Millar Pressure-Volume Systems MPVS-300/400 Series Owner's Guide



AT THE HEART OF SENSOR INNOVATION®

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Product:	MPVS-300 (P/N 753-2057)
	MPVS-400 (P/N 753-2058)

Hardware: Millar Instruments

Documentation: Millar Instruments

Document Number: 004-2142 Rev. B Copyright © 2004, 2005.

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Chapter

Safety Notes

Intended Use

Millar Instruments Pressure-Volume Systems (MPVS) have been designed for use in animal research applications. They are not intended for clinical or critical life-care use and should never be used for these purposes, or for the prevention, diagnosis, curing, treatment, or alleviation of disease, injury, or handicap.

Emergency Access to Power Cord

The power cord must not be blocked by other equipment or any panel or fixture. In case of an emergency, the user must be able to reach the cord to disconnect it.

Applicable Safety Standards

The MPVS-300 and MPVS-400 meet the requirements of the standards listed in the table below. Further information is available upon request.

Standard	Title
EN61010-1:2001	Product Safety for Laboratory Equipment
EN61326-1: A1:1998 and A2:2001	EMC Requirements for Measurement, Control and Laboratory Use

Explanation of Symbols

♦	Refers to earth ground connection. This connection is made through the power cord.
CE	EU Declaration of conformity
	Waste Electrical and Electronic Equipment

Preventative Inspection and Maintenance

Millar's MPVS-300/400 series hardware is maintenance free and does not require internal calibration or adjustment. There are no user-serviceable hardware parts inside the cabinet. The end user should make no attempts to service the hardware. There is no need for the end user to open the cabinet for inspection or maintenance, and doing so within the warranty period will void the warranty.

Chapter

Product Overview

This manual describes the basic features of the Millar Pressure-Volume System (MPVS), along with instructions on how to connect the system directly to a computer or a data acquisition system. Hardware models MPVS-300 and MPVS-400 are covered in this manual.

There are other manuals that may be required for use with the complete system. The PVAN user's manual describes the use of Millar's PVAN pressure-volume post analysis software, specifically with this system. The ADInstruments Chart user's manual describes the use of ADInstruments Chart[®] data acquisition software. The ADInstruments PowerLab[®] user's manual describes the use of the integrated data acquisition capabilities of the MPVS-400.

The MPVS simultaneously and continuously measures high-fidelity left ventricular pressure and relative volume from the intact beating hearts of small laboratory animals such as transgenic mice and rats. Using the MPVS, cardiovascular pressure and volume signals can be plotted against each other in real time, generating the characteristic pressure-volume (P-V) loops that are an excellent illustration of the cardiac cycle in normal or diseased conditions of the living heart. With the MPVS, P-V loops may be captured during pharmacological, therapeutic, and hemodynamic interventions, allowing comprehensive, beat-by-beat evaluation of the fundamental mechanical properties of the heart.

The MPVS supports the use of Millar's ultra-miniature P-V catheters that couple a high-fidelity pressure sensor with either four or six electrodes. The electrodes allow volume data to be acquired across a single segment of the left ventricle.

The MPVS-300/400 has an internal 20 KHz fixed-frequency oscillator which generates a 20µA RMS sinewave current, creating an electric field within the left ventricle of the heart. The system measures a corresponding voltage that is proportional to the electric field. The system continuously measures this voltage across a pair of receiving electrodes located near the catheter tip. The measured voltage changes during the cardiac cycle as the left ventricle blood volume rises (diastolic filling phase) and falls (systolic ejection phase). The measured voltage decreases as volume increases (resistance of blood pool decreases, conductance of blood pool increases) and the measured voltage increases as volume decreases (resistance of blood increases, conductance of blood pool decreases). The system then converts measured voltages into ventricular conductance which is proportional to the changes in ventricular volume as stated above. The MPVS-300/400 also measures high fidelity pressure signals using a Millar catheter. The solid-state pressure sensor uses the same technology found in Millar's extensive line of Mikro-Tip[®] catheter pressure transducers. The use of the pressure sensor located between the conductance/volume electrodes allows the simultaneous measurement of pressure and conductance/volume, enabling a researcher to obtain high quality P-V loops from the rapidly beating hearts of small animals. The MPVS system should ONLY be used with a Millar catheter.

The MPVS-400 has built-in data acquisition capability. The system incorporates hardware from the ADInstruments PowerLab[®] 4/25 four-channel data acquisition system. The operation of the data acquisition hardware is thus identical to that of the PowerLab 4/25 system. Two channels of the MPVS-400 are dedicated to acquisition of pressure signals. A third channel is dedicated to the acquisition of the conductance/volume signal. The fourth channel is available for the acquisition of an auxiliary voltage signal from a peripheral measuring device. This system does not require a separate data acquisition system since it can connect directly to the USB port of a computer. The required ADInstruments Chart data acquisition software is included with the purchase of the MPVS-400 signal conditioning hardware.

Chapter

Computer Requirements

The MPVS-300/400 hardware is designed for use with computers that run on a Microsoft[®] Windows[®] operating system.

