(X_{j+1}) - g_j(X_j)).$$

Hence

$$\begin{split} E(f(X_n) | \mathscr{F}_m) - T_{(n-m)\tau} f(X_m) &= \sum_{j=m}^{n-1} E[g_{j+1}(X_{j+1}) - g_j(X_j) | \mathscr{F}_m] \\ &= \sum_{i=m}^{n-1} E[E(g_{j+1}(X_{i+1}) | \mathscr{F}_i) - g_j(X_j) | \mathscr{F}_m] \,. \end{split}$$

Taking absolute values and expectations on both sides of this equality, and noting that $g_i = T_r g_{i+1}$, we obtain (6).

Let L be the subspace of C^2 consisting of those functions whose second derivatives satisfy the Lipschitz condition

$$M(g'') = \sup_{x \neq y} \frac{|g''(x) - g''(y)|}{|x - y|} < \infty.$$

For $g \in L$, let

$$||g|| = |g'|_{\infty} + |g''|_{\infty} + M(g'')$$
,

where $|\cdot|_{\infty}$ is the supremum norm. Any function g in L possesses a Taylor expansion

$$g(y) = g(x) + (y - x)g'(x) + 2^{-1}(y - x)^{2}g''(x) + \lambda |y - x|^{3}M(g''),$$

where $|\lambda| \leq \frac{1}{6}$. For the remainder in the first order Taylor expansion is

$$g(y) - g(x) - \delta g'(x) = \delta^2 \int_0^1 (1 - s)g''(x + s\delta) ds$$

= $2^{-1}\delta^2 g''(x) + \delta^2 \int_0^1 (1 - s)(g''(x + s\delta) - g''(x)) ds$,

where $\delta = y - x$, and the last term on the right has absolute value at most

$$|\delta|^3 M(g'') \int_0^1 (1-s) s \, ds = 6^{-1} |\delta|^3 M(g'')$$
.

Hence, in view of (1),

$$E(g(X_{j+1}) \mid \mathscr{F}_j) = g(X_j) + \tau \Gamma g(X_j) + \lambda ||g|| \sum_{i=1}^3 |e_{i,j}|,$$

where $|\lambda| \leq 1$. By (3), $T_{\tau}g$ has a similar expansion, so

(7)
$$E[|E(g(X_{j+1})|\mathscr{F}_j) - T_{\tau}g(X_j)|] \le ||g|| \sum_{i=1}^3 [E(|e_{i,j}|) + \sup_{x \in I} |e_i(\tau, x)|].$$

The following lemma is a by-product of the proof of Theorem 9.1.1 (see [5] (3.8) page 150). We shall have more to say about it at the end of the section.

LEMMA 1. T_t maps C^3 into L. Moreover, for any $f \in C^3$ and $K < \infty$, $\sup_{t \le K} ||T_t f|| < \infty$.

Applying (7) to $g_{j+1} = T_{(n-j-1)\tau} f$ for $f \in C^3$, using Lemma 1 to estimate $||g_{j+1}||$, and combining the result with (6), we obtain

(8)
$$E[|E(f(X_n^N)|\mathcal{F}_m^N) - T_{(n-m)\tau}f(X_m^N)|] \\ \leq K' \sum_{i=1}^3 [(n-m) \sup_{x \in I} |e_i(\tau_N, x)| + \sum_{j=m}^{n-1} E(|e_{i,j}^N|)]$$

for some constant K', provided that $(n-1)\tau_N \leq K$.

Suppose now that $0 \le s \le t$ and let $n = [t/\tau_N]$ and $m = [s/\tau_N]$. As a consequence of (2) and (4), the quantities on the right and left in (8) approach 0 as $N \to \infty$. But $(n-m)\tau \to t-s$ as $N \to \infty$, and, as noted in Section 1, the semigroup T_t on C is strongly continuous with respect to the supremum norm, so $T_{(n-m)\tau}f(x) \to T_{t-s}f(x)$, uniformly over x, as $N \to \infty$. Thus

(9)
$$E[|E(f(X^{N}(t))| \mathcal{F}^{N}(s)) - T_{t-s}f(X^{N}(s))|] \to 0$$

as $N \to \infty$, where $\mathscr{F}^N(s) = \mathscr{F}^N_{[s/\tau]}$. Since C^3 is dense in C, (9) holds for all $f \in C$. It follows from (9) that

$$E(f(X^{N}(t))) - E(T_{t-s}f(X^{N}(s)))$$

$$= E[E(f(X^{N}(t)) | \mathscr{F}^{N}(s)) - T_{t-s}f(X^{N}(s))] \to 0.$$

Taking s = 0, and noting that $T_t f \in C$, so that

$$E(T_t f(X^N(0))) \to E(T_t f(X(0)))$$

= $E(f(X(t)))$,

we obtain $E(f(X^N(t))) \to E(f(X(t)))$ as $N \to \infty$.

