Improved Signal Processing Methods for Velocity Selective Neural Recording Using Multi-Electrode Cuffs

Assad Al-Shueli, Chris Clarke, Nick Donaldson and John Taylor

Abstract—This paper describes an improved system for obtaining velocity spectral information from electromyogram (ENG) recordings using multi-electrode cuffs (MECs). The starting point for this study is some recently published work that considers the limitations of conventional linear signal processing methods (‘delay-and-add’) with and without additive noise. By contrast to earlier linear methods, the present paper adopts a fundamentally non-linear velocity classification approach based on a type of artificial neural network (ANN). The new method provides a unified approach to the solution of the two main problems of the earlier delay-and-add technique, i.e. a damaging decline in both velocity selectivity and velocity resolution at high velocities. The new method can operate in real-time, is shown to be robust in the presence of noise and also to be relatively insensitive to the form of the action potential waveforms being classified.

Index Terms—Biomedical signal processing, Biomedical transducers, Microelectronic implants, Neural prosthesis, Artificial neural networks

I. INTRODUCTION

One component of neuroprosthetic systems is the neural input through which information is transferred from a physiological environment into the prosthesis. Tripolar recording from peripheral nerves cannot generally extract much of the information in the neural traffic, so we and others have been investigating velocity selective recording (VSR) since it allows the possibility of increasing the information obtainable from peripheral nerves (electromyogram-ENG) by carrying out a spectral analysis in the velocity domain [1]-[7]. The resulting spectrum shows not only the direction of action potential (AP) propagation (afferent or efferent) but also provides a measure of the differential levels of excitation of the fibre populations in the nerve. Since there is a well-established relationship between AP propagation velocity and fibre diameter that is approximately linear for myelinated fibres, VSR provides a method for assessing the level of activity in nerve fibre populations of different diameter as well as establishing the direction of propagation. This information potentially allows more information to be extracted from an intact nerve for use in applications requiring sensory feedback such as neuroprostheses [8]-[11].

In principle the velocity of a propagating AP can be calculated by timing the appearance of the signal at two or more points along the nerve and then dividing the distance between the points by the delay. Various researchers have investigated practical adaptations of this simple idea in the past (e.g. [12]-[14]). These methods were made both possible and practical by the introduction of electrode cuffs in 1974-75 [15] that provided for the first time a stable, chronically implantable interface for neural stimulation and recording. However, each cuff contained typically only 2-3 electrodes limiting the available velocity selectivity. The invention of the multi-electrode cuff (MEC) [16] has been very influential in improving this situation, enabling recordings to be made with increased velocity selectivity (each MEC contains typically about 10-15 electrodes). Using this approach it should be possible to access slower (i.e. smaller diameter) fibres than was the case with the earlier, simpler cuffs whilst at the same time offering the potential for incorporation in compact, integrated implantable systems. In addition, considerable progress has been made in the design of implantable integrated signal processing systems featuring low-noise, low-power amplifiers intended to be mounted directly on the MEC for optimum signal-to-noise performance [2],[3].

The method currently in use to estimate the velocity spectrum of ENG data captured by an MEC-based system is based on linear signal processing principles and is referred to