Millar recommends the use of a reliable, name brand computer with a Pentium[®] 1GHz or faster processor, a hard disk with at least 40MB of available memory, Windows 2000, XP or later operating system, at least 256MB RAM, USB interface, CD-RW drive, an accelerated color Super VGA card (or equivalent), and a 16-inch or larger monitor.

Chapter

Descriptions of Controls and Indicators – Front Panels



MVPS-300 Front Panel



MVPS-400 Front Panel

Pressure Transducer/Calibration Switch

This switch allows the user to obtain the pressure signal from the transducer (catheter-tip pressure sensor) or a verification pressure signal of 0, 25, or 100 mmHg. The verification settings can be used to provide pressure calibration reference points when configuring the data acquisition software. In verification modes, the transducer is still powered and the balance control knobs have no effect on the 0, 25, or 100mmHg verification pressure signal output. The approximate fixed voltage outputs for the verification pressure signals are 0 volts for 0 mmHg, 0.25 volts for 25 mmHg, and 1.0 volts for 100 mmHg.

Pressure Transducer Balance Controls

The pressure transducer balance controls allow the user to ensure the transducer's pressure signal indicates 0 mmHg when the transducer is exposed to a zero pressure environment. Before inserting the catheter into the experimental subject, use the knob to control the balance of the transducer output while dipping the transducer just below the surface of sterile water or saline. Turn the transducer balance control knob until the output signal reads 0 mmHg (as displayed by an indicator in the data acquisition software or similar monitor) then lock the knob into position. Remove the catheter from the water or saline and insert it into the experimental subject.

For best stability and zero adjustment, shield the sensor from bright ambient light during the balancing procedure and allow the catheter-tip to soak in body temperature sterile water or saline for 30 minutes prior to insertion into the biological environment. For best results, zero balancing of the pressure signal should be done before each catheterization procedure.

Lighted Pressure Output Indicators

These indicators provide visual references for the magnitude of the pressure output signals. The indicators use the signals that are being sent to the output connectors, so they can be valuable trouble-shooting tools if no signal is seen on the computer that is being used to acquire and display the data. The indicators are not intended to give an accurate quantitative value for the pressure output signals.

Volume Transducer/Relative Volume Unit (RVU) Calibration Switch

This switch allows the user to obtain a volume signal from the transducer or a verification signal of 5, 25 or 50 RVUs. The verification settings can be used to provide volume calibration reference points when setting up the data acquisition software. The verification settings *simulate approximate volumes of 5, 25, and 50 µL*. Conductance values were measured in the volumes listed above using a four-electrode catheter with 4.5 mm spacing between electrode pairs and 0.9% saline solution.¹ Resistors were then selected to provide an equivalent system output, which may be used to estimate volumes at these settings. The relationship between RVUs and true volume depends on electrode spacing. Longer spacing between electrode pairs (i.e. larger measuring segment) will mean more volume per RVU. The approximate output voltages for the verification volume signal outputs are approximately -8 volts for 5 RVUs, -0.2 volts for 25 RVUs and 9 volts for 50 RVUs.

¹If a 0.9% saline solution is not available, dissolve 9.0 g of sodium chloride in water to make 1000 mL. These are the directions given in the United States Pharmacopoeia (USP XXII) National Formulary (XVII).

Lighted Conductance/Volume Output Indicator

This indicator provides a visual reference for the magnitude of the conductance/volume output signal. The indicator uses the signal that is being sent to the output connector, so it can be a valuable troubleshooting tool if no signal is seen on the computer that is being used to acquire and display the data. The indicator is not intended to give an accurate quantitative value for the conductance/volume output signal.

Segment Select Switch

This switch allows the user of a six-electrode selectable segment length catheter to activate different electrode pairs in order to take measurements from either a short or long measuring segment. A majority of Millar's small animal P-V catheters models have four electrodes. *When using a four-electrode catheter, leave the segment select switch in the <u>Short position</u>. When using a selectable segment length catheter, please remember to consider calibration changes if the segment selection is changed during the procedure. As noted above, the relationship between RVUs and true volume depends on electrode spacing (i.e. segment length). Longer spacing between electrode pairs (i.e. larger segment length) will mean more volume per RVU.*

Lighted Power Indicator

The green indicator will be lit when the system is receiving power.

Lighted Status Indicator (MPVS-400 only)

This indicator provides information about the status of the data acquisition technology inside the MPVS-400. It will be green when the system is operating properly but not communicating with a computer. It will be yellow when sampling or communicating with the computer. If the indicator flashes red, refer to the Connecting to a Computer section below for more information. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Chapter 5

Descriptions of Connections and Features – Back Panels



MPVS-300 Back Panel



MPVS-400 Back Panel

Catheter Connections

The MPVS-300/400 signal conditioning hardware is designed specifically for use with the Millar's SPR-series four (or six) electrode ultra-miniature P-V catheters. Cables are required to connect the catheter(s) to the hardware.

Either pressure input channel (CH1-P1 or CH2-P2) on the back panel can be used for the P-V catheter, provided the data acquisition system is properly configured. The hardware will also accept a second Millar Mikro-Tip pressure transducer input.

