Capacity and Expressiveness of Genomic Tandem Duplication

Siddharth Jain, *Student Member, IEEE*, Farzad Farnoud (Hassanzadeh), *Member, IEEE*, and Jehoshua Bruck, *Fellow, IEEE*

***Abstract*— The majority of the human genome consists of repeated sequences. An important type of repeated sequences common in the human genome are tandem repeats, where identical copies appear next to each other. For example, in the sequence $AGTCTGTGC$, $TGTG$ is a tandem repeat, that may be generated from $AGTCTGC$ by a tandem duplication of length 2. In this paper, we investigate the possibility of generating a large number of sequences from a *seed*, i.e. a small initial string, by tandem duplications of bounded length. We study the capacity of such a system, a notion that quantifies the system's generating power. Our results include *exact capacity* values for certain tandem duplication string systems. In addition, motivated by the role of DNA sequences in expressing proteins via RNA and the genetic code, we define the notion of the *expressiveness* of a tandem duplication system as the capability of expressing arbitrary substrings. We then *completely* characterize the expressiveness of tandem duplication systems for general alphabet sizes and duplication lengths. In particular, based on a celebrated result by Axel Thue from 1906, presenting a construction for ternary squarefree sequences, we show that for alphabets of size 4 or larger, bounded tandem duplication systems, regardless of the seed and the bound on duplication length, are not fully expressive, i.e. they cannot generate all strings even as substrings of other strings. Note that the alphabet of size 4 is of particular interest as it pertains to the genomic alphabet. Building on this result, we also show that these systems do not have full capacity. In general, our results illustrate that duplication lengths play a more significant role than the seed in generating a large number of sequences for these systems.**

***Index Terms*— Capacity, expressiveness, tandem repeats, tandem duplication, finite automaton, irreducible strings.**

I. Introduction

MORE than 50% of the human genome consists of repeated sequences [9]. Two important types of common repeats are i) interspersed repeats and ii) tandem repeats. Interspersed repeats are caused by transposons. A transposon, also known as a jumping gene, is a segment of DNA that can copy or cut and paste itself into new positions of the genome. Tandem repeats are caused by slipped-strand mispairings [13]. Slipped-strand mispairings occur when one DNA strand in the duplex becomes misaligned with the other.

Tandem Repeats are common in both prokaryote and eukaryote genomes. They are present in both coding and non-coding regions and are believed to be the cause of several genetic disorders. The effects of tandem repeats on several biological processes is understood by these disorders. They can result in generation of toxic or malfunctioning proteins, chromosome fragility, expansion diseases, silencing of genes, modulation of transcription and translation [16] and rapid morphological changes [5].

A process that leads to tandem repeats, e.g. through slipped-strand mispairing, is called *tandem duplication*, which allows substrings to be duplicated next to their original position. For example, from the sequence $AGTCGTCGCT$, a tandem duplication of length 2 can give $AGTCGT\underline{CGCG}CT$, which, if followed by a duplication of length 3 may give $AGTCG\underline{TCGTCG}\ CGCT$. The prevalence of tandem repeats in the human genome [9] motivates us to study the capacity and expressiveness of string systems with tandem duplication, as defined below.

The model of a *string duplication system* consists of a *seed*, i.e., a starting string of finite length, a set of duplication rules that allow generating new strings from existing ones, and the set of all sequences that can be obtained by applying the duplication rules to the seed a finite number of times. The notion of *capacity*, introduced in [4] represents the average number of m-ary symbols per sequence symbol that are asymptotically required to encode a sequence in the string system, where m is the *alphabet* size (for DNA sequences the alphabet size is 4). The maximum value for capacity is 1. A duplication system is *fully expressive* if all strings with the alphabet appear as a substring of some string in the system. As we will show, if a system is not fully expressive, then its capacity is strictly less than 1.

Before presenting the notation, definitions, and the results more formally, in the rest of this section, we present two simple examples to illustrate the notions of expressiveness and capacity for tandem duplication string systems. Furthermore, we outline some useful tools as well as some of the results of the paper.

Example 1: Consider a string system on the binary alphabet $\Sigma = \{0, 1\}$ with 01 as the seed that allows tandem duplications of length up to 2. It is easy to check that the strings generated by this system start with 0 and end with 1. In fact, it can be

Manuscript received September 16, 2015; revised May 15, 2017; accepted July 7, 2017. Date of publication July 19, 2017; date of current version September 13, 2017. This work was supported by the NSF Expeditions in Computing Program–The Molecular Programming Project. This paper was presented at the 2015 IEEE International Symposium on Information Theory, Hong Kong.

S. Jain and J. Bruck are with the Electrical Engineering Department, California Institute of Technology, Pasadena, CA 91125 USA (e-mail: sidjain@caltech.edu; bruck@caltech.edu).

F. F. (Hassanzadeh) was with the Electrical Engineering Department, California Institute of Technology, Pasadena, CA 91125 USA. He is now with the Department of Electrical and Computer Engineering and the Department of Computer Science, University of Virginia, Charlottesville, VA 22903 USA (e-mail: farzad@virginia.edu).

Communicated by E. Tuncel, Associate Editor for Source Coding.

Digital Object Identifier 10.1109/TIT.2017.2728079

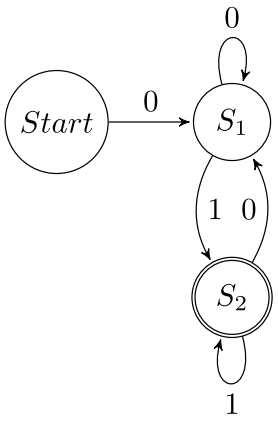

Fig. 1. The finite automaton for the systems $S = (\{0, 1\}, 01, \mathcal{T}^{tan}_{\leq k})$, where $k \geq 2$, including the system of Example 1. Notation used here is described in detail in Section II.

proved that all binary strings of length n which start with 0 and end with 1 can be generated by this system. The proof is based on the fact that every such string can be written as $0^{r_1}1^{r_2}\cdots 0^{r_{v-1}}1^{r_v}$, where each $r_i \geq 1$ and v is even. A natural way to generate this string is to duplicate 01 $\frac{v}{2}$ times and then duplicate the 0s and 1s as needed via duplications of length 1.

Expressiveness: From the preceding paragraph, every binary sequence s can be generated as a substring in this system as $0s1$. For example, although 11010 cannot be generated by this system, it can be generated as a substring of 0110101 in the following way:

$$01 \to 0101 \to 010101 \to 0\underline{110101}.$$

Hence this system is fully expressive.

Capacity: The number of length-n strings in this system is 2^{n-2}. Thus, encoding sequences of length n in this system requires $n-2$ bits. The capacity, or equivalently the asymptotic average number of bits (since the alphabet Σ is of size 2) per symbol, is thus equal to 1. This is not surprising as the system generates almost all binary sequences. □

Observing these facts for an alphabet of size 2, one can ask related questions on expressiveness and capacity for other alphabet sizes and duplication lengths. However, counting the number of length-n sequences for capacity calculation and characterizing fully expressive systems for larger alphabets are often not straightforward tasks. In this paper, we study these questions and develop methods to answer them.

A useful tool in this study is the theory of finite automata. As a simple example note that the string system over the binary alphabet {0, 1} in the preceding example can be represented by the finite automaton given in Figure 1. The regular expression for the language defined by the finite automaton is

$$R_{01} = (0^+1^+)^+, \tag{1}$$

which represents all binary strings that start with 0 and end with 1. For definitions of finite automata and regular expression, the reader can refer to Section II.

One can use the Perron-Frobenius theory [7], [12] to count the number of sequences which can be generated by a finite automaton. This enables us to use finite automata as a tool to calculate capacity for some string duplication systems with tandem repeats over larger alphabet.