Suppose, inductively, that $(X^N(t_1), \dots, X^N(t_k)) \Rightarrow (X(t_1), \dots, X(t_k))$ for some $k \ge 1$. If $f_i \in C$ for $1 \le i \le k+1$, and $0 \le t_1 < t_2 < \dots < t_{k+1}$, then

$$E[\prod_{i=1}^{k+1} f_i(X^N(t_i))] = E[\prod_{i=1}^k f_i(X^N(t_i)) E(f_{k+1}(X^N(t_{k+1})) | \mathscr{F}^N(t_k))]$$

= $E[\prod_{i=1}^k f_i(X^N(t_i)) T_{t_{k+1}-t_k} f_{k+1}(X^N(t_k))] + \delta_N$,

where $\delta_N \to 0$ by (9),

$$\to E[\prod_{i=1}^k f_i(X(t_i)) T_{t_{k+1}-t_k} f_{k+1}(X(t_k))],$$

by the induction hypothesis,

$$= E\left[\prod_{i=1}^{k+1} f_i(X(t_i))\right].$$

Thus k in the induction hypothesis can be replaced by k + 1, and the proof of Theorem 1 is complete.

Since the boundedness assertion of Lemma 1 is the key to the proof, it is worthwhile to review briefly how this boundedness was established in [5]. This review also gives some insight concerning the origin of the assumption concerning b in Theorem 1. Let $V_{\tau}f(x)=E(f(X_{n+1}^{\tau})|X_n^{\tau}=x)$ for the discrete parameter Markov process X_n^{τ} that moves from x to $x+\tau a(x)+\tau^{\frac{1}{2}}\sigma_1(x)$ with probability $p(x)=\sigma_0(x)/(\sigma_0(x)+\sigma_1(x))$ and from x to $x+\tau a(x)-\tau^{\frac{1}{2}}\sigma_0(x)$ with probability 1-p(x). It was shown by direct calculation that there is a constant γ such that $||V_{\tau}^nf|| \le e^{\gamma n\tau}||f||$ for $\tau>0$, $n\ge 0$, and $f\in C^3$ ([5] Lemma 2.2, page 142). But $V_{\tau}^nf\to T_tf$ uniformly as $\tau\to 0$ and $n\tau\to t$, and it follows that $||T_tf||\le e^{rt}||f||$. We remark that it would be desirable to have an alternative proof of Lemma 1 based on semigroup and differential equation theory. For an illustration of such methods in a similar context, see [2].

3. Weak convergence of the distribution of $X^N(\cdot)$. For any K > 0, let D_K be the space of real-valued functions on [0, K] that are right-continuous and have left-hand limits. Let D_K be equipped with the Skorohod J_1 topology ([1] Section 14), and let \Rightarrow denote convergence in distribution for random elements of D_K .

Theorem 2. Suppose that the hypotheses of Theorem 1 hold, and that, in addition, there are constants G_i such that

$$|E(\Delta X_n^N | \mathscr{F}_n^N)| \leq G_1 \tau_N$$

and

(11)
$$\operatorname{Var}\left(\Delta X_{n}^{N} \mid \mathscr{F}_{n}^{N}\right) \leq G_{2} \tau_{N}$$

a.s., for $N \ge 1$ and $n \ge 0$. Then, for any K > 0, $\{X^N(t), t \le K\} \Longrightarrow \{X(t), t \le K\}$ as $N \to \infty$.

PROOF. According to Theorem 15.6 of [1], it suffices to show that there is a constant $H = H_K$ such that

$$E[(X^{N}(t) - X^{N}(t_{1}))^{2}(X^{N}(t_{2}) - X^{N}(t))^{2}] \leq H(t_{2} - t_{1})^{2}$$

for all $0 \le t_1 \le t \le t_2 \le K$. For this it is sufficient that

(12)
$$E[(X_n^N - X_m^N)^2 | \mathscr{F}_m^N] \le H'(n-m)\tau$$

for $0 \le m \le n \le K/\tau_N$.