The pin-out of the pressure input connectors is shown in **Figure 1**. Refer to the Technical Specifications section of this manual for transducer requirements. All of Millar's Mikro-Tip pressure transducer catheters are compatible with the MPVS pressure input channels with the appropriate cable. Pressure cables differ in lengths and Millar catheter connector types (ex. low-profile, Redel, and Viking). Pressure cables are available from Millar Instruments or the authorized Millar distributor in your area.

The pin-out of the conductance/volume input connector (CH3-Vol.) is shown in Figure 2.

Redel connectors, such as those on the pressure and volume input cables, are locked when mated. Pull on the plastic collar to release the connector. BNC connectors must be twisted when mating and un-mating. Do not pull on the flexible portion of the cable to disconnect.

Transducer	Channel	MPVS-400 DIN 8 Pin	Input Cable
Pressure 1	1	7	PEC-4D
Pressure 2	2	5	PEC-4D
Volume	3	1	CEC-4B
Auxilary Analog Input	4	6	BNC termination

The auxiliary analog input of the MPVS-400 can accept voltage signals meeting the conditions described in the Technical Specifications section of this manual.

	Pin 1	+Excitation		Pin 1	E1
					(Tip)
	Pin 2	+Signal		Pin 2	E2
	Pin 3	-Signal		Pin 3	E3
	Pin 4	-Excitation		Pin 4	E4
	Pin 5	Shield		Pin 5	E5
				Pin 6	E6
Figure 1			Figure 2		
Pressure connector pin-out, viewed			pin-out, viewed from rear of		
trom rear of enclosure			enclosure		

Monitor/Analog Output (MPVS-400)

The MPVS-400 provides analog output of all four signals through an 8-pin DIN connector. The pin-out of the connector is shown in Figure 3. The pressure signals (CH1-P1 & CH2-P2) are scaled to 1V/100mmHg. The pressure signals' maximum output compliance is $\pm 5V$. The volume (CH3-Vol.) signal is scaled to approximately 0.4V/RVU. The volume signal maximum output compliance is $\pm 10V$. To disconnect, pull on the connector, not the flexible cable.

	Pin 1	Volume (CH3)
	Pin 2	Analog Ground
	Pin 3	Analog Ground
	Pin 4	Analog Ground
	Pin 5	Pressure 2 (CH2)
	Pin 6	Analog Input (CH4)
	Pin 7	Pressure 1 (CH1)
	Pin 8	No Connection
	Shield	Analog Ground
Figure 3		
Monitor/Analog output connector pinout, viewed from rear of enclosure		

Analog Output (MPVS-300)

The MPVS-300 provides analog output for all three channels at individual BNC connectors. The pressure signals (CH1-P1 & CH2-P2) are scaled to 1V/100mmHg. The pressure signals' compliance is $\pm 5V$. The volume (CH3-Vol.) signal is scaled to approximately 0.4V/RVU. The volume signal compliance is $\pm 10V$.

Analog Input (MPVS-400)

The MPVS-400 offers an additional channel of analog input (BNC connector) that allows for data acquisition of external signals. Therefore, the MPVS-400 can accept analog voltage outputs from other peripheral transducers (ex. temperature, flow, EKG, pH, etc.) that have this capability. Using the Analog Input connector, the signal from a peripheral measuring device can be recorded simultaneously with the P-V signals. The auxiliary analog input can record signals from micro volts to a maximum of ± 10 V. *Applying more than* $\pm 15V$ to the analog input can damage the internal circuitry

I²C Bus (MPVS-400)

The I²C Bus is an extra port designed to connect the MPVS-400 to specialized devices. The I²C Bus is not required for normal operation of the MPVS-400.

USB Connector (MPVS-400)

When using the ADInstruments Chart data acquisition software, the computer can communicate with the MPVS-400 directly through the USB connection. While a separate data acquisition system is not required, the analog output signals can still be accessed using the Monitor/Analog Output connector located on the back panel of the MPVS-400.

Earth Ground Terminal

This terminal connects directly to earth ground through the ground pin of the power cord. The outer conductors of the MPVS-300 BNC output connectors are grounded. The outer conductor of the MPVS-400 Analog Input BNC connector is not grounded.

Power Switch

This switch controls power to the MPVS-300/400 hardware. The indicators on the front panel should light up when the MPVS hardware is plugged in and Power Switch is flipped to the on position (|).

Power Input

This standard connector accepts grounded power cords. Grounding of the MPVS-300/400 hardware is important for signal integrity and user safety. There are no internal fuses on the power supply and there are no user-serviceable parts inside the cabinet.

The MPVS-300/400 hardware accepts 'universal' power as stated in the Technical Specifications section of this manual. A three-pin IEC power socket is used to connect the MPVS-300/400 to a three pin earthed (grounded) power cable. You should ensure that your power outlet is also a three-pin and that the grounded power cable is actually connected.

Millar Instruments provides standard U.S. and European style power cords with the MPVS-300/400 hardware. The end user may be responsible for providing his/her own power cords if the cords supplied by Millar are not appropriate for the country of operation. Cords must connect to a grounded power outlet for safety. To disconnect, pull on the connector, not the flexible cable.