TABLE I

CAPACITY VALUES TANDEM DUPLICATION STRING SYSTEMS $(\Sigma, s, \mathcal{T}^{tan}_{\leq k})$. HERE $x, y \in \Sigma^*$, AND $a, b, c \in \Sigma$ ARE DISTINCT

Σ	s	k	Capacity
$\{0, 1, 2\}$	012	3	$\simeq 0.876036$
arbitrary	$xabcy$	3	$\simeq 0.876036 \log_{\lvert\Sigma\rvert} 3$

In our results, we find that the exact capacity of the tandem duplication string system over the ternary alphabet {0, 1, 2} with seed 012 and duplication length at most 3 equals $\log_3 \frac{3+\sqrt{5}}{2} \simeq 0.876036$. Moreover, we generalize this result by characterizing the capacity of tandem duplication string systems over an arbitrary alphabet and a seed with maximum duplication length of 3. Namely, we show that if the maximum duplication length is 3 and the seed contains abc as a substring, where a, b, and c are distinct symbols, then the capacity $\simeq 0.876036 \log_{|\Sigma|} 3$. If such a substring does not exist in the seed, then the capacity is given by $\log_{|\Sigma|} 2$, unless the seed is of the form a^m, in which case the capacity is 0. Some of these results are highlighted in Table I.

Our next example presents a system that, unlike that of Example 1, is not fully expressive.

Example 2: Consider a tandem duplication string system over the ternary alphabet {0, 1, 2} with seed 012 and maximum duplication length 3. This system is not fully expressive as it cannot generate 210, 102, or 021, even as a substring. We provide a simple proof.

Proof: Let $z = \alpha\gamma\beta$, where α, γ and β are strings over {0, 1, 2} with $|\alpha|, |\beta| \geq 0$, and $1 \leq |\gamma| \leq 3$. Suppose z does not contain 210, 102 or 021 as a substring. We will now show that $z^* = \alpha\gamma\gamma\beta$ does not contain 210, 102 or 021 as a substring either. We have three cases:

- For $\gamma = a_1$, with $a_1 \in \{0, 1, 2\}$, the only possible new substrings generated in z^* which may not occur in z are the ones with suffix a_1a_1 or prefix a_1a_1.
- For $\gamma = a_1a_2$, with $a_1, a_2 \in \{0, 1, 2\}$, the only possible new substrings of length 3 generated in z^* are $a_1a_2a_1$ and $a_2a_1a_2$.
- For $\gamma = a_1a_2a_3$, with $a_1, a_2, a_3 \in \{0, 1, 2\}$, there is no substring of length 3 in z^* which does not occur in z.

Hence, if z does not contain 210, 102 or 021 as a substring, neither will z^*. Since the seed 012 does not contain 210, 102 or 021 as a substring, neither will any other string in the system. ■

Therefore, this tandem duplication string system is not fully expressive. □

Based on the previous example, one may ask what happens if we start with a seed that contains one of the strings 210, 102, or 021, e.g., if we let the seed be 01210? Does the system become fully expressive? While this system can generate all strings of length 3 as substrings, the answer is still no as shown in Theorem 2: Regardless of the seed, a ternary system with maximum duplication length of 3 is not fully expressive. We show in Theorem 4, that a maximum duplication length

TABLE II
EXPRESSIVENESS OF TANDEM DUPLICATION STRING SYSTEMS $\left(\Sigma, s, \mathcal{T}^{tan}_{\leq k}\right)$

Σ	s	k	fully expressive
$\{0,1,2\}$	arbitrary	≤ 3	No
$\{0,1,2\}$	012	≥ 4	Yes
Size ≥ 4	arbitrary	arbitrary	No

of at least 4 is needed to arrive at a fully expressive ternary system.

While for alphabets of size 2 or 3, increasing the maximum length on duplications turns a system that is not fully expressive into one that is, for alphabets of size 4 or more, duplication systems are not fully expressive regardless of how large the bound on duplication length is. The main tool in constructing quaternary strings that do not appear independently or as substrings in these systems is Thue's result proving the existence of ternary *squarefree* sequences of any length. A string is called squarefree if it does not have a tandem repeat of any length (Note that unary and binary squarefree sequences of arbitrarily large length do not exist.). The existence of such sequences underlies the significant shift in the behavior of tandem duplication systems with regards to expressiveness as a function of alphabet size. Some of our results on expressiveness are summarized in Table II.

As part of this paper, we also study regular languages for tandem duplication string systems. In [11], it was shown that the tandem duplication string system is not regular if the maximum duplication length is 4 or more when the seed contains 3 consecutive distinct symbols as a substring. However for maximum duplication length 3, this question remained open. In this paper, we show in Theorem 5 that if the maximum duplication length is 3, a tandem duplication string system is regular irrespective of the seed and the alphabet size. Moreover, we characterize the exact capacity for all these systems.

A. Related Work

Tandem duplications have already been studied in [2], [3], and [10]. However the main concern of these works is to determine the place of tandem duplication rules in the Chomsky hierarchy of formal languages. A study related to our work can be found in[4] and [11]. String systems with different duplication rules, namely end duplication, tandem duplication, reversed duplication and duplication with a gap, are defined and studied in [4]. In end duplication, a substring of certain length k is appended to the end of the previous string - for example, $A\underline{CT}GT \rightarrow ACTGT\underline{CT}$. In reversed tandem duplication, the reverse of a substring is appended in tandem in the previous string - for example, $A\underline{CT}GT \rightarrow ACTT\underline{C}GT$. In duplication with a gap, a substring is inserted after a certain gap g from its position in the original string, for example, $A\underline{CT}GT \rightarrow ACTG\underline{CT}T$.

For tandem duplication string systems, the authors in [4] show that for a fixed duplication length the capacity is 0. Further, they find a lower bound on the capacity of these systems, when duplications of all lengths are allowed. In this paper, we consider tandem duplication string systems, where we restrict the maximum size of the block being tandemly duplicated to a certain *finite* length.

In the rest of the paper, the term tandem duplication string system refers to string duplication systems with bounded duplication length.

The rest of the paper is organized as follows. In Section II, we present the preliminary definitions and notation. In Section III, we derive our main results on capacity and expressiveness. In Section IV, we show that if the maximum duplication length is 3, then the tandem duplication string system is regular irrespective of the seed and alphabet size. Further, using the regularity of the systems, we extend our capacity results. We present our concluding remarks in Section V.

II. PRELIMINARIES

Let Σ be some finite alphabet. An n-string $x = x_1x_2\cdots x_n \in \Sigma^n$ is a finite sequence where $x_i \in \Sigma$ and $|x| = n$. The set of all finite strings over the alphabet Σ is denoted by Σ^*. For two strings $x \in \Sigma^n$ and $y \in \Sigma^m$, their concatenation is denoted by $xy \in \Sigma^{n+m}$. For a positive integer m and a string s, s^m denotes the concatenation of m copies of s. A string $v \in \Sigma^*$ is a substring of x if $x = uvw$, where $u, w \in \Sigma^*$.

A string system $S \subseteq \Sigma^*$ is represented as a tuple $S = (\Sigma, s, \mathcal{T})$, where $s \in \Sigma^*$ is a finite string called seed, which is used to initiate the duplication process, and $\mathcal{T}$ is a set of rules that allow generating new strings from existing ones [4]. In other words, the string system $S = (\Sigma, s, \mathcal{T})$ contains all strings that can be generated from s using rules from $\mathcal{T}$ a finite number of times.

A tandem duplication map $T_{i,k}$,

$$T_{i,k}(x) = \begin{cases} uvvw, & x = uvw,\ |u| = i,\ |v| = k, \\ x, & \text{else}, \end{cases}$$

creates and inserts a copy of the substring of length k which starts at position $i+1$. We use $\mathcal{T}^{tan}_k : \Sigma^* \rightarrow \Sigma^*$ and $\mathcal{T}^{tan}_{\leq k}$ to denote the set of tandem duplications of length k, and tandem duplications of length at most k, respectively,

$$\mathcal{T}^{tan}_k = \left\{T_{i,k} : i \in \mathbb{N} \cup \{0\}\right\},$$
$$\mathcal{T}^{tan}_{\leq k} = \left\{T_{i,j} : i \in \mathbb{N} \cup \{0\}, j \in \mathbb{N},\ j \leq k\right\}.$$

With this notation, the system of Example 1 can be written as $(\{0,1\}, 01, \mathcal{T}^{tan}_{\leq 2})$.

The *capacity* of the string system $S = (\Sigma, s, \mathcal{T})$ is defined as

$$\text{cap}(S) = \limsup_{n\rightarrow\infty} \frac{\log_{|\Sigma|} |S \cap \Sigma^n|}{n}. \tag{2}$$

Furthermore, it is *fully expressive* if for each $y \in \Sigma^*$, there exists a $z \in S$, such that y is a substring of z.

