High-speed visualization of disturbed pathlines in axial flow ventricular assist device under pulsatile conditions

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ABSTRACT

Objective: To investigate potentially prothrombotic flow patterns within an axial flow ventricular assist device under clinically relevant pulsatile hemodynamic conditions.

Methods: A transparent replica of the HeartMate-II left ventricular assist device (Thoratec, Pleasanton, Calif) was visualized using a high speed camera at both low and high frame rates (125 and 3000 fps). Three steady-state conditions were studied: nominal (4.5 lpm), low flow (3.0 lpm), and high flow (6.0 lpm). Time-varying conditions were introduced with an external pulsatile pump that modulated the flow rate by approximately $\pm 50\%$ of the mean, corresponding to a pulsatility index of 1.0.

Results: At nominal and high flow rates, the path lines within the upstream region were generally stable, well attached, and streamlined. As the flow rate was reduced below 3.8 lpm, a rapid transition to a chaotic velocity field occurred, exhibiting a large toroidal vortex adjacent to the upstream bearing. The pathlines in the downstream stator section were consistently chaotic for all hemodynamic conditions investigated. It was common to observe tracer particles trapped within recirculation bubbles and drawn retrograde, causing repeated contact with the bearing surfaces. The addition of pulsatility caused the flow field to become periodically chaotic during the diastolic portion of the cardiac cycle depending on the instantaneous flow rate and acceleration.

Conclusions: The contribution of pulsatility by the native heart may induce a periodic disturbance to an otherwise stable flow field within an axial flow ventricular assist device, particularly during the diastolic and decelerating portion of the cardiac cycle. Potentially prothrombotic flow features were found to occur periodically in the region of the upstream bearing. (J Thorac Cardiovasc Surg 2015;150:938-44)

Path lines within axial flow ventricular assist device during the decelerating phase of the cardiac cycle.

Central Message

Excessively slow pump speed causes chaotic and potentially prothrombotic flow patterns in an axial flow pump.

Perspective

Recent practice in managing continuous flow VADs has promoted reducing the pump speed to allow opening of the aortic valve. This study illustrates the adverse effect of excessively slow speed on the flow surrounding the bearings, particularly during the diastolic phase of the cardiac cycle. This may partially explain the recent rise in incidence of device-related thrombosis.

See Editorial Commentary page 945.

Video clip is available online.

Rotodynamic (ie, continuous flow) ventricular assist devices (VADs) generate pressure and flow by rapid rotation of an impeller, imparting momentum to the blood.¹⁻⁷ The resulting rotational flow pattern is susceptible to the formation of vortices and other disturbances that may

promote thrombosis.^{8,9} These conditions may be exacerbated by the flow pulsatility contributed by the native heart. In other words, under nominal conditions, although the average flow may be within an acceptable range, the instantaneous flow at discrete points within the cardiac cycle will be less than (and greater than) the average displayed by the controller. In addition, the rate of change of flow rate causes acceleration and deceleration, which independently influences the flow field. Hence it is possible, in fact likely, that the flow patterns within the

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Abbreviations and Acronyms

dQ/dt = rate of change of flow rate
 LVAD = left ventricular assist device
 VAD = ventricular assist device

pump will be disturbed during portions of the cardiac cycle depending on the intensity of pulsatility. These factors have become increasingly relevant with the noted observance of a rising rate of left VAD (LVAD) pump thrombosis in the HeartMate II implantable LVAD (Thoratec, Pleasanton, Calif)¹⁰⁻¹² requiring device exchange¹³⁻¹⁶ and/or resulting in adverse cerebrovascular events.^{10,12,15,17} A number of mechanisms have been described as triggers of this complication that relate to the principles of Virchow's triad.¹⁸ This study focuses explicitly on the hemodynamic component of the triad, independently from properties of the blood and surface.