Cooling Fan (MPVS-400)

The MPVS-400 can generate a fair amount of heat, so it is fitted with a cooling fan. The cooling fan is visible on the back panel of the MPVS-400. It maintains sufficient airflow in the enclosure to keep the circuitry within operating tolerances. The fan outlet must be kept unobstructed, and far enough from walls or other devices (at least three inches, 8cm) so as not to impede ventilation. Blocking the fan outlet may result in the MPVS-400 heating up enough internally to affect the performance of the system and reduce reliability. Periodically inspect the fan grate and the various vents around the edges of the front and rear panels for dust buildup. A buildup of dust can impede air flow and cause overheating of the equipment.

Chapter

Setup and Use

Please do not attempt to connect the MPVS-300/400 series hardware to a power outlet or computer, or turn the units on, until you have read this section. Please refer to the other manuals provided with this system for operation of the associated software and data acquisition hardware.

MPVS-300 Quick Start

- 1. Unpack the hardware, software, and cables.
- 2. Inspect the equipment for any obvious signs of external and/or internal damage. If anything is missing or the equipment seems to be damaged in any way, contact Millar Instruments or the authorized Millar distributor in your area immediately.
- 3. Power up the computer that will be used to acquire and analyze the P-V data and install the necessary software (DAQ software, Millar PVAN).
- 4. Connect the P-V channel outputs on the back panel of the MPVS-300 to the input channels on the data acquisition system using BNC-BNC cables or BNC cables that terminate at a connector that is compatible with the data acquisition system. Turn on the data acquisition system.
- 5. Connect the power cord to the power input jack on the back of the MPVS-300. Please ensure the power cord conforms to the standards in the country of operation. Millar Instruments provides standard U.S. and European style power cords with the hardware. The end user may be responsible for providing his own power cords if the cords supplied by Millar are not sufficient for the country of operation. Cords must connect to a grounded power outlet for safety.
- 6. Turn on the MPVS-300 using the power switch located on the back panel. Ensure that the Power Indicator located on the front panel lights up.
- Connect the pressure cable (PEC-4D or similar) and volume cable (CEC-4B or similar) to the appropriate input connectors on the back of the MPVS-300. Connect the second pressure input cable if desired.
- 8. Unpack a P-V catheter, being careful to handle it with a high degree of care.
- 9. Connect the pressure transducer connector on the P-V catheter to the pressure input cable (PEC-4D or similar).

- 10. Connect the conductance/volume transducer connector on the P-V catheter to the volume input cable (CEC-4B or similar).
- 11. Using a shallow dish, beaker, or syringe, soak the catheter tip in body temperature sterile water or saline for at least 30 minutes prior to insertion into the biological environment.
- 12. Set the Pressure Transducer/Calibration Switch on the front panel of the MPVS-300 to 0 mmHg.
- 13. Set the Volume Transducer/Relative Volume Units (RVU) Calibration Switch on the front panel of the MPVS-300 to 5 RVU.
- 14. Verify that the appropriate segment length has been selected using the Segment Select switch on the front panel of the MPVS-300. For standard four-electrode P-V catheters, select the "Short" position. For six-electrode P-V catheters, select either the "Short" or "Long" position depending on the desired electrode spacing (segment length).
- 15. Open the data acquisition software on the computer.
- 16. Configure the data acquisition software according to user preferences so that it can effectively acquire, display, and record the P-V data produced by the MPVS-300.
- 17. *Calibration Setup*: Follow the instructions in the data acquisition software user's manual as well as Millar's PVAN user's guide to perform two-point calibration procedures for both the pressure and volume channels. For simplified electronic calibration verification, use the transducer/calibration controls on the front of the MPVS-300 to provide fixed voltage signal references for pressure (0, 25, 100 mmHg) and volume (5, 25, 50 RVU) that may be used to quickly perform the two-point calibration procedure. Remember that RVU verification is only for Relative Volume Units, not absolute volume. Conversion of Relative Volume Units into true volume units requires additional steps (please refer to Millar's PVAN User's Guide for additional instructions regarding volume calibration techniques). Volume calibration for a given catheter may be performed using small tubes of known diameters (a calibration cuvette, which can be supplied by Millar Instruments) filled with fresh, slightly heparinized blood drawn from an experimental subject. Validation of the volume estimates provided by the Millar hardware can be performed by other means, e.g. by thermodilution, flow probes, hypertonic saline dilution, echo, etc.
- 18. *Pressure Transducer Balancing:* After the catheter tip has soaked for at least 30 minutes, set the Pressure Transducer/Calibration switch on the front panel of the MPVS-300 to Transducer. Unlock and adjust the Balance control knob for the pressure channel the catheter is connected to. Use the Balance control knob to adjust the pressure transducer output while dipping the transducer just below the surface of sterile water or saline (essentially a zero pressure environment). Adjust the Balance control knob until the output signal reads approximately 0 mmHg (as displayed by an indicator in the data acquisition software or similar monitor) then lock the knob into position. Repeat the procedure if an additional pressure transducer is connected to the MPVS-300. Pressure calibration may be verified using an external manometer (digital or mercury).

For best stability and zero adjustment, shield the sensor from bright ambient light during the balancing procedure and allow the catheter-tip to soak in body temperature sterile water or saline for 30 minutes prior to insertion into the biological environment. Zero balancing of the pressure transducer should be done prior to each catheterization procedure.