A useful tool in calculating capacity of tandem duplication string systems is deterministic finite automaton (DFA) which consists of:

- A finite set of states Z.
- Alphabet Σ.

- Transition Rule $\delta : Z \times \Sigma \to Z$.
- Start state $z_o \in Z$.
- A set of accept states Y.

There also exist non-determinisitc finite automata. In this paper however, all the automata considered are deterministic. Henceforth, we will be using finite automaton to refer to a DFA. An example of a finite automaton is given in Figure 1. For this finite automaton we have,

- $Z = \{Start, S_1, S_2\}$.
- $\Sigma = \{0, 1\}$.
- $\delta(Start, 0) = S_1$, $\delta(S_1, 0) = S_1$, $\delta(S_1, 1) = S_2$, $\delta(S_2, 0) = S_1$, $\delta(S_2, 1) = S_2$.
- $z_o = Start$.
- $Y = \{S_2\}$.

The set of all possible strings that can be generated by a given DFA represent the language described by the finite automaton. This language L_R can be represented by a regular expression R. Formal definitions of regular expression can be found in [6]. For the purpose of this paper, we define:

- $R = s^*$: represents the language L_R which consists of all strings with 0 or more concatanated copies of $s \in \Sigma^*$, i.e., $L_R = \{s^m : m \geq 0\}$.
- $R = s^+$: represents the set of all strings with 1 or more concatanated copies of $s \in \Sigma^*$, i.e., $L_R = \{s^m : m \geq 1\}$.
- $R = R_1 R_2$: represents the language L_R formed by the concatenation of L_{R_1} and L_{R_2}, i.e. $L_R = \{s_1 s_2 : s_1 \in L_{R_1}, s_2 \in L_{R_2}\}$.

III. Capacity and Expressiveness

In this section, we present our results on the capacity and expressiveness of tandem duplication system with bounded duplication length. The section is divided into two parts; the first part focuses on capacity and the second on expressiveness.

A. Capacity

Our first result is on the capacity of a tandem duplication string system over ternary alphabet.

Theorem 1: For the tandem duplication string system $S = \left(\{0, 1, 2\}, 012, \mathcal{T}^{tan}_{\leq 3}\right)$, *we have*

$$\text{cap}(S) = \log_3 \frac{3 + \sqrt{5}}{2} \simeq 0.876036.$$

Proof: We prove this theorem by showing that the finite automaton given in Figure 2 accepts precisely the strings in S, and then finding the capacity using the Perron-Frobenius theory [7], [12].

The regular expression R for the language defined by this finite automaton is given by (see [6] for details on how to find a regular expression given a finite automaton)

$$R = (0^+1^+)^+2^+(1^+2^+)^* [0^+(2^+0^+)^*1^+(0^+1^+)^*2^+(1^+2^+)^*]^*. \quad (3)$$

Let L_R be the language defined by the regular expression R (and by the finite automaton). We first show that $L_R \subseteq S$. The direct way of doing so is to start with 012 and generate all the sequences in L_R via duplication. For simplicity of presentation, however, we take the reverse route: We show that every sequence in L_R can be transformed to 012 by a sequence *deduplications*. A deduplication of length k is an operation that replaces a substring $\alpha\alpha$ by α if $|\alpha| = k$. For two regular expressions R_1 and R_2, we use $R_1 \xrightarrow{dd_{\leq k}} R_2$ to denote that *each* sequence in L_{R_1} can be transformed into *some* sequence in L_{R_2} via a sequence of deduplications of length at most k.

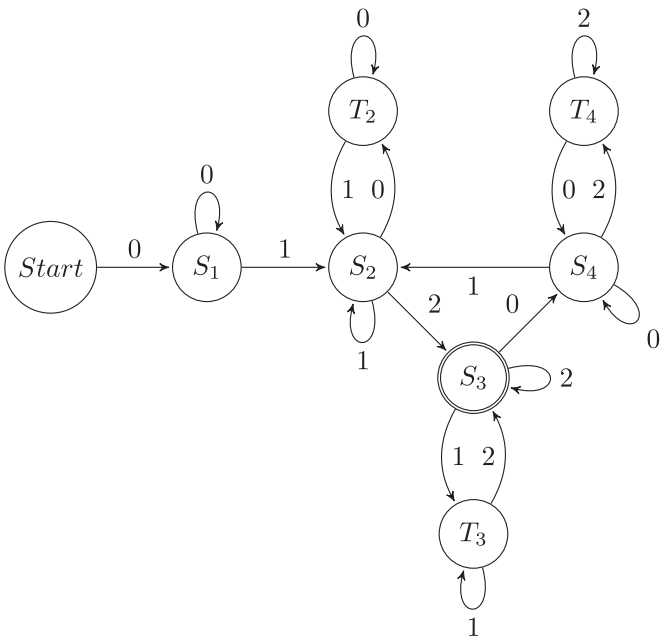

Fig. 2. Finite automaton for $S = (\{0, 1, 2\}, 012, \mathcal{T}^{tan}_{\leq 3})$.

Note that $R = B_1 B_2^*$, where

$$B_1 = (0^+1^+)^+2^+(1^+2^+)^*,$$
$$B_2 = 0^+(2^+0^+)^*1^+(0^+1^+)^*2^+(1^+2^+)^*.$$

We have $B_1 \xrightarrow{dd_{\leq 3}} 012\,(12)^* \xrightarrow{dd_{\leq 3}} 012$, since $a^+ \xrightarrow{dd_{\leq 3}} a$ and $(ab)^+ \xrightarrow{dd_{\leq 3}} ab$ for all $a, b \in \Sigma$. Furthermore,

$$B_2 \xrightarrow{dd_{\leq 3}} 0(20)^*1(01)^*2(12)^* \xrightarrow{dd_{\leq 3}} 0(20)^*1(01)^*2 \xrightarrow{dd_{\leq 3}} 0(20)^*12 \xrightarrow{dd_{\leq 3}} \{02012, 012\}. \quad (4)$$

Note for example that $1(01)^*\underline{2}(12)^* \xrightarrow{dd_{\leq 3}} 1(01)^*2$ as the underlined 2 is always preceded by a 1.

We thus have $R = B_1 B_2^* \xrightarrow{dd_{\leq 3}} \{01202012, 012012, 012\} \xrightarrow{dd_{\leq 3}} 012$, proving that $L_R \subseteq S$.

To complete the proof of $L_R = S$, we now show that $S \subseteq L_R$. In what follows, we say a finite automaton generates a sequence s, if there is a path with label s from $Start$ to an accepting state. If an automaton generates uvw, with $u, v, w \in \Sigma^*$, we may use v to refer both to the string v itself and to the part of the path that generates v. The meaning will be clear from the context.

We show $S \subseteq L_R$, by proving the following for the finite automaton in Figure 2:

i) It can generate 012.

ii) If the automaton can generate pqr, with $p, q, r \in \Sigma^*$ and $|q| \leq 3$, it can also generate pq^2r.

Condition i) holds trivially (see the path $Start - S_1 - S_2 - S_3$ in Figure 2). In order to prove ii), we define:

- *Path Label*: Given a path a in a finite automaton, the path label $l_a \in \Sigma^*$ is defined as the sequence obtained by concatenating the labels on the edges forming the path.
- *Path Length* is the number of edges of the path.
- *Duplicable Path*: Let q be a path that ends at state C. The path q is said to be *duplicable* if there exists path q' that **starts and ends at state** C such that the path labels of q and q' are the same.

Suppose a finite automaton can generate pqr. If q is duplicable, then pq^2r can also be generated by the finite automaton. As a result, to prove ii), it suffices to show that for each state C in Figure 2, all paths of length 1, 2 or 3 ending in C are duplicable.

Now, we show that all paths ending in $\{S_1, S_2, S_3, S_4, T_2, T_3, T_4, \}$ with length ≤ 3 are duplicable. Note that there are no nontrivial paths ending in the $Start$ state.

Given a state C and $j \in \{1, 2, 3\}$, let P_j^C be the set of all length-j paths ending in C and let Q_j^C be the set of all length-j paths starting and ending in C. If

$$\bigcup_{a \in P_j^C} l_a = \bigcup_{a \in Q_j^C} l_a, \tag{5}$$

then all length-j paths ending in C are duplicable.