We previously demonstrated that the stability and coherence of the fluid pathlines within the HeartMate II axial-flow blood pump under steady hemodynamic conditions could be classified in terms of 3 ranges of operation: nominal, low-flow, and high-flow. The latter 2 conditions were characterized by recirculation regions believed to be prothrombotic.¹⁹ The purpose of this study was to extend this investigation to include clinically relevant pulsatile conditions, focusing primarily on anomalies that may be seen within the low-flow range of operation.

METHODS

The flow field within a transparent replica of the HeartMate II was visualized by high-speed videography (See Figure 1). The pump consisted of a flow straightener having 3 airfoil-shaped stay vanes followed by an axial rotor wrapped with 3 blades, and an outlet stator also wrapped with 3 blades in the reverse direction, all within a 12-mm diameter cylindrical channel. The rotor is radially and axially supported by 2 ball-and-cup bearings located at each end at the interface with their respective stator. The rotor was actuated by an integrated motor composed of electromagnetic coils, located outside the flow tube, that cooperate with permanent magnets within the rotor. The pump was placed in a simplified circulatory simulator that included a pneumatic pump to simulate the pulsatile contribution of the native left ventricle (see Figure 2). The differential pressure across the pump was measured by a pair of high-fidelity, sealed stainless steel transducers (EW-07356-54; Cole-Parmer, Vernon Hills, Ill). The instantaneous flow through the pump was measured by transit-time flowmeter (ME-PXN Sensors; Transonic Systems, Ithaca, NY). A transparent blood analog fluid consisted of a solution of water/glycerin (33% by volume) that was titrated to a viscosity of 3.5 cP and a density of 1.08 g/cm³ at room temperature to approximate the asymptotic viscosity of whole blood. Neutrally buoyant 30 μ m fluorescent microspheres (Thermo Scientific, Waltham, Mass) were seeded into the circuit as tracer particles for visualization of pathlines.

The regions of interest for visualization were centered at the upstream and downstream bearings (indicated in Figure 1), encompassing the upstream and downstream stator blades. The flow field was illuminated with a laser sheet (approximately 1 mm thick) using a diode-pumped solid-state laser (532 nm wavelength, DPSSL-1000; CNI Optoelectronics Technology Co Ltd, China), created by cylindrical lens (5 mm diameter,

3.7 mm focal length) in series with a convex lens (5 mm focal length). Background glare was attenuated using an optical longpass filter (550 nm) (FGL550s; Thorlabs, Newton, NJ) providing excellent contrast of the fluorescent particles. Video was acquired using a high speed CMOS camera (Fastcam SA4; Photron USA, Inc, San Diego, Calif) at 2 different frame rates: 125 fps and 3000 fps. The experimental matrix encompassed 3 average flow rates: 4.5 lpm (nominal), 3.0 lpm (low), and 6.0 lpm (high) with the rotor speed maintained at 9000 rpm. For each of these average flow rates, pulsatile conditions were also visualized in which the flow rate was modulated by approximately 50% of the average flow; that is, 2.0 to 6.8 lpm (nominal), 1.6 to 4.3 lpm (low), and 3.0 to 9.0 lpm (high). This corresponded to a pulsatility index of approximately 1.0.²⁰

RESULTS

The flow fields in the upstream and downstream bearing regions exhibited markedly different characteristics. These observations are summarized below for the conditions studied.

Steady-State Flow

The pathlines in the region of the upstream bearing were observed to be generally stable, well attached, and coherent at the nominal steady-state flow condition (4.5 lpm at 9000 rpm). Figure 3, A, provides a representative video frame within a visualization plane approximately 2.6 mm anterior of the centerline (1 mm along the span on the blade). When the flow rate was reduced below 3.8 lpm (9000 rpm, 68 mm Hg), a transition in the character of the flow field was observed wherein toroidal vortices appeared between the trailing edge of the upstream stay vane and the leading edge of the impeller, encompassing the front bearing. This is illustrated in Figure 3, B, for a flow rate of 3.0 lpm (9000 rpm, 72 mm Hg), which was also representative of the pathlines observed between 1.0 and 3.8 lpm. As the flow was reduced below 2.0 lpm, the vortex was observed to migrate further upstream within the bladed region of the stator. The path lines in aft stator region were observed to be chaotic for all conditions studied, with virtually indistinguishable differences between 1.0 and 6.0 lpm.