19. The system is now ready to begin conducting P-V experiments. Prepare the experimental subject for catheter insertion according to the lab's established experimental protocols. When the experimental subject is ready, insert the catheter into the left ventricle and begin recording P-V data [Please Note:

Millar Instruments recommends a data acquisition sampling rate of <u>1 kHz (1000</u> <u>samples/second)</u> in order to accurately capture all of the features of the P-V waveforms produced by the fast beating hearts of mice and rats]. To ensure activation of the catheter tip transducers, make sure that both the Pressure Transducer/Calibration Switch and Volume Transducer/Relative Volume Units (RVU) Calibration Switch on the front panel of the MPVS-300 are switched away from the calibration reference settings and are set to Transducer. After the P-V data has been collected by the data acquisition software, please refer to the PVAN software user's guide for instructions on how to export P-V data selections out of the data acquisition software to perform a comprehensive data analysis in PVAN.

20. When necessary, clean the MPVS-300 hardware with a cloth and isopropyl alcohol or water. Do not allow liquids to enter the chassis as this will damage the sensitive electronics housed inside. Air vents may be cleaned with suction (i.e. a vacuum cleaner). The MPVS-300 hardware has not been tested for compatibility with any method of sterilization.

MPVS-400 Quick Start

- 1. Unpack the hardware, software, and cables.
- 2. Inspect the equipment for any obvious signs of external and/or internal damage. If anything is missing or the equipment seems to be damaged in any way, contact Millar Instruments or the authorized Millar distributor in your area immediately.
- 3. Power up the computer that will be used to acquire and analyze the P-V data and install the provided software (ADInstruments Chart, Millar PVAN).
- 4. Connect the MPVS-400 to the computer using the USB cable provided. Please see the "Connecting the MPVS-400 to a Computer" section of this manual (page 17) for more information.
- 5. Connect the power cord to the power input jack on the back of the MPVS-400. Please ensure the power cord conforms to the standards in the country of operation. Millar Instruments provides standard U.S. and European style power cords with the hardware. The end user may be responsible for providing his/her own power cords if the cords supplied by Millar are not appropriate for the country of operation. Cords must connect to a grounded power outlet for safety.
- 6. Turn on the MPVS-400 using the power switch located on the back panel. Ensure that the Power Indicator and Status Indicator located on the front panel are both illuminated.
- 7. Connect the pressure cable (PEC-4D or similar) and volume cable (CEC-4B or similar) to the appropriate input connectors on the back of the MPVS-400. Connect the second pressure input cable and the auxiliary analog input BNC cable if desired.
- 8. Unpack a P-V catheter, being careful to handle it with a high degree of care.
- 9. Connect the pressure transducer connector on the P-V catheter to the pressure input cable (PEC-4D or similar).
- 10. Connect the conductance/volume transducer connector on the P-V catheter to the volume input cable (CEC-4B or similar).
- 11. Using a shallow dish, beaker, or syringe, soak the catheter tip in body temperature sterile water or saline for at least 30 minutes prior to insertion into the biological environment.

- 12. Set the Pressure Transducer/Calibration Switch on the front panel of the MPVS-400 to 0 mmHg.
- 13. Set the Volume Transducer/Relative Volume Units (RVU) Calibration Switch on the front panel of the MPVS-400 to 5 RVU.
- 14. Verify that the appropriate segment length has been selected using the Segment Select switch on the front panel of the MPVS-400. For standard four-electrode P-V catheters, select the "Short" position. For six-electrode P-V catheters, select either the "Short" or "Long" position depending on the desired electrode spacing (segment length).
- 15. Open the ADInstruments Chart data acquisition software on the computer.
- 16. Configure the Chart software according to user preferences (please refer to the ADInstruments user's manual(s) for more details). The PVAN CD contains a suggested Chart data acquisition interface to help investigators understand how to utilize the Chart software to record P-V data from the MPVS-400.
- 17. *Calibration Setup*: Follow the instructions in the ADInstruments Chart user's manual(s) as well as Millar's PVAN user's guide to perform two-point calibration procedures for both the pressure and volume channels. For simplified electronic calibration verification, use the transducer/calibration controls on the front of the MPVS-400 to provide fixed voltage signal references for pressure (0, 25, 100 mmHg) and volume (5, 25, 50 RVU) that may be used to quickly perform the two-point calibration procedure. Remember that RVU verification is only for Relative Volume Units, not absolute volume. Conversion of Relative Volume Units into true volume units requires additional steps (please refer to Millar's PVAN user's guide for additional instructions regarding volume calibration techniques). Volume calibration for a given catheter may be performed using small tubes of known diameters (a calibration cuvette, which can be supplied by Millar Instruments) filled with fresh, slightly heparinized blood drawn from an experimental subject. Validation of the volume estimates provided by the Millar hardware can be performed by other means, e.g. by thermodilution, flow probes, hypertonic saline dilution, echo, etc.
- 18. *Pressure Transducer Balancing:* After the catheter tip has soaked for at least 30 minutes, set the Pressure Transducer/Calibration switch on the front panel of the MPVS-400 to Transducer. Unlock and adjust the Balance control knob for the pressure channel the catheter is connected to. Use the Balance control knob to adjust the pressure transducer output while dipping the transducer just below the surface of sterile water or saline (essentially a zero pressure environment). Adjust the Balance control knob until the output signal reads approximately 0 mmHg (as displayed by an indicator in the data acquisition software or similar monitor) then lock the knob into position. Repeat the procedure if an additional pressure transducer is connected to the MPVS-400. Pressure calibration may be verified using an external manometer (digital or mercury).