We prove that (5) holds for all states and all $j \in \{1, 2, 3\}$. This is done by computing $\mathcal{A}$, $\mathcal{A}^2$ and $\mathcal{A}^3$, where $\mathcal{A}_1$ is the (labeled) adjacency matrix of the finite automaton given in Figure 2. Here in computing the matrix products, symbols do not commute, e.g. $xy \neq yx$. The adjacency matrix $\mathcal{A}$ and its square $\mathcal{A}^2$, where x, y and z represent edges labeled by 0, 1, and 2, respectively, and where rows and columns correspond in order to $S_1, S_2, S_3, S_4, T_2, T_3, T_4$, are given by

$$\mathcal{A} = \begin{bmatrix} x & y & 0 & 0 & 0 & 0 & 0 \\ 0 & y & z & 0 & x & 0 & 0 \\ 0 & 0 & z & x & 0 & y & 0 \\ 0 & y & 0 & x & 0 & 0 & z \\ 0 & y & 0 & 0 & x & 0 & 0 \\ 0 & 0 & z & 0 & 0 & y & 0 \\ 0 & 0 & 0 & x & 0 & 0 & z \end{bmatrix},$$

$$\mathcal{A}^2 = \begin{bmatrix} x^2 & y^2{+}xy & yz & 0 & yx & 0 & 0 \\ 0 & y^2{+}xy & z^2{+}yz & zx & x^2{+}yx & zy & 0 \\ 0 & xy & z^2{+}yz & x^2{+}zx & 0 & y^2{+}zy & xz \\ 0 & y^2{+}xy & yz & x^2{+}zx & yx & 0 & z^2{+}xz \\ 0 & y^2{+}xy & yz & 0 & x^2{+}yx & 0 & 0 \\ 0 & 0 & z^2{+}yz & zx & 0 & y^2{+}zy & 0 \\ 0 & xy & 0 & x^2{+}zx & 0 & 0 & z^2{+}xz \end{bmatrix}.$$

Each entry in these matrices lists the paths of specific length from the state identified by its row to the state identified by its column. For example, the entry $(6, 3)$ of $\mathcal{A}^2$, which equals $z^2 + yz$, indicates that there are two paths of length 2 from T_3 to S_3 with labels $z^2 = 22$ and $yz = 12$.

To check (5), we need to verify that the nonzero terms in the non-diagonal elements of each column also appear in its diagonal element. For $\mathcal{A}$ and $\mathcal{A}^2$, this can be easily done by observing the matrices. For example, the entry $(3, 3)$ of $\mathcal{A}^2$ equals $z^2 + yz$ and contains all terms appearing in column 3 of $\mathcal{A}^2$, which are yz and $z^2 + yz$. We verified using a computer that $\mathcal{A}^3$ also satisfies the same condition. Hence, we have shown that all paths of length at most 3 ending in $\{S_1, S_2, S_3, S_4, T_2, T_3, T_4\}$ are duplicable.

This completes the proof of $S \subseteq L_R$.

Now that we have shown $S = L_R$, we use the Perron-Frobenius theory [7], [12] to count the number of sequences which can be generated via the finite automaton in Figure 2. The accepting state S_3 is reachable from every other state in the finite automaton, therefore we can compute the capacity by calculating the maximum absolute eigenvalue e^* of the (unlabeled) adjacency matrix B of the strongly connected component of the finite automaton (i.e. the subgraph induced by $S_2, S_3, S_4, T_2, T_3, T_4$).

$$B = \begin{bmatrix} 1 & 1 & 0 & 1 & 0 & 0 \\ 0 & 1 & 1 & 0 & 1 & 0 \\ 1 & 0 & 1 & 0 & 0 & 1 \\ 1 & 0 & 0 & 1 & 0 & 0 \\ 0 & 1 & 0 & 0 & 1 & 0 \\ 0 & 0 & 1 & 0 & 0 & 1 \end{bmatrix}. \tag{6}$$

The maximum absolute eigenvalue of B is $e^* = \frac{3+\sqrt{5}}{2} \simeq 2.618034$. By the Perron-Frobenius Theory, $\text{cap}(S) = \log_3 e^* \simeq 0.876036$. ■

While the proof of the preceding theorem providing the exact capacity of the system under study is somewhat involved, it is easy to see why the capacity is strictly less than 1. One can observe from the regular expression for the finite automaton that it cannot generate a string that has 210, 021 or 102 as a substring, implying that the system is not fully expressive. As we will see in Lemma 4, such systems cannot have capacity 1. It is worth noting that the set of strings that avoid 210, 021, and 102 can be shown to have capacity $\simeq 0.914838$, which is slightly larger than the capacity of the system of the theorem.

B. Expressiveness

We now turn to study the expressiveness of tandem duplication systems with bounded duplication length. For completeness we start with binary systems, which is indeed the simplest case.

Lemma 3: The system $S = \left(\{0, 1\}, s, \mathcal{T}_{\leq 1}^{tan}\right)$, *for any* s *is not fully expressive.*

Proof: The system cannot generate $(01)^m$ as a substring of any string in S for $2m > |s|$. ■

As shown in Example 1, to obtain fully expressive binary systems, it suffices to increase the maximum duplication length to 2.

The next theorem is concerned with the expressiveness of $S = (\{0, 1, 2\}, s, \mathcal{T}_{\leq 3}^{tan})$. Larger alphabets and larger duplication lengths are considered in Theorems 3 and 4.

Theorem 2: Consider $S = (\{0, 1, 2\}, s, \mathcal{T}_{\leq 3}^{tan})$, *where* s *is any arbitrary starting string* $s \in \{0, 1, 2\}^*$. *Then,* S *is not fully expressive.*

Proof: A *k-irreducible* string is a string that does not have a tandem repeat $\alpha\alpha$, such that $|\alpha| \leq k$. For example, 01201, 01210, 02101, and 01210121 are 3-irreducible strings,

while 01212, 021021 and 01112 are not 3-irreducible. To prove the theorem, we identify certain properties in new 3-irreducible strings that may appear after a duplication and then construct a 3-irreducible string that is neither a substring of s, nor it satisfies the properties that every new 3-irreducible substring must satisfy.

Consider a duplication event that transforms a sequence $z = uvw$ to $z^* = uvvw$, where $|v| \leq 3$. Let x be a 3-irreducible string of length at least 4 that is present in z^* but not in z. The string x must intersect with both copies of v in z^* or else it is also present in z. Furthermore, it cannot contains vv, since otherwise it would not be 3-irreducible. To determine the properties of x, we consider three case: $|v| = 1, 2, 3$. In what follows assume $a_1, a_2, a_3 \in \Sigma$.

First, suppose $|v| = 1$, say $v = a_1$. In this case, a string x with the aforementioned properties does not exist as all new substrings contain the square a_1a_1.

Second, assume $|v| = 2$, say $v = a_1a_2$. Then $z^* = ua_1a_2a_1a_2w$ and x either ends with $a_1a_2a_1$ or starts with $a_2a_1a_2$.

Third, suppose $|v| = 3$, say $v = a_1a_2a_3$. So $z^* = ua_1a_2a_3a_1a_2a_3w$. Recall that $|x| \geq 4$. The string x either ends with $a_1a_2a_3a_1$ or $a_2a_3a_1a_2$, or starts with $a_2a_3a_1a_2$ or $a_3a_1a_2a_3$.

So for any new 3-irreducible substring $x = x_1 \cdots x_j$, $x_i \in \Sigma$, $j \geq 4$, we have $x_1 = x_3$, $x_1 = x_4$, $x_j = x_{j-2}$, or $x_j = x_{j-3}$. Now consider the string $(0121)^\ell 0$, where $\ell > |s|$. This sequences is 3-irreducible but does not satisfy any of the 4 properties stated for x. Since it is not a substring of s and it cannot be generated as a new substring, it is not a substring of any $y \in S$. ∎

Next we consider the system $\left(\Sigma, s, \mathcal{T}^{tan}_{\leq k}\right)$, $|\Sigma| \geq 4$ in Theorem 3. The proof of the theorem, uses the following lemma, which states that the expressiveness of a system also has a bearing on its capacity.

Lemma 4: If a string system S with alphabet Σ is not fully expressive, then $\text{cap}(S) < 1$.