Pulsatile Flow

Activation of the in-series pneumatic pump introduced an approximately sinusoidal modulation of the volumetric flow rate and differential pressure. Representative waveforms are shown in Figure 4, A. To evaluate the influence of pulsatility, the characteristics of the flow field were assessed in terms of the instantaneous flow rate and rate of change of flow rate (dQ/dt) as reported previously by Shu and colleagues²¹ (see Figure 4, B). In general, the stability of the flow field in the upstream bearing region was observed to vacillate between 2 states, designated as streamlined flow (Figure 4, panel I) and disturbed flow (Figure 4, panel II), closely resembling the high-flow and low-flow steady-state conditions reported in Figure 3, A and B. By observing the video recordings frame by frame, it was possible to identify the approximate point in the cardiac cycle at which the

FIGURE 1. Transparent replica of the HeartMate II ventricular assist device (Thoratec, Pleasanton, Calif) and schematic of the visualization area.

flow field transitioned between these 2 states—indicated by an open circle in [Figure 4, B](#). It was found that the transition between coherent to disturbed flow was a function of both instantaneous flow rate and dQ/dt . During ventricular systole, in which the flow was accelerating ($dQ/dt > 0$), the transition point occurred at a lower instantaneous flow rate than when the flow was decelerating ($dQ/dt < 0$). This was most apparent for the nominal flow case (4.5 lpm average), for which approximately one-third of the cardiac cycle was observed to be chaotic. [Figure 5 \(top\)](#) provides representative video frames for the accelerating and decelerating flow at the same instantaneous flow of 4.5 lpm (indicated by closed circles in [Figure 4, B](#).)

The low-flow condition (3.0 lpm average) exhibited chaotic pathlines for approximately one-half of the cardiac cycle. At an average flow of 2.0 lpm (not shown), the flow was disturbed for the majority of the cardiac cycle but exhibited a brief period of stable flow during the accelerating portion of the cycle. Conversely, the high-flow case (6.0 lpm

average) exhibited a brief period of unstable flow during the decelerating portion of the cycle. In all cases, the initial wave of acceleration appeared to displace the vortex structures, washing them downstream. The pathlines in the stator region were highly chaotic throughout the cardiac cycle of all hemodynamic conditions studied. (See [Figure 5, bottom panels](#).) High speed videography (3000 fps) revealed the same general behavior but made it possible to observe the trajectory of individual particles. The latter results are best appreciated in [video](#) format, provided in the [Online Data Supplement](#). Under low-flow conditions, it was observed that particles were commonly trapped within recirculation bubbles and drawn antegrade along the nose cone causing repeated contact with the bearing surface. In the stator region, for all hemodynamic conditions, the particle pathlines exhibited large swirling trajectories. Various coherent structures were periodically observed rotating both clockwise and counterclockwise, at times reversing direction during the cardiac cycle.

DISCUSSION

Despite great advances in the development and use of continuous flow VADs, the incidence of device-related thrombosis remains high.^{10,11,22-25} The problem is understandably multifactorial and complex.^{10,26} Known contributing factors can be categorized as relating to the design of the device, intrinsic properties of the patient's blood, anticoagulation strategy, and hemodynamic parameters. More recently, variations in pump implant techniques, influencing the inflow cannula positioning within the left ventricle, have been shown to be related to the incidence of pump thrombosis.²⁷

Practice patterns can also modify the operating conditions of the pump. This was most relevant as reported by several investigators who have recommended operating the HeartMate II at lower speed and flow range in an attempt to encourage opening of the aortic valve to prevent the development of aortic insufficiency.²⁸ This has the effect of reducing flow across the inflow bearing and also introducing variable amounts of pulsatility as the native ventricle contributes more to the total cardiac output. Until recently, the encouragement of myocardial recovery on an LVAD has not been acknowledged as a risk factor for pump thrombosis. However, in a recent report,²⁹ pump thrombosis was noted in association with operating the pump at a lower speed during the weaning phase of the HeartMate II in a myocardial recovery trial (Remission from Stage D Heart Failure). Finally, 1 of the most disconcerting complications seen in continuous flow LVAD recipients is chronic gastrointestinal bleeding, which often results in withholding anticoagulation therapy.³⁰ This practice may also have implications for pump thrombosis under circumstances where flow patterns across the pump bearing are not optimized.