<table>
<thead>
<tr>
<th>Acronym</th>
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<tr>
<td>ADC</td>
<td>Analogue to Digital Converter</td>
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<td>ANN</td>
<td>Artificial Neural Network</td>
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<td>AP</td>
<td>Action Potential</td>
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<td>FPGA</td>
<td>Field Programmable Gate Array</td>
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<td>IVS</td>
<td>Intrinsic Velocity Spectrum</td>
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<td>MEC</td>
<td>Multi-Electrode Cuff</td>
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<td>SNR</td>
<td>Signal to Noise Ratio</td>
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<td>SPU</td>
<td>Signal Processing Unit</td>
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<td>TMAP</td>
<td>Trans-Membrane Action Potential</td>
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<td>VSR</td>
<td>Velocity Selective Recording</td>
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TABLE I
A LIST OF ACRONYMS USED
In this paper we propose a nonlinear approach based upon a type of artificial neural network (ANN) to tackle both the difficulties mentioned above without increasing either the length of the MEC or the sampling rate. It is shown that good results can be achieved with a very simple optimized nonlinear system (although we retain the term ANN due to the training methods used in the optimization). In addition, the new method achieves significantly higher velocity selectivity than any other currently available method and since it requires supervised learning, once trained it runs in real-time. Although ANNs have been employed in related applications (e.g. AP classification [17]) this is the first time they have been proposed as a method to discriminate between the velocities of propagating ENG. The approach is somewhat analogous to that employed in speech recognition and hence the chosen type of ANN, a time-delay neural network, was first employed in that application [18]. This paper consists of a study based on ENG data generated using a specially designed synthesised source [19] that permits the addition of controlled amounts of white Gaussian noise. The behaviour of the ANN-based velocity classifier is examined as the noise level is increased and the form of the AP input is varied from the pattern used for training. The results are compared to the performance of a ‘conventional’ delay-and-add system. Finally, the hardware cost of an ANN-based classifier relative to an equivalent delay-and-add system is assessed.

II. METHODS

A. System Description

1) Conventional VSR signal processing: ‘delay-and-add’

The overall plan of the complete system is shown in Fig 1 and consists of three separate stages. The first stage on the left (block 1) is an FPGA-based programmable AP and noise signal generator (synthesiser) whose output is eleven unipolar signals delayed so as to simulate an AP moving through the cuff [19]. The signals are generated using a modified pulse-width modulation output direct from the FPGA. In this study we consider an MEC of length 3 cm and propagation velocities in the range 10 – 100 m/s although operation at lower velocity is also possible. To these signals controlled amounts of uncorrelated additive white Gaussian noise were added [3]. Blocks 2 and 3 are the two principal sections of a neural recording system intended for ultimate implantation as a pair of remotely powered modules connected together using a multicores implantable cable [7]. Block 2 is referred to as the electrode unit and is intended to be mounted on the MEC for optimum performance. It consists of a set of specially-designed low noise differential amplifiers (nominal gain 10,000) whose 10 bipolar (or single differential) outputs are digitized (10 bits resolution) and multiplexed into one channel for transmission along the implantable cable [6]. In a distributed sensor architecture, several electrode units are placed at different sites in the peripheral nervous system and connected by implanted cables to a single monitoring unit [7]. The monitoring unit is a demultiplexer (DEMUX) and digital signal processing system that interfaces with an RF telemetry system (not described in this paper). In addition to processing and transmitting the recorded data for external logging, the monitoring unit processes the system commands and provides power supplies.
Although a single monitoring unit can process the data from and control and power several electrode units [7], in the experiments described in this paper only one of each type of unit is used as shown in Fig 1. In addition, the electrode unit employed is a complete prototype custom integrated circuit whereas the monitoring unit is realised in FPGA. Although for ultimate implantation the monitoring unit will be a second integrated circuit, the comparison of different VSR signal processing strategies that lies at the heart of this investigation requires easy flexibility of design suggesting the use of FPGA. The signal processing circuitry is contained in the unit on the far right of block 3 in Fig 1 (SPU) and is preceded by stages of de-multiplexing into 10, parallel bipolar channels and conversion into 9 tripolar (or double differential) channels. Tripolar signals are generally employed in ENG recording applications since they have enhanced immunity to common mode signal contamination (such as electromyogram) compared to monopolar or bipolar ones [20]. Finally the captured and recorded signals are processed in MATLAB. As already noted, the fundamental comparison to be made is between the conventional delay-and-add method [1] and the proposed new method, which employs an ANN as the core of the SPU.