Proof: Since S is not fully expressive, there exists a $z \in \Sigma^*$ that does not appear as a substring of any $y \in S$. Let $|z| = m$ and $\mu = n - m\lfloor \frac{n}{m} \rfloor$. We have

$$|S \cap \Sigma^n| \leq (|\Sigma|^m - 1)^{\lfloor \frac{n}{m} \rfloor} |\Sigma|^\mu.$$

Since m is finite, $\text{cap}(S) < 1$. ∎

Theorem 3: Consider $S = \left(\Sigma, s, \mathcal{T}^{tan}_{\leq k}\right)$, where $|\Sigma| \geq 4$, s is any arbitrary seed $\in \Sigma^*$ and k is some finite natural number. Then S is not fully expressive, which also implies $\text{cap}\ (S) < 1$.

Proof: Suppose $z = uvw \in S$, where $|v| \leq k$, and let $z^* = uvvw$ be the result of a duplication applied to z. Furthermore, suppose that $x = x_1 \cdots x_j$, where $x_i \in \Sigma$ and $j > k$, is a squarefree substring of z^* but not z. Similar to the proof of Theorem 2, x intersects both copies of v but does not contain both. As a result, either $x_1 = x_{1+i}$ or $x_j = x_{j-i}$, for some $2 \leq i \leq k$.

For definiteness assume Σ contains the symbols $\{0, 1, 2, 3\}$. The sequence $0t0$, where t is a squarefree sequence over the alphabet $\{1, 2, 3\}$ and $|t| > \max\{|s|, k\}$, is not a substring of s and cannot be generated as a substring since it does not satisfy the conditions stated for x above. Note that such a t exists since as shown by Thue [15], for an alphabet size ≥ 3, there exists a squarefree string of any length. Hence S is not fully expressive. The second part of the theorem follows from Lemma 4. ∎

Theorem 4: Consider $S = (\{0, 1, 2\}, 012, \mathcal{T}^{tan}_{\leq 4})$. Then S is a fully expressive string system.

Proof: Let $S' = \left(\{0, 1, 2\}, 012, \mathcal{T}^{tan}_{\leq 3}\right)$. Clearly, $S' \subseteq S$. From the proof of Theorem 1, we know that the automaton of Figure 2 gives the same language as S'. By checking this automaton, we find that all strings of lengths 1, 2, and 3, except 021, 210, and 102, appear as a substring of some string in S' and, as a result, some string in S. To generate 021, 210, and 102 as substrings of some string in S, we proceed as follows:

$$012 \to 0\underline{1212} \to \underline{01\mathbf{210}1212}$$
$$012 \to \underline{012012} \to 01\underline{202}012 \to 012\mathbf{021}202012$$
$$012 \to \underline{012012} \to 01\underline{202}012 \to 012020\underline{\mathbf{102}012}$$

where the repeats are underlined.

We have shown that all strings of length 3 appear in S as substrings. Now we show the same for every string $w = w_1w_2w_3w_4$ of length 4. To do so, we study 3 cases based on the structure of w:

I) First, suppose that w_4 is the same as w_1, w_2, or w_3. For generating such w as a substring, we first generate $w' = w_1w_2w_3$ as a substring of some string and then do a tandem duplication of w_3 if $w_4 = w_3$, of w_2w_3 if $w_4 = w_2$ and of $w_1w_2w_3$ if $w_4 = w_1$.

II) Suppose I) does not hold but $w_1 = w_2$ or $w_2 = w_3$. If the former holds, first generate $w_1w_3w_4$ and then duplicate w_1, and if the latter hold, generate $w_1w_2w_4$ and duplicate w_2.

III) If neither I) nor II) holds, then $w = 1210$, up to a relabling of the symbols. In this case, we first generate $w' = 0121$ and then do a tandem duplication of w' to get w. Note that w' is of type considered in I).

Until now, we have shown that all strings w of length at most 4 appear as a substring of some string in S. We use induction to complete the proof. Suppose all strings of length at most m appear as a substring of some string in S, where $m \geq 4$. We show that the same holds for strings of length $m + 1$.

Consider an arbitrary $w = a_1a_2 \cdots a_ma_{m+1}$. We now consider two cases:

i) If all three letters in the alphabet occur at least once in $a_{m-3}a_{m-2}a_{m-1}a_m$, then a_{m+1} equals a_{m-3}, a_{m-2}, a_{m-1}, or a_m, and w can be generated as a substring by a tandem duplication of some suffix of size ≤ 4 of $w' = a_1a_2 \cdots a_m$. Note that by the induction hypothesis w' can be generated as a substring of some string.

ii) If at least one letter in the alphabet does not occur in $a_{m-3}a_{m-2}a_{m-1}a_m$, then $a_{m-3}a_{m-2}a_{m-1}a_m$ is a sequence over binary alphabet and so it has a tandem repeat. Therefore w can be generated as a substring by tandem duplication. Hence, we have proved the Theorem. ∎

Table III summarizes the result of this subsection. It can be observed from the table that a change of behavior in

TABLE III
EXPRESSIVENESS OF TANDEM DUPLICATION STRING SYSTEMS $(\Sigma, s, \mathcal{T}^{tan}_{\leq k})$

Σ	s	k	fully expressive	Reason
$\{0\}$	0	≥ 1	Yes	Trivial
$\{0,1\}$	arbitrary	1	No	Lemma 3
$\{0,1\}$	01	≥ 2	Yes	Example 1
$\{0,1,2\}$	arbitrary	≤ 3	No	Theorem 2
$\{0,1,2\}$	012	≥ 4	Yes	Theorem 4
$\lvert\Sigma\rvert \geq 4$	arbitrary	arbitrary	No	Theorem 3

expressiveness occurs when the size of the alphabet increases to 4. If the size of the alphabet is 1, 2, or 3, for sufficiently large maximum duplication length, the systems are fully expressive. However, if the size of the alphabet is at least 4, then regardless of the maximum duplication length, the system is not fully expressive. This change is related to the fact that for alphabets of size 1 and 2, all squarefree strings are of finite length, but for alphabets of size 3 and larger, there are squarefree strings of any length. Specifically, in case ii) in the proof of Theorem 4, we used the fact that the binary string $a_{m-3}a_{m-2}a_{m-1}a_m$ has a tandem repeat. To adapt this proof for $|\Sigma| \geq 4$, we would need to show that the $(|\Sigma|-1)$-ary string $a_{m-3}a_{m-2}a_{m-1}a_m$ has a tandem repeat. But this is not in general true, since there are squarefree strings over alphabets of size at least 3 per Thue's result [15] and indeed we showed in Theorem 3, again using Thue's result, that the system $\left(\Sigma, s, \mathcal{T}^{tan}_{\leq k}\right)$ is not fully expressive for $|\Sigma| \geq 4$ and any k.

IV. REGULAR LANGUAGES FOR TANDEM DUPLICATION STRING SYSTEMS

Tandem duplication string systems that define regular languages are easier to study due to the fact that one can use tools from the Perron-Frobenius theory [7], [12] to calculate capacity. It was proved in [11] that for $|\Sigma| \geq 3$ and maximum duplication length ≥ 4, the language defined by tandem duplication string systems is not regular, if the seed contains abc as a substring such that a, b and c are distinct. However, if the maximum duplication length is 3, this question was left unanswered. In Theorem 5, we show that the language resulting from a tandem duplication system with the maximum duplication length of 3 is regular regardless of the alphabet size and seed. Further, in Corollary 5 we characterize the exact capacity of such tandem duplication string systems.

Theorem 5: Let $S = (\Sigma, s, \mathcal{T}^{tan}_{\leq 3})$, where Σ and s are arbitrary. The language defined by S is regular.

Proof: We first assume that $s = a_1 \cdots a_m$, where a_i are distinct. The case in which a_i are not distinct is handled later.

For $3 \leq j \leq m$, let

$$\begin{aligned} R_{a_1\cdots a_j} = {} & a_1^+ a_2^+ \left(a_1^+ a_2^+\right)^* a_3^+ \left(a_2^+ a_3^+\right)^* {B_{a_1a_2a_3}}^* \\ & a_4^+ \left(a_3^+ a_4^+\right)^* {B_{a_2a_3a_4}}^* \\ & \cdots \\ & a_i^+ \left(a_{i-1}^+ a_i^+\right)^* {B_{a_{i-2}a_{i-1}a_i}}^* \\ & \cdots \\ & a_j^+ \left(a_{j-1}^+ a_j^+\right)^* {B_{a_{j-2}a_{j-1}a_j}}^*, \end{aligned}$$

where, for $a, b, c \in \Sigma$,

$$B_{abc} = a^+(c^+a^+)^* b^+ (a^+b^+)^* c^+ (b^+c^+)^*.$$

We already know from Theorem 1 that $S = (\Sigma, s, \mathcal{T}^{tan}_{\leq 3})$ with $s = a_1 \cdots a_m$ is a regular language if $m = 3$. We show that for $m \geq 4$, S represents a regular language whose regular expression is given by $R_{a_1a_2\cdots a_m}$. Let L_R be the language defined by $R_{a_1a_2\cdots a_m}$. It suffices to show $L_R = S$.