FIGURE 2. Transparent replica of HeartMate II (HM II) ventricular assist device (Thoratec, Pleasanton, Calif) inserted into a simplified circulatory simulator, illuminated by a laser sheet.

FIGURE 3. Fluorescent particle pathlines within the upstream and downstream sections of the pump (inset) under steady conditions. A, At the nominal flow rate of 4.5 lpm the pathlines near the upstream bearing are generally aligned and coherent. B, Below 3.8 lpm, secondary flow disturbances become evident; shown here for 3.0 lpm. C and D, Pathlines in the aft stator are consistently chaotic for all flow rates studied (1.0-6.0 lpm).

Because of the many factors contributing to pump thrombosis, it is virtually impossible to consider all combinations, variations, and permutations in preclinical or clinical trials. Therefore, flow visualization studies such as those presented here might serve to interpolate (and possibly extrapolate) the relationships observed clinically. These experiments are not intended as a substitute for formal verification and validation, but as a supplementary tool to gain insight to the internal hemodynamic parameters as influenced by flow rate and pulsatility. This study employed flow visualization techniques reported previously^{19,31-33} that provide only a qualitative assessment of the flow fields. Although more sophisticated and quantitative methods are available, we found that a great deal of insight was provided by direct observation of the tracer pathlines. Additional limitations of the protocol

should be acknowledged, such as the possible disparity in hemorheology of the blood analog fluid versus whole human blood, and the use of neutrally buoyant microspheres to visualize pathlines of the flow. However it is not likely that errors introduced by these approximations would alter the conclusions presented here.

It is well known that rotodynamic pumps, such as the axial-flow pump studied here, are designed to operate at 1 specific, steady-state, best efficiency point defined in terms of pressure, flow, and impeller speed. For example, the inflow angle of the impeller blade is designed to match the incipient flow angle at 1 particular combination of flow and impeller speed. This is rarely the case in practice owing to the variability of patient size, volume status, afterload, and other factors. Further deviation from the ideal operating point is caused by the pulsatile contribution

FIGURE 4. A, Representative hemodynamic parameters for pulsatile condition, shown here for 4.5 lpm average flow. B, Same data plotted in terms of acceleration (dQ/dt) versus flow (Q) for 1 cardiac cycle, at 3 average hemodynamic conditions. The shaded region of panel II indicates conditions wherein disturbed flow was observed in the inlet stator region.

FIGURE 5. Unsteady (pulsatile) hemodynamic conditions for average flow of 4.5 lpm. Left panel, Representative flow fields for the accelerating phase of the cardiac cycle (*panel I* in [Figure 4](#)). Right panel, Corresponding flow field for the decelerating phase (*panel II* in [Figure 4](#)).

of the native heart. Nevertheless, it is desirable to maintain blood flow within the device that is streamlined, absent of stagnation or recirculation regions to minimize the exposure of blood elements to the artificial surfaces and non-physiologic shear. From the perspective of the end-user, this is particularly difficult to achieve consistently, because the internal fluid dynamics are neither visible nor measurable. We aimed to provide some insight to the interrelationship of the observable variables, namely flow rate, pump speed, and pulsatility with the flow stability within the device. Specifically, using the flow-acceleration loops of [Figure 4](#), it is possible to intuit the proportion of the cardiac cycle in which the flow is laminar versus disturbed. With minor modifications to

the VAD controller, such loops could readily be displayed on a monitor screen.