For best stability and zero adjustment, shield the sensor from bright ambient light during the balancing procedure and allow the catheter-tip to soak in body temperature sterile water or saline for 30 minutes prior to insertion into the biological environment. Zero balancing of the pressure transducer should be done prior to each catheterization procedure.

19. The system is now ready to begin conducting P-V experiments. Prepare the experimental subject for catheter insertion according to the lab's established experimental protocols. When the experimental subject is ready, insert the catheter into the left ventricle and begin acquiring and recording P-V data [Please Note: Millar Instruments recommends a data acquisition sampling rate of <u>1 kHz</u> (1000 samples/second) in order to accurately capture all of the features of the P-V waveforms produced by the fast beating hearts of mice and rats]. To ensure activation of the catheter tip

transducers, make sure that both the Pressure Transducer/Calibration Switch and Volume Transducer/Relative Volume Units (RVU) Calibration Switch on the front panel of the MPVS-400 are switched away from the calibration reference settings and are set to Transducer. Please refer to the ADInstruments Chart software guide for additional details on operating the data acquisition software. After the P-V data has been collected, please refer to the PVAN software user's guide for instructions on how to export P-V data selections out of the Chart software to perform a comprehensive data analysis in PVAN.

20. When necessary, clean the MPVS-400 hardware with a cloth and isopropyl alcohol or water. Do not allow liquids to enter the chassis as this will damage the sensitive electronics housed inside. Air vents may be cleaned with suction (i.e. a vacuum cleaner). The MPVS-400 hardware has not been tested for compatibility with any method of sterilization.

Connecting the MPVS-400 to a Computer

Before the MPVS-400 is connected to a computer, it should be checked to make sure it is working properly.

The MPVS-400 performs a self-test each time it is switched on, whether or not it is connected to a computer. To check that the MPVS-400 is functioning properly, follow the instructions below.

- 1. Connect the MPVS-400 to a 3-pin earthed power outlet using the power cable that came with the unit (choose the appropriate power cord for the region of operation). Please ensure that the power outlet is properly earthed. Turn the power on at the wall.
- 2. Turn on the power switch located on the back panel of the unit.

When an MPVS-400 is turned on, the Power indicator light on the front panel should glow blue, while the Status indicator light should flash yellow and then remain green. *(See Table for Status Indicator states for the MPVS-400.)*

If the Power indicator light of the MPVS-400 is off when the power switch is on, then check that the unit is plugged into a power source, that the power outlet is working correctly (try connecting a different device to the power outlet), and that the power cable has no obvious problem (try a different power cable if possible). If there is no apparent cause for the lack of power then a fuse may have blown inside the MPVS-400 unit and you will need to contact Millar Instruments or the authorized Millar distributor in your area for technical support.

If the Status indicator light of the MPVS-400 flashes red, switch the MPVS-400 unit off, wait five seconds and turn it back on. If the red flashes continue, turn the unit off and contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Status Indicator	Meaning
Off	Idle and not yet initialized by the software.
Green	Idle, initialized and waiting for a command from the computer.
Yellow	Sampling or communicating with the computer.
Four red flashes and	The MPVS-400 has detected a temporary fault. Turn the MPVS-400
one yellow one	unit off, wait ten seconds and turn on again.
Red flashes	The MPVS-400 has detected an internal fault during the power-up
	test. It will repeat until the power to the MPVS-400 is turned off.

USB Connection

The MPVS-400 can connect to a computer using a USB (universal serial bus) port if the ADInstruments Chart software will be used to acquire the P-V data. USB is supported by computers running Windows 98, Me, 2000 or later operating systems (Windows 95 and NT4 do not support USB).

Connect the USB port on the MPVS-400 back panel to the USB port on the computer using the USB cable supplied with the MPVS-400 hardware (**Figure 4**). USB ports and cables are usually marked with a trident shaped icon,

Please observe the following points when using USB connection:

- Cable length should be less than 5 meters (16 feet) between devices.
- Never attempt to make your own USB cable or modify one.
- Do not disconnect the MPVS-400 from your computer while the ADInstruments Chart software is active.



Figure 4 Connecting the MPVS-400 with a USB cable

New Hardware Wizard

The first time a Windows computer is started up with the MPVS-400 connected, the New Hardware wizard may appear as shown in **Figure 5**. This occurs whether or not the ADInstruments Chart Software has been installed. Leave the wizard on its default settings and click the OK button. Insert the ADInstruments Chart Installer CD. The PowerLab setup information file will have been selected in the dialog box that appears. Click the OK button and Windows will continue starting up. This operation should only have to be performed once.