We first show that $L_R \subseteq S$ by proving $R_{a_1a_2\cdots a_m} \xrightarrow{dd_{\leq 3}} s$. To do so, we show by induction that $R_{a_1a_2\cdots a_i} \xrightarrow{dd_{\leq 3}} a_1a_2\cdots a_i$. First note that this holds for $i = 3$, from the proof of Theorem 1. Assuming that it holds for i, to show that this also holds for $i+1$, where $i \geq 3$, we write

$$\begin{aligned} R_{a_1a_2\cdots a_{i+1}} = {} & R_{a_1a_2\cdots a_i} a_{i+1}^+ \left(a_i^+ a_{i+1}^+\right)^* {B_{a_{i-1}a_ia_{i+1}}}^* \\ & \xrightarrow{dd_{\leq 3}} a_1a_2\cdots a_ia_{i+1}\,(a_ia_{i+1})^*\,{B_{a_{i-1}a_ia_{i+1}}}^* \\ & \xrightarrow{dd_{\leq 3}} a_1a_2\cdots a_ia_{i+1}\,{B_{a_{i-1}a_ia_{i+1}}}^* \\ & \xrightarrow{dd_{\leq 3}} \{a_1a_2\cdots a_ia_{i+1}\,(a_{i-1}a_ia_{i+1})^*, \\ & \qquad a_1a_2\cdots a_ia_{i+1}\,(a_{i-1}a_{i+1}a_{i-1}a_ia_{i+1})^*\} \\ & \xrightarrow{dd_{\leq 3}} a_1a_2\cdots a_ia_{i+1}. \end{aligned}$$

Here we have used the fact that $cB_{abc} \xrightarrow{dd_{\leq 3}} cabc$ which follows from (4). Hence, $L_R \subseteq S$.

We now show that $S \subseteq L_R$. The finite automaton for L_R is given in Figure 3. Note that the seed s is in L_R. It thus suffices to show that if $x = pqr \in L_R$, then $y = pq^2r \in L_R$, where $p, q, r \in \Sigma^*$ and $|q| \leq 3$.

We prove this by showing that any length-1, 2 or 3 path ending in any state of the finite automaton in Figure 3 is duplicable, or in other words (5) holds for all the states in Figure 3. The finite automaton in Figure 3 is a generalization of the one in Figure 2. Note that in Figure 3, the states $\{S_1, S_2, S_3, S_4, T_2, T_3, T_4\}$ are exactly the same as those in Figure 2. More precisely, there is no additional path ending in these states in Figure 3. So, from the proof of Theorem 1, (5) holds for these states.

Now we show for the newly added states, i.e. $\{S_i, T_i : i \geq 5\}$, (5) holds. Consider a set $Q_k = \{S_{3k-1}, S_{3k}, S_{3k+1}, T_{3k-1}, T_{3k}, T_{3k+1}\}$ for some $k \geq 2$. The labelled adjacency matrix $\mathcal{A}$ for the subgraph induced by these states is given by

$$\mathcal{A} = \begin{bmatrix} y & z & 0 & x & 0 & 0 \\ 0 & z & x & 0 & y & 0 \\ y & 0 & x & 0 & 0 & z \\ y & 0 & 0 & x & 0 & 0 \\ 0 & z & 0 & 0 & y & 0 \\ 0 & 0 & x & 0 & 0 & z \end{bmatrix},$$

where x is used as a label for a_k, y for a_{k+1} and z for a_{k+2}. $\mathcal{A}^2$ is given by

$$\mathcal{A}^2 = \begin{bmatrix} y^2{+}xy & z^2{+}yz & zx & x^2{+}yx & zy & 0 \\ xy & z^2{+}yz & x^2{+}zx & 0 & y^2{+}zy & xz \\ y^2{+}xy & yz & x^2{+}zx & yx & 0 & z^2{+}xz \\ y^2{+}xy & yz & 0 & x^2{+}yx & 0 & 0 \\ 0 & z^2{+}yz & zx & 0 & y^2{+}zy & 0 \\ xy & 0 & x^2{+}zx & 0 & 0 & z^2{+}xz \end{bmatrix},$$

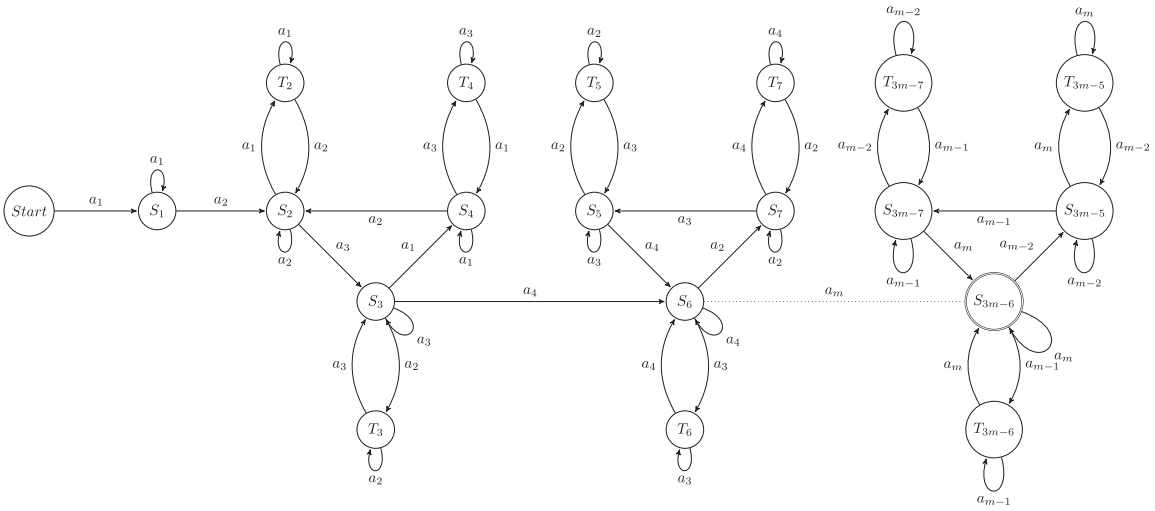

Fig. 3. Finite automaton for $S = (\Sigma, a_1a_2a_3\cdots a_m, \mathcal{T}^{tan}_{\leq 3})$.

The non-zero terms in the non diagonal entries of each column also appear in the diagonal entry of that column for $\mathcal{A}$ and $\mathcal{A}^2$. This can also be verified for $\mathcal{A}^3$. Hence, we have shown that any length-1, 2, 3 path starting in some state $C \in Q_k$ and ending in some state $D \in Q_k$ is duplicable for all $k \geq 2$.

We also need to show that any length-1, 2, 3 path that ends in Q_k but starts in a state that is not in Q_k is duplicable. Note that the states in Q_k are not reachable from states $\in Q_{k'}$ with $k' > k$, any possible path of length-1, 2 or 3 ending in some state $D \in Q_k$ and starting in a state $C \in Q_{k'}$ with $k' < k$, has to pass through state S_{3k-3}. Now, we enumerate the labels of all length-1, 2 and 3 paths ending in some state in Q_k but starting in some state $\in Q_{k'}$ with $k' < k$.