Following Guess ([4] page 294), we write

(13)
$$X_n - X_m = \sum_{j=m}^{n-1} V_j + \sum_{j=m}^{n-1} W_j,$$

where

$$V_j = E(\Delta X_j | \mathcal{F}_j)$$

and

$$W_{j} = \Delta X_{j} - E(\Delta X_{j} | \mathcal{F}_{j}) .$$

By (10),

(14)
$$(\sum_{j=m}^{n-1} V_j)^2 \le G_1^2 (n-m)^2 \tau^2$$

$$\le G_1^2 K(n-m)\tau.$$

By (11)

$$E(W_j^2 | \mathscr{F}_m) = E(\operatorname{Var}(\Delta X_j | \mathscr{F}_j) | \mathscr{F}_m) \leq G_2 \tau$$

hence

(15)
$$E((\sum_{j=m}^{n-1} W_j)^2 | \mathscr{F}_m) = \sum_{j=m}^{n-1} E(W_j^2 | \mathscr{F}_m) \\ \leq G_2(n-m)\tau.$$

Combining (13), (14), and (15) we obtain (12).

4. Two genetic models. The models considered by Watterson [7] are relevant to a population of N diploid individuals. Of these, N_1 are males and N_2 are females. The three genotypes, aa, aA, and AA have frequencies k, $N_1 - k - l$, and l among males, and r, $N_2 - r - s$, and s among females. The successive values of the vector (k, l, r, s) form a Markov process in both models. However, interest centers on the average

$$X = 2^{-1} + 4^{-1}N_1^{-1}(k-l) + 4^{-1}N_2^{-1}(r-s)$$

of the relative frequencies of the a gene in the two sexes, and the trajectory of this variate is non-Markovian.

Variations in X are controlled by two mutation parameters, α_1 and α_2 , two selection parameters, ν_1 and ν_2 , and a nonrandom-mating parameter f. The latter is fixed as $N \to \infty$, but the former are assumed to be inversely proportional to N; $N\alpha_i = \bar{\alpha}_i \ge 0$ and $N\nu_i = \bar{\nu}_i$ are constant. Moreover, $N_i/N = r_i > 0$ is fixed. Let X_n be the value of X after n steps, and let \mathscr{F}_n be the σ -field generated by the values of k, l, r, and s after j steps, $j \le n$.

The significance of a single step is different in the two models. One is of the Moran type, with overlapping generations. Each step of the process corresponds to the death of a single individual and the birth of another. The other model is of the Wright-Fisher type. Generations are non-overlapping and the entire population is replaced at each step of the process. The step-size parameters for the overlapping and non-overlapping generation models are, respectively, $\tau_N = N^{-2}$ and $\tau_N = N^{-1}$.

In both models, the function a of Theorem 1 is

$$a(x) = \tilde{\alpha}_2(1-x) - \tilde{\alpha}_1 x - x(1-x) \{ \tilde{\nu}_1[(1-f)x + f] + \tilde{\nu}_2[(1-f)(1-x) + f] \}.$$

For the overlapping generation model,

$$b(x) = 4^{-1}(r_1^{-1} + r_2^{-1})(1+f)x(1-x),$$

while b(x) is half the quantity on the right for the other model. Lemmas 2 and 3 give estimates of the error terms $e_{i,n}^N$ in (1).

LEMMA 2. For the overlapping generation model,

$$E(|e_{i,n}^N|) \leq c\tau_N^{\frac{5}{4}} + c'e^{-n/2N}\tau_N$$

for i = 1 and 2, $n \ge 0$, and all N. Also

$$E(|e_{3,n}^N|) \leq c \tau_N^{\frac{3}{2}}.$$

LEMMA 3. For the non-overlapping generation model, $|e_{i,n}^N| \leq G\tau_N$ a.s. for i=1 and 2 and $n \geq 0$, $E(|e_{1,n}^N|) \leq c\tau_N^{\frac{3}{2}}$ for $n \geq 2$, $E(|e_{2,n}^N|) \leq c\tau_N^{\frac{3}{2}}$ for $n \geq 1$, and $E(|e_{3,n}^N|) \leq c\tau_N^{\frac{3}{2}}$ for $n \geq 0$. In all cases these estimates hold uniformly over N.

These estimates can be obtained by lengthy but, for the most part, straightforward calculations, using the suggestions in [7]. It follows immediately from these estimates that (2) is satisfied. Moreover it is very easy to show that (10) and (11) hold. Thus Theorems 1 and 2 apply to both models. We conclude that the gap in Watterson's proof does not alter the correctness of his conclusion that diffusion approximation is applicable to these models.

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