New Hardware Found	x
Unknown Device	
Select which driver you want to install for your new hardware:	
O Windows default driver	
• Driver from disks provided by hardware manufacturer	
O Do not install a driver (Windows will not prompt you again)	
O Select from a list of alternative drivers	
OK Cancel <u>H</u> elp	





Troubleshooting

This appendix describes most of the common problems that may occur when using the MPVS-300/400 series hardware. It covers how these problems are caused and what may be done to alleviate them. If the solutions here do not work, the ADInstruments Chart software guide or transducer Instructions for Use may contain remedies for other possible problems. If none of the solutions here, in the software guide, or the transducer IFU appear to help, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Power Indicator does not light up.

Make sure the power cord is firmly plugged into the back panel of the MPVS-300/400 hardware and that the unit is switched on (|).

Make sure that the power cord is plugged into a live power outlet.

If a power strip or uninterruptible power supply is used, be sure that this equipment is turned on.

If the Power Indicator flashes or is red, switch the unit off immediately. Check the voltage and frequency output of the power source to be sure that they are within the ranges specified on the back panel of the MPVS-300/400.

If the unit stopped operating after a long period of continuous operation, check the top and bottom of the chassis to see whether the unit might have overheated. Check the vent holes under the front panel and at the top of the rear panel. Make sure that the fan in the MPVS-400 is blowing.

If the power source and all connections are good, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

MPVS-400 Status indicator does not light up.

If the Status indicator is not lit at all, this may indicate a problem that will require factory maintenance. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

MPVS-400 Status Indicator flashes red and/or yellow.

An internal problem has been discovered by the diagnostic self-test. (The diagnostic self-test is performed by the MPVS-400 every time it powers up.) Turn the MPVS-400 off, wait at least five seconds and turn it back on again. This should clear a temporary problem. If the problem persists, the MPVS-400 may need repair. Take note of the colors and flashing pattern (please see the "Connecting the MPVS-400 to a Computer"

section of this manual for more information, page 17) and contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Pressure Output Indicator does not light up.

The Pressure Output Indicators light up at pressure inputs above approximately 20 mmHg. Pressures lower than this will not be indicated on the display.

If the pressure input is higher than 20 mmHg, check the connections from the transducer to the cable and the cable to the rear panel.

Turn the Pressure Control knob to the 25 mmHg and 100 mmHg calibration settings. The first and fourth LED's in both displays should light in these positions, respectively.

If the displays work properly in the calibration settings, but not when a transducer is connected, switch the transducer to the other pressure channel to see if the other display works. If there is still no response, try a different cable or a different transducer to determine whether the cable or transducer may be damaged.

If the displays do not function for any setting of the Pressure Control knob, then observe the output of the system on a voltmeter or data acquisition system at each of the calibration settings on each channel.

If the problem persists, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Volume Output Indicator does not light up.

The Volume Output Indicator should be lit for any volume input. Given a large enough change in input during a study, the indicator should change with each heart beat. However, for smaller stroke volumes, the change may not be sufficient enough to change the indicator. Stroke volumes of about 5 RVU or smaller may not cause the indicator to change with each heart beat.

If the MPVS-300/400 is on but the indicator is not lit, observe the output of the Volume channel at all three calibration settings with a voltmeter or data acquisition system.

If the Volume Output Indicator lights up, but does not change with changing calibration input, measure the output of the channel at each calibration setting.

If the problem persists, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

When Windows starts up it does not recognize the MPVS-400 hardware.

This should happen only the first time the MPVS-400 system is connected to the computer. Windows will bring up the New Hardware wizard and ask you if you want to install a driver. Leave the wizard on its default settings and click the OK button. Insert the ADInstruments Chart Installer CD. The PowerLab setup information file will have been selected in the dialog box that appears. Click the OK button and continue.

Please see the "New Hardware Wizard" section of this manual for more information (page 19).

If the problem persists, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

The computer cannot locate the MPVS-400 hardware when ADI Chart is launched.

The MPVS-400 may not be powered on. Check the power switch, the power cord, and the power source.

There may be a poor connection between the MPVS-400 and the computer or there may be a bad USB cable. Make sure that the cable is firmly attached at both ends and try again. If there is still a problem, try a new cable.

The MPVS-400 may have an internal problem or temporary freeze. Turn the computer and MPVS-400 off. After at least five seconds, turn the MPVS-400 back on. Then, turn on the computer and try using the software again.

The computer may not have adequate specifications. Check the minimum requirements listed in the "Computer Requirements" section of this manual (page 5) or on the ADInstruments Chart software package.

If the problem persists, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

The computer hangs up while recording from the MPVS-400, or there is data loss

There may be a poor connection between the MPVS-400 and the computer or there may be a bad USB cable. Make sure that the cable is firmly attached at both ends and try again. If there is still a problem, try a new cable.

If the problem persists, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

No signal appears to be coming from the USB output

If the computer and the ADInstruments Chart software have recognized the MPVS-400 and yet there does not appear to be a signal coming from the system via the USB connection, try the following.

The settings in the Chart software interface may be incorrect. Open the preconfigured data acquisition interface that can be found on the PVAN CD (this interface should be automatically installed onto your desktop by running the PVAN installer on the CD. There is one interface configured specifically for the MPVS-400 and a general interface for the MPVS-300 and previous versions of the Millar Pressure-Volume hardware). If this interface is able to acquire data, compare the settings in the Channel Settings dialog box for the pre-configured interface to those in the other interface.