2) Action potential (AP) and noise generation

The AP generator was described in [19] and so only a summary will be given here. The input to the MEC is a transmembrane action potential function (TMAP), \( V_m(t) \) and the resulting single fibre action potential is a propagating wave with the time dependence of the underlying TMAP function and the spatial properties of the MEC [2]. The TMAP can therefore be regarded as the driving potential resulting in a propagating electric field. APs appear at electrodes placed in this field. We represent the TMAP function by the following time-varying function [1]:

\[
V_m(t) = A t e^{-Bt}
\]

where \( A \), \( B \) and \( n \) are constants. These constants are chosen to generate APs with typical mammalian shape, the duration of the resulting signals being on the order of 1 ms. In this paper we use one of the TMAP functions that formed the basis of the analysis in [2]. The parameter values in eqn (1) are: \( A = 4.08 \times 10^{-4}, B = 1.5 \times 10^{-4} \) and \( n = 1 \). In section III C a second function (TMAP2) is introduced as an example of other parameter sets that can also be used. The signal TMAP2 is used to investigate the effect of varying the TMAP on the performance of the system. In this case the parameters are: \( A = 7.44 \times 10^{-11}, B = 1.0 \times 10^{-6} \) and \( n = 3 \). The form of both of these functions is shown in Fig 2.

The AP signal generation system is implemented as an FPGA controller plus a few passive components. It employs pulse-width modulation and related methods to achieve a very smooth representation of \( V_m(t) \) without the need for excessively high sample rates and/or high resolution digital components[19]. For the purposes of this study, the system is modified to allow variable amounts of noise to be added to each channel. Linear feedback shift registers with different tap positions were added to provide the necessary uncorrelated noise sources, the noise level being controlled by an 8-bit counter and a comparator. The controller has a USB interface to a desktop PC that allows easy programmability of all the main parameters. A simplified block diagram of the AP generator is given in Fig 3 showing the modifications necessary for the inclusion of noise.

B. VSR Signal Processing Methods

1) Conventional VSR signal processing: 'delay-and-add'

Fig 4 is the block diagram of the signal-processing unit
(SPU) in a conventional VSR system. One complete set of fixed delays is required for each AP propagation velocity to be classified [2]. In an implanted system the SPU will be connected to the mixed analogue/digital signal acquisition integrated circuits by means of implanted cables and so, as already noted, the multiplexed input is first de-multiplexed into 10 parallel (dipole) channels. After the signals are converted to tripolar form, the next steps are to add programmable delays to counteract the naturally occurring delays and then sum the outputs. In the presence of an excited population with propagation velocity \( v \), as the delays are stepped through the appropriate range, the output passes through a maximum when the delay \( T \) is given by:

\[
T = \frac{d}{v}
\]  

where \( d \) is the inter-electrode spacing. There are two fundamental problems with this approach, however. Firstly, for a practical cuff length (i.e. about 2-3 cm) the available velocity selectivity is generally too low to be useful. This issue can be mitigated to some extent by placing bandpass filters at the output [2]. Secondly, the velocity resolution, that is to say the number of available points on the velocity axis, declines at high velocities making the velocity spectral content difficult to measure. This is because the inter-channel delays \( T \) generated by a typical MEC are small at high velocities which limits the number of possible velocity steps. Since the relationship between delay and velocity is reciprocal (see eqn 2) the available points are concentrated at low velocities, leaving few higher up the scale. For example, for an MEC where \( d = 3 \) mm, for \( v = 100 \) m/s, from eqn (2), \( T = 30 \) μs, i.e. about one sample period. So in this case an inter-electrode delay of 1 sample period corresponds to a velocity of 100 m/s, 2 samples delay represents 50 m/s, with no possible velocity values between the two.

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Tuning curves were introduced as a compact method to display the velocity spectral behavior of a VSR system by analogy to techniques familiar from the frequency domain analysis of linear systems [1]. The magnitude of the output of the system is plotted as a function of velocity and so the method is somewhat analogous to computing the frequency response of a linear system. This analogy has been expanded to propose a metric $Q$, to quantify velocity selectivity, the idea being based on the quality factor familiar from the study of linear resonant circuits [3]. $Q$ is defined as $v_v/(v_1 - v_2)$ where $v_v$ is the matched velocity, $v_1$ and $v_2$ are the velocities where the normalized amplitude of the tuning curve drops to $1/\sqrt{2}$ either side of the matched velocity ($v_v$, above and $v_2$, below). More recently, an additional parameter, $R_v$, has been introduced in an attempt to quantify velocity resolution [6]. $R_v$ is defined as $\Delta v/v$, where $\Delta v$ is minimum available velocity step at propagation velocity $v$. It was suggested in [6] that a reasonable maximum value for $R_v$ across the velocity band of interest would be about 0.1. For example, with the values in the previous paragraph, at a propagation velocity of 10 m/s, $R_v = 0.11$, which is satisfactory, but at 100 m/s, this has risen to 0.5, which is clearly not. These parameters are employed in the analysis and comparison of systems in the following paragraphs.