It was not surprising that unstable, disturbed flow was exhibited at 3.0 lpm, which is below the typical use of this device. However it was revealing that under nominal conditions of 4.5 lpm average output, the flow field resembled 3.0 lpm for approximately one-third of the cardiac cycle. Under steady conditions, the approximate breakpoint below which vortices began to appear was 3.8 lpm. Under pulsatile conditions, this breakpoint was dependent on the degree of acceleration or deceleration of the flow; that is, lower during systole and greater during diastole.

This study was not intended to be a definitive nor exhaustive analysis of all possible combinations of pump speed,

pressure rise, flow rate, and pulsatility. However the results reported here may be translated to a wide range of conditions through the use of nondimensional analysis. For steady state flow, Wu and colleagues³¹ illustrated that the number of independent variables may be reduced to

$$\text{head coefficient : } \Psi = \frac{gH}{R^2\Omega^2}$$

$$\text{and flow coefficient : } \Phi = \frac{Q}{AR\Omega}$$

where H and Q are the pressure head and flow rate, respectively, R is the radius of the pump impeller, A is the flow discharge area, Ω is the rotational speed of the impeller, and g is the acceleration due to gravity. Not only does this simplify the experimental matrix, but also the extrapolation of these results to other combinations of flow rate, pressure head, and speed.

When the hemodynamic parameters are pulsatile, additional independent variables are introduced that can affect the flow field, including heart rate, stroke volume, and contractility. We previously demonstrated that these independent factors can be simplified by also considering the acceleration or rate of change of flow rate, dQ/dt , or more generally, the first derivative of the flow coefficient, $d\Phi/dt$.²¹ Therefore, all possible combinations of unsteady hemodynamic conditions can be described by 2 variables, depicted on the $\dot{\Phi}$ to Φ plane. For example, the transition point was found to be $\Phi = 0.08$ for steady-state flow, whereas the transition occurs at a higher instantaneous value when the flow is decelerating (eg, $\Phi = 0.12$ when $d\Phi/dt = -1.53 \times 10^{-7}$). Conversely, acceleration of the flow tends to provide a stabilizing effect, allowing a lower instantaneous flow rate that is free from upstream vortices.

Despite the above simplifications, the translation to clinical practice is not entirely obvious. In particular, what degree (percentage) of chaotic flow is tolerable is unknown. This ambiguity is further confounded by the coagulability of a patient's blood. Furthermore, the transition from streamlined to unstable to flow is not entirely abrupt.

On the other hand, these studies clearly illustrated that excessively low flow is more precarious than high flow. Extremely high flow (and impeller speed) is also undesirable due to shear stress on the formed blood elements,¹⁹ although our investigation did not detect any observable high-flow transition in the stability of the flow within the range studied.

The upstream bearing is known to be particularly vulnerable to thrombosis¹¹; therefore, it is reasonable to believe that sustained vortical flow, causing repeated cell contact with the bearing and extending the dwell time in this region, should be avoided. The fact that unstable flow persists in the aft stator for all conditions is also disconcerting. However, it

is reasonable to surmise that platelet activation and/or agonist generation produced in the downstream vortices are washed out of the pump, whereas those occurring upstream would be more problematic. These hypotheses warrant further investigation and are the topic of ongoing numerical simulation.

CONCLUSIONS

Flow patterns in an axial flow VAD observed in vitro by high-speed flow visualization revealed flow conditions under which potentially prothrombotic, chaotic flow features may occur. Under pulsatile conditions caused by the contribution of the native heart, disturbed flow patterns within the VAD depend on both average flow and pulsatility. Therefore, streamlines that are stable at a specified flow rate under steady-state conditions may become chaotic under pulsatile conditions during the diastolic portion of the cardiac cycle. Translation of these results to clinical practice may be simplified through the use of nondimensional indices; however, future research is necessary to develop definitive guidelines, particularly in situations where increased pulsatility may not be desirable in a pump that was designed to operate optimally under continuous flow conditions.

Conflict of Interest Statement

Authors have nothing to disclose with regard to commercial support.

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