- Label of a path of length 1: a_{k+2} (ends in S_{3k}).
- Label of a path of length 2: i) ending in S_{3k}: $a_{k+1}a_{k+2}$, $a_{k+2}a_{k+2}$, ii) ending in S_{3k+1}: $a_{k+2}a_k$, iii) ending in T_{3k}: $a_{k+2}a_{k+1}$.
- Label of a path of length 3:
  - i) ending in S_{3k-1}: $a_{k+2}a_ka_{k+1}$,
  - ii) ending in S_{3k}: $a_{k+1}a_{k+1}a_{k+2}$, $a_ka_{k+1}a_{k+2}$, $a_{k+1}a_{k+2}a_{k+2}$, $a_{k+2}a_{k+2}a_{k+2}$, $a_{k+2}a_{k+1}a_{k+2}$,
  - iii) ending in S_{3k+1}:$a_{k+1}a_{k+2}a_k$, $a_{k+2}a_ka_k$, $a_{k+2}a_{k+2}a_k$,
  - iv) ending in T_{3k}: $a_{k+1}a_{k+2}a_{k+1}$, $a_{k+2}a_{k+1}a_{k+1}$, $a_{k+2}a_{k+2}a_{k+1}$,
  - v) ending in T_{3k+1}: $a_{k+2}a_ka_{k+2}$.

All the path labels enumerated above are duplicable which can be verified by inspecting $\mathcal{A}$, $\mathcal{A}^2$, $\mathcal{A}^3$, for paths of length 1, 2 and 3 respectively. This completes the proof of $S \subseteq L_R$.

We have proved the statement of Theorem 5 assuming all a_i's in the seed s to be distinct. Now assume the symbols of s are not distinct. We color the symbols of s so that they become distinct and obtain the system $\tilde{S} = \left(\tilde{\Sigma}, \tilde{s}, \mathcal{T}^{tan}_{\leq 3}\right)$. Applying the preceding proof for distinct symbols to $\tilde{S}$, we find that $\tilde{S}$ is regular. Let $h : \tilde{\Sigma} \to \Sigma$ be a mapping that removes the colors. This mapping is called a morphism. By [14], we have that $S = h(\tilde{S})$ is also regular. ■

An immediate corollary on the capacity of tandem duplication string system considered in Theorem 5 is stated next.

Corollary 5: If for S in Theorem 5, s contains abc as a substring such that a, b, and $c \in \Sigma$ are distinct, then $\text{cap}(S) = \log_{|\Sigma|} \frac{3+\sqrt{5}}{2} \simeq 0.876036 \log_{|\Sigma|} 3$. *Otherwise, except for the seed s of the form a^m,* $\text{cap}(S) = \log_{|\Sigma|} 2$. *If $s = a^m$,* $\text{cap}(S) = 0$.

Proof: If abc occurs as a substring of the seed s such that a, b and $c \in \Sigma$ are distinct, then the adjacency matrix of the finite automaton for B_{abc} (strongly connected component of the finite automaton for $R_{a_1a_2\cdots a_m}$) has the maximum eigenvalue. Therefore, the $\text{cap}(S) = \log_{|\Sigma|} \frac{3+\sqrt{5}}{2} \simeq 0.876036 \log_{|\Sigma|} 3$ (see (6) in the proof of Theorem 1).

If no 3 consecutive symbols in the seed s are all distinct and $s \neq a^m$, then the maximum capacity component is a finite automaton only over 2 distinct symbols as in Figure 1. In other words, terms of the form $(a_i^+ a_{i+1}^+)^*$ determine the capacity. Hence the capacity is $\log_{|\Sigma|} 2$.

When seed $s = a^m$, there is exactly one sequence of any given length in the system. Hence $\text{cap}(S) = 0$. ■

The following examples illustrate the statement of Theorem 5 and an application of its proof method.

Example 6: The string system $S = (\{0, 1, 2, 3\}, 0123, \mathcal{T}^{tan}_{\leq 3})$ is regular by Theorem 5 and the regular expression is given by

$$R_{0123} = 0^+1^+(0^+1^+)^*2^+(1^+2^+)^*B_{012}{}^*3^+(2^+3^+)^*B_{123}{}^*.$$

By Corollary 5, the capacity of this system $\simeq 0.876036 \log_4 3 \simeq 0.694242$. □

Example 7: The string system $S = (\{0, 1, 2\}, 0112, \mathcal{T}^{tan}_{\leq 3})$ is regular by Theorem 5, and the regular expression is given by

$$R_{0112} = 0^+1^+(0^+1^+)^*1^+(1^+1^+)^*B_{011}{}^*2^+(1^+2^+)^*B_{112}{}^*.$$

By Corollary 5, the capacity of this system is given by $\log_3 2 \simeq 0.63093$. □

When a_i's are assumed to be distinct it can be verified from the regular expression $R_{a_1\cdots a_j}$ in the proof of Thereom 5 that

TABLE IV

CAPACITY AND EXPRESSIVENESS FOR DIFFERENT TANDEM DUPLICATION STRING SYSTEMS ($\Sigma, s, \mathcal{T}^{tan}_{\leq k}$)

Σ	s	k	Capacity	Fully Expressive
$\{0\}$	0^m for some $m \geq 1$	1	1	Yes
$\{0,1\}$	01	1	0	No
$\{0,1\}$	arbitrary but not a^m for some $a \in \{0,1\}$	≥ 2	1	Yes
$\lvert\Sigma\rvert \geq 3$	arbitrary but not a^m for some $a \in \Sigma$	2	$\log_{\lvert\Sigma\rvert} 2$	No
$\lvert\Sigma\rvert \geq 3$	$xabcy$ (x and $y \in \Sigma^*$, a, b and $c \in \Sigma$ and $a \neq b \neq c \neq a$)	3	$\log_{\lvert\Sigma\rvert} \frac{3+\sqrt{5}}{2}$	No
$\lvert\Sigma\rvert \geq 3$	No 3 consecutive symbols in the seed are all distinct and $s \neq a^m$ for $a \in \Sigma$	3	$\log_{\lvert\Sigma\rvert} 2$	No
$\{0,1,2\}$	012	≥ 4	?	Yes
$\lvert\Sigma\rvert \geq 4$	arbitrary	≥ 4	?	No

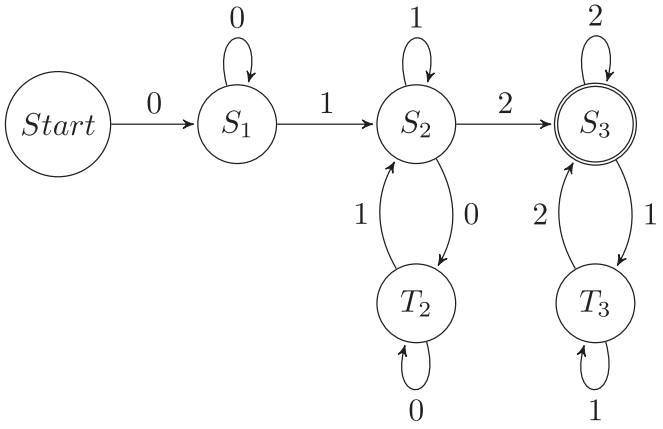

Fig. 4. Finite automaton for $S = (\{0,1,2\}, 012, \mathcal{T}^{tan}_{\leq 2})$. The regular expression $R = 0^+1^+(0^+1^+)^*2^+(1^+2^+)^*$.

the last occurence of a_i is before the first occurence of a_{i+3} for any $i = 1, 2, \cdots, j-3$ for all $z \in S$.

The following corollary follows for maximum duplication length 2 using the same idea as in Theorem 5

Corollary 8: The capacity for $S = (\Sigma, a_1a_2\cdots a_m, \mathcal{T}^{tan}_{\leq 2})$ *is given by* $\log_{|\Sigma|} 2$*, except for the case in which seed* $s = a^m$ *for* $a \in \Sigma$*. In that case, the capacity is* 0*.*

Proof: The string system $S = (\Sigma, a_1a_2\cdots a_m, \mathcal{T}^{tan}_{\leq 2})$ is regular. This can be proved using the same method as used in the proof of Theorem 5. The regular expression $Q_{a_1a_2\cdots a_m}$ for $m \geq 2$ is given by

$$Q_{a_1a_2\cdots a_m} = a_1^+a_2^+(a_1^+a_2^+)^*a_3^+(a_2^+a_3^+)^*\cdots a_m^+(a_{m-1}^+a_m^+)^*.$$

As in Proof of Corollary 5, the capacity is determined by term(s) of the form $(a_i^+a_{i+1}^+)^*$, except for the case when seed $s = a^m$. Therefore, the capacity for language represented by $Q_{a_1a_2\cdots a_m}$ is $\log_{|\Sigma|} 2$, when $s \neq a^m$ and 0 when $s = a^m$. ■

The finite automaton for a special case of Corollary 8 with $|\Sigma| = 3$ is given in Figure 4.