It has been shown that the first problem (low velocity selectivity) can be resolved by placing bandpass filters at the output of the system [2]. However without a solution to the resolution issue, this would be ineffective since no data at high velocities can be recorded. Conversely, although the resolution problem can be solved using some form of interpolation, this does not improve the situation at high velocities if the underlying selectivity is inadequate. In the next section we propose an alternative process to delay-and-add that offers an effective solution to both problems.

![Fig. 7. A section of tripolar training data with APs that are 400 samples apart, corresponding to a duration of 12ms. This allows for the maximum delay encountered by an AP in this system (1.2ms). The section contains four APs with propagation velocities of 20, 21, 22 & 23 m/s and the SNR is 1 in all cases. For the system shown here, the MEC has 11 electrodes with 3mm separations and the signal is sampled at 32.5 kSa/s. The signals Ch1 to Ch9 correspond to tripolar signals from the 11 unipolar signals of the MEC as shown inset in Fig. The target shown is that required for training the ANN to detect APs with velocities of 22 m/s. In the full training sequence, this target amplitude is zero throughout the rest of the training sequence whilst the AP velocities increase to 120m/s.](image-url)
2) Artificial neural network method for VSR

Figure 5 shows the network structure used in this study. The network selected is a feed-forward type, specifically a time-delay neural network. As already noted this network has been applied in analogous applications such as speech recognition [18] and time series prediction [21]. As in the case of the delay-and-add method, one complete structure is required for each velocity to be classified. As in Fig 4, the inputs to the ANN derive from the outputs of the DEMUX but in this case the inputs are normalised (variable spike amplitudes are a potential issue in assessing the performance of the ANN but have not been considered in this study). Each of the nine tripolar output channels (1 ≤ i ≤ 9) forms the input to a shift register whose outputs are multiplied by a set of programmable weights w_j. The length of the shift registers is determined by the maximum required delay and this is set by the minimum velocity to be considered (in this case 10 m/s) and the inter-electrode spacing, d. Using eqn (2) with d = 3 mm, the corresponding inter-channel delay at this velocity is 150 µs or 1.2 ms for the complete 9 channel tripolar system. For a sampling period T_s of 30 µs, each shift register therefore requires 40 stages (1 ≤ j ≤ 40) and hence for each training set a matrix of (9 x 40) weights is computed and stored. The output consists of the sum of 9 overlaid weighted sets of 40 samples each, the values at each sample point being different from set to set reflecting the time shift along the MEC. The summed output is then passed through a second layer consisting of a single neuron, which is realised as a linear or nonlinear (Hyperbolic Tangent Sigmoid) function.

3) ANN training method

The time-delay neural network requires supervised learning using pre-determined targets after which the system can be run in real-time. Two types of target were tested: (i) a complete tripolar AP scaled appropriately for d and v (see eqn 2 and [2]) and (ii) a rectangular pulse derived from the AP. The pulse width and amplitude of the pulse target were derived from the positive phase of the AP at the points where its amplitude had fallen below the peak value by 3 dB, as illustrated in Fig 6. The pulse therefore expresses the delay, amplitude and duration of the AP. The recording duration required to capture the entire transit of an AP through the cuff is a function of propagation velocity, the electrode spacing d, the number of electrodes and the TMAP model function itself. The pulse as derived expresses the propagation velocity since the other parameters are constants. For the training process itself, a set of 9 tripolar input signals of normalised amplitude are applied in parallel and the summed output is then compared to the target waveform generated from the analytical TMAP model function (eqn (1)) as described above. The input signals are generated by the synthesiser, which, as already noted, adds noise to each channel as required.

Each of the records from the 9-tripolar channels is divided into time segments and each segment contains a single AP of normalised amplitude propagating at a particular velocity. For a complete set of 9 segments the relative delays between the pulses define the unique propagation velocity of the AP in each segment. Note that the minimum length of the segments is set by the total length of 9 APs delayed to represent the slowest propagation velocity under consideration. For example, in the cases considered in this paper, the data were sampled at f_s = 32.5 kSa/s. For a typical 3 cm separation between the end-electrodes of the MEC and a minimum AP velocity of 20m/s the delay is 1.2 ms (8 times T). The chosen delay line length of 40 samples (corresponding to a duration of 1.2 ms) is therefore more than adequate for the full range of propagation velocities under consideration. Fig 7 shows four example time segments taken from the training set to which noise was added (signal-to-noise ratio (SNR) = 1-see below). The tripolar APs in these segments have propagation velocities ranging from 20 to 23 m/s.

The summed output is compared to the target representing one velocity and coincident with the final (i.e. the most delayed) channel input at the matched velocity. At all other times the target amplitude is zero. The difference between the
training data and the template forms the basis of the backpropagation training process [22], [23]. Note that the designer can effectively choose the scale on the velocity axis as part of the training process, to some extent removing the problem of reduced velocity resolution at high propagation velocities encountered by the delay-and-add method.

III. RESULTS

A. Noiseless tuning curves

As an initial step, the system was set up to detect three velocities in the appropriate range (20 m/s, 50 m/s and 90 m/s). This requires three networks of the type shown in Fig 4 (for delay-and-add) or Fig 5 (for ANN), one for each target velocity. Tuning curves were then computed for each trained velocity [1]. The results are shown in Fig 8. Figs 8(a) and (b) show the outputs obtained using the conventional linear delay-and-add method and display the characteristic difficulties of this approach as described above in Section II.B.1 (Conventional VSR signal processing: ‘delay-and-add’). Note particularly the decline in velocity selectivity as the matched velocity increases. This is shown by the increasing width of the peaks as the peak position moves to a higher velocity and eventually leads to the absence of a peak in the 90 m/s curve (Fig 8(a)). This is an artefact of the method and when interpolation (1 to 8) is introduced (Fig 8(b)), the peak is clearly visible, albeit at an incorrect velocity. By comparison, consider the plots shown in Figs 8(c) and (d) which are the outputs from the ANN structure shown in Fig 5. In Fig 8(c), the ANN was trained using a complete AP derived from TMAP1 while in Fig 8(d) the target was a rectangular pulse derived from TMAP1 as described above. The improvement in the response is dramatic, both in terms of selectivity (the narrowness of the response) and the accuracy of the detected velocities. This is especially true for the curves shown in Fig 8(d). Quantitatively, calculating the velocity Q factor ($Q_v$) discussed earlier, for the curves in Fig 9 [1]–[3], for the 50 m/s case, the comparative values are 1.2 (delay-and-add: both cases); 7.1 (ANN with AP target) and 12.5 (ANN with pulse target).

The ability to discriminate the direction of travel of the AP through the MEC was also tested and it was found that for a 90m/s trained system, the maximum normalised response to an AP travelling in the opposite direction to the training was 0.1. For a 20m/s trained system, the maximum normalised response was just 0.02.

B. The addition of noise.

The tuning curves shown in Fig 8 are ideal in the sense that they were generated using noiseless training data and a noiseless target. A practical system is required to reconstruct the original signal in the presence of noise and/or partial or incomplete data (generalisation) and so the network was retrained with data to which noise had been added. Six sets of 116 (= 696) APs were generated by the synthesiser with a range of conduction velocities from 4 m/s to 120 m/s in steps of 1 m/s, each set having a different SNR value (0.1, 1, 5, 10, 25, 50, 100).