Table IV lists the capacity and expressiveness results presented in this paper and also the open question on capacity when $k \geq 4$. The expressiveness results follows from Table III.

V. CONCLUSION

In this paper, we showed that for tandem duplication string systems with bounded duplication length if the maximum duplication length is 3 or less, the language described by the string system is regular. Further, we computed exact capacities for these systems. Computing the capacities for bounded tandem duplication string systems with maximum duplication length greater than 3 remains an open problem.

Using Thue's result [15], we showed that a tandem duplication string system cannot be fully expressive if the alphabet size is ≥ 4. However, for an alphabet of size 3 or less such systems can be fully expressive. Therefore, we have completely characterized fully expressive and non-fully expressive tandem duplication string systems with bounded duplication lengths. As future work, we would like to generalize the notion of expressiveness by counting the asymptotic number of *substrings* of length n that a string system can generate. Mathematically, we may define the expressiveness $Exp(S)$ of a string system S as

$$Exp(S) = \limsup_{n\to\infty} \frac{\log_{|\Sigma|} E_n(S)}{n}.$$

Here $E_n(S)$ represents the number of substrings of length n that can be generated by S. It is notable here that with this definition of expressiveness, a fully expressive string system S has $Exp(S) = 1$.

In this paper, we studied questions related to the generation of a diverse set of sequences from a seed given a tandem duplication rule. One can also study the minimum number of steps required to generate a given sequence of length n from a squarefree seed and therefore define the notion of distance between a sequence and its seed given a tandem duplication rule. For the special case of binary sequences, we have studied this distance in [1].

It is notable here that the same sequence can be deduplicated to (or equivalently, generated from) more than one squarefree seed given a tandem duplication rule. For example: the sequence 012101212 can be deduplicated to 012 as well as 0121012 under bounded tandem duplication with maximum duplication length 4 in the following way

$$\underline{012101212} \xrightarrow{dd_{\leq 4}} 0\underline{1212} \xrightarrow{dd_{\leq 4}} 012.$$
$$0121\underline{01212} \xrightarrow{dd_{\leq 4}} 0121012.$$

Here the underlined portion represents the repeat that is being deduplicated in a given step. This raises the question of the uniqueness of squarefree seeds for strings generated by a given tandem duplication rule. We have studied this question in [8] in the context of duplication errors.

ACKNOWLEDGEMENTS

The authors would like to thank the anonymous reviewers for the thorough reviews and valuable comments. The reviews helped us make the finite automata and proofs presented in this paper simpler and more clear.

REFERENCES

[1] N. Alon, J. Bruck, F. Farnoud, and S. Jain, "On the duplication distance of binary strings," in *Proc. IEEE Int. Symp. Inf. Theory (ISIT)*, Barcelona, Spain, Jul. 2016, pp. 260–264.

[2] J. Dassow, V. Mitrana, and G. Păun, "On the regularity of duplication closure," *Bull. Eur. Assoc. Theor. Comput. Sci.*, vol. 69, pp. 133–136, Oct. 1999.

[3] J. Dassow, V. Mitrana, and A. Salomaa, "Operations and language generating devices suggested by the genome evolution," *Theor. Comput. Sci.*, vol. 270, nos. 1–2, pp. 701–738, 2002.

[4] F. F. Hassanzadeh, M. Schwartz, and J. Bruck, "The capacity of string-duplication systems," *IEEE Trans. Inf. Theory*, vol. 62, no. 2, pp. 811–824, Feb. 2016.

[5] J. W. Fondon and H. R. Garner, "Molecular origins of rapid and continuous morphological evolution," *Proc. Nat. Acad. Sci. USA*, vol. 101, no. 52, pp. 18058–18063, 2004.

[6] J. E. Hopcroft, R. Motwani, and J. D. Ullman, *Introduction to Automata Theory, Languages, and Computation*. Boston, MA, USA: Addison-Wesley, 2001.

[7] K. A. S. Immink, *Codes for Mass Data Storage Systems*. Denver, CO, USA: Shannon Foundation, 2004.

[8] S. Jain, F. Farnoud, M. Schwartz, and J. Bruck, "Duplication-correcting codes for data storage in the DNA of living organisms," in *Proc. IEEE Int. Symp. Inf. Theory (ISIT)*, Barcelona, Spain, Jul. 2016, pp. 1028–1032.

[9] E. S. Lander *et al.*, "Initial sequencing and analysis of the human genome," *Nature*, vol. 409, no. 6822, pp. 860–921, 2001.

[10] P. Leupold, C. Martín-Vide, and V. Mitrana, "Uniformly bounded duplication languages," *Discrete Appl. Math.*, vol. 146, no. 3, pp. 301–310, 2005.

[11] P. Leupold, V. Mitrana, and J. M. Sempere, "Formal languages arising from gene repeated duplication," in *Aspects of Molecular Computing*. Berlin, Germany: Springer, 2003, pp. 297–308.

[12] D. Lind and B. Marcus, *An Introduction to Symbolic Dynamics and Coding*. Cambridge, U.K.: Cambridge Univ. Press, 1985.

[13] N. Mundy and A. J. Helbig, "Origin and evolution of tandem repeats in the mitochondrial DNA control region of shrikes (*Lanius* spp.)," *J. Mol. Evol.*, vol. 59, no. 2, pp. 250–257, 2004.

[14] J. Shallit, *A Second Course in Formal Languages and Automata Theory*. Cambridge, U.K.: Cambridge Univ. Press, 2008.

[15] A. Thue, "Über unendliche Zeichenreihen," in *Skrifter Udgivne af Videnskabsselskabet i Christiania. I, Mathematisk-Naturvidenskabelig Klasse*, vol. 7. Cristiana, 1906.

[16] K. Usdin, "The biological effects of simple tandem repeats: Lessons from the repeat expansion diseases," *Genome Res.*, vol. 18, no. 7, pp. 1011–1019, 2008.

Siddharth Jain (S'15) is a PhD Candidate in the department of Electrical Engineering at Caltech. His research interests include information and coding theory, machine learning, information theoretic and statistical analysis of genomic data, pattern recognition, data compression and computational biology. Siddharth received Bachelors and Masters degree from Indian Institute of Technology (IIT) Kanpur, India in 2013. He was awarded the proficiency medal at IIT Kanpur for excellent academic performance in Electrical Engineering.

Farzad Farnoud (Hassanzadeh) (M'13) is an Assistant Professor in the Department of Electrical and Computer Engineering and the Computer Science Department at the University of Virginia. Previously, he was a postdoctoral scholar at the California Institute of Technology. He received his MS degree in Electrical and Computer Engineering from the University of Toronto in 2008. From the University of Illinois at Urbana- Champaign, he received his MS degree in mathematics and his Ph.D. in Electrical and Computer Engineering in 2012 and 2013, respectively. His research interests include the information-theoretic and probabilistic analysis of genomic evolutionary processes; rank aggregation and gene prioritization; and coding for flash memory and DNA storage. Dr. Farnoud is the recipient of the 2013 Robert T. Chien Memorial Award from the University of Illinois for demonstrating excellence in research in electrical engineering and the recipient of the 2014 IEEE Data Storage Best Student Paper Award.

Jehoshua Bruck (S'86–M'89–SM'93–F'01) is the Gordon and Betty Moore Professor of computation and neural systems and electrical engineering at the California Institute of Technology (Caltech). His current research interests include information theory and systems and the theory of computation in nature. Dr. Bruck received the B.Sc. and M.Sc. degrees in electrical engineering from the Technion-Israel Institute of Technology, in 1982 and 1985, respectively, and the Ph.D. degree in electrical engineering from Stanford University, in 1989. His industrial and entrepreneurial experiences include working with IBM Research where he participated in the design and implementation of the first IBM parallel computer; cofounding and serving as Chairman of Rainfinity (acquired in 2005 by EMC), a spin-off company from Caltech that created the first virtualization solution for Network Attached Storage; as well as cofounding and serving as Chairman of XtremIO (acquired in 2012 by EMC), a start-up company that created the first scalable all-flash enterprise storage system. Dr. Bruck is a recipient of the Feynman Prize for Excellence in Teaching, the Sloan Research Fellowship, the National Science Foundation Young Investigator Award, the IBM Outstanding Innovation Award and the IBM Outstanding Technical Achievement Award.