The trained network was tested using noisy input data and the results are expressed as the tuning curves shown in Fig 9, for SNRs of 0.1 and 1. The figure compares the effect of the noise on the interpolated delay-and-add system and on the ANN trained with both AP and pulse targets. Comparing these results with the noiseless ones in Fig 8 it is clear that the delay-and-add method is more susceptible to noise than the ANN. This is seen in the increase in relative response to non-matched APs which makes the matched APs less easy to distinguish. For the SNR = 1 example, $Q_v = 1.1$ for delay-and-...
**IV. DISCUSSION**

A. The need for the new methods proposed in this paper

The delay-and-add system in its current form was first proposed as a method to provide velocity spectral analysis of multi-channel propagating AP in 2004 [1]. Its properties have subsequently been discussed [1]-[6]. Although it provides a very robust approach to the problem of velocity spectral analysis, there are a number of key weaknesses, as noted in section II.B.1 (Conventional VSR signal processing: ‘delay-and-add’). In particular, both the velocity selectivity and velocity resolution are limited, especially at high velocities and the various methods that have been proposed to deal with these problems have met with limited success.

B. Choice of ANN and its effectiveness in this application

In order to achieve more effective velocity spectral analysis than is possible using the delay-and-add method, a non-linear approach has been adopted here using, specifically, a time-delay neural network [18], [21]. The choice of this type of network was motivated by clear analogies between speech and time-series prediction and multichannel VSR, when the latter is treated as a velocity classification process. In addition, since a time-delay neural network requires supervised learning, once trained it can run in real-time, which is a significant advantage in this application. Note that all of these processes (i.e. rather than some of them) could have been carried out in MATLAB but it was decided to use real amplifiers and filters so as to be as close as possible to a final system suitable for implantation.
The performance of the ANN shown in Figs 9 and 10 significantly outperforms a delay-and-add system having the same number of channels in terms of both velocity selectivity ($Q_v$) and resolution ($R_v$). This improvement can be explained as follows. A delay-and-add network of the type shown in Fig 3 can be regarded as an array of $N$ ($N$ is 9 in this case) FIR filters each having a single weight of fixed value (unity). By contrast the $N$ filters that form the basis of the ANN described here each have 40 weights, all independently programmable, offering many more degrees of freedom for shaping the overall response than is possible for simple delay-and-add. When used in combination with a nonlinear transfer function, a very effective velocity classifier is obtained. It should also be noted that when the network is trained using a rectangular pulse derived from the target AP as described in section II.B.3 (ANN training method), a further, very significant increase in selectivity is obtained. There are two main reasons for this. Firstly, since the AP is replaced by a pulse whose width is less than the AP itself, there is inevitably an increase in selectivity [2]. Secondly, since in the training process noise is only fed back when the pulses (i.e. the training data and the target) are non-zero, the noise power will be significantly less than when the complete AP is used, increasing the accuracy of the training algorithm.

The high selectivity and robustness of the ANN approach offers considerable promise for future therapeutic applications. For example, the two main bladder afferents in humans (stretch and tension) have propagation velocities of 44 and 38 m/s respectively [24]. In order to separate these using VSR, we require $Q_v$ to be about 41 m/s divided by 6 m/s or about 7. The data in Fig 10 suggests that this is possible for the pulse-trained ANN even in the presence of significant levels of noise.

C. Hardware considerations

Table II shows the device utilization summary and estimated power consumption for both approaches. Compared with the delay-and-add system, the ANN requires approximately double the number of gates and double the power consumption. However this increase should be considered in the context that our design is implemented using FPGA\(^1\), which is well known to have significantly higher power consumption than a comparable custom device of the type that would be required in an implantable system. For example we estimate that moving to a (conservative) 0.18um process would be enough to counteract the impact of the additional processing in the ANN-based system.

In an implantable system based on the type of ANN described in this paper the only change needed to detect different velocities is a change in the weights. The system is therefore adaptable to different situations and applications. For example whilst a complete set of ANNs could be constructed to detect APs with propagation velocities in the range 10 – 120 m/s with, say, 1 m/s resolution, this would not be necessary in a system designed to detect human bladder afferents which occur in the range 30 – 50 m/s. This allows considerable flexibility in the system design in terms of size and power consumption. Furthermore, in all these instances the weights required to realize ANNs sensitive to particular velocities can be generated off-line (i.e. unsupervised) and simply downloaded from a database as required.

REFERENCES


\(^1\) The exact device used was a Xilinx Spartan 3E: XC3S1200E-5FG320C.