If no data can be acquired with the standard interface, verify that no signal is seen when the calibration settings on the Pressure Control and Volume Control knobs are changed. If the indicators on the front panel change, but there is still no sign of change in the digital data, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

No signal appears to be coming from the Monitor/Analog output

Use a voltmeter and the wiring diagram in the Catheter Connections section of this manual (page 9) to verify that the cable is wired properly. The following voltages should be seen when checking the wiring:

Channel	Calibration Setting	Approximate Voltage
Pressure 1 and 2	0 mmHg	0 V
Pressure 1 and 2	25 mmHg	0.25 V
Pressure 1 and 2	100 mmHg	1 V
Volume	5 RVU	-8 V
Volume	25 RVU	-0.2 V
Volume	50 RVU	9 V

These signals should also be seen at the connector itself, using the pin-out in this manual as a guide.

No signal appears to be coming from the BNC outputs

Check the wiring of the output cable. The center conductor must be the signal (+) and the outer conductor must be the ground/shield.

Measure the voltage directly at the BNC output connector using a voltmeter. The voltage outputs should be as follows:

Channel	Calibration Setting	Approximate Voltage
Pressure 1 and 2	0 mmHg	0 V
Pressure 1 and 2	25 mmHg	0.25 V
Pressure 1 and 2	100 mmHg	1 V
Volume	5 RVU	-8 V
Volume	25 RVU	-0.2 V
Volume	50 RVU	9 V

Verify that each signal is functional by observing the respective indicators on the front panel. If the lights on the indicator respond to changes in the calibration settings, but there is no change in the voltage output, contact Millar Instruments or the authorized Millar distributor in your area for technical support.

The Transducer Balance Control does not appear to have any impact on the signal output

The Pressure Transducer Balance controls do not affect the pressure signals when the control switches are set to the calibration settings.

Be sure that you are adjusting the appropriate Balance knob. Changes to one channel's balance setting will not affect the other channel.

If a pressure transducer cannot be balanced by adjusting the Balance control for the appropriate channel, the transducer may be malfunctioning. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Signal output does not change when switching through the Pressure and/or Conductance/Volume calibration settings

Check the cable connections from the MVPS-300/400 hardware to the data acquisition equipment or computer.

Observe the lighted indicators on the front panel as the calibration settings are changed. If the indicators do not change, then the equipment may need factory service.

If the indicators do change with alterations to the calibration settings, and yet the signal output(s) do(es) not, then check the analog signal at the back panel connectors. If there is no signal at the rear panel connectors, then the equipment may need factory service. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

No conductance/volume data is coming from a four-electrode transducer

Verify that the Segment Select switch is in the "Short" position.

Verify the continuity of the CEC-4B cable following the schematic in the cable's Instructions for Use.

The electrodes on the P-V catheter may be coated with biological material. After *every* use, be sure to follow the cleaning instructions in the P-V catheter's Instructions for Use.

If the Segment Select switch is set correctly and the cable is functional and properly connected, check the P-V catheter's functionality. If no signal change is seen when the tip (all four electrodes) of the P-V catheter is dipped in a beaker of saline, then the P-V catheter may need factory service. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

The transducer balance changed significantly even after soaking the pressure sensor for more than 30 minutes.

Be sure to use the locking lever on the Transducer Balance controls. It may be necessary to hold the knob to keep it from turning as the lock is activated.

Note that the pressure transducer is sensitive to ambient light and there will often be bright light in the vicinity of the surgical field. When balancing the transducer and checking its balance at a later time, be sure to shield the pressure sensor from bright ambient light.

If the problem persists, then the transducer may need factory service. Contact Millar Instruments or the authorized Millar distributor in your area for technical support.

The MPVS-300/400 became unusually hot during operation.

The MPVS-400 has a fan for cooling the internal electronics. Be sure the fan vent hole is clean and unobstructed.

Be sure the vent holes located under the front panel and at the top of the back panel are clean and unobstructed.

Be sure the fan is blowing. If the fan is not blowing, the equipment may require factory service.

The MPVS-300 does not have a fan for cooling. Be sure the vent holes located under the front panel and at the top of the rear panel are clean and unobstructed.

If the problem persists, please contact Millar Instruments or the authorized Millar distributor in your area for technical support.

Appendix

Technical Specifications

Pressure Transducer Characteristics		
Specification	Value	
Transducer Sensitivity	$5 \mu\text{V/V/mmHg}$, nominal	
Bridge Excitation Load Resistance	1000 ohms, nominal	
	350 ohms, minimum	
Transducer Bridge Excitation	5.0 V _{DC} , nominal	
Signal Input Resistance	50 mega-ohms, nominal	
Input connector for system	Five-pin Redel	

Pressure Outputs	
Specification	Value
Sensitivity	1 V/100 mmHg, nominal.
Accuracy Error Band	<±1 mmHg or 1 % of reading, whichever is greater
Frequency Response	DC to 600 Hz (-3 dB), minimum
Output Resistance	1000 ohms, nominal
Noise	<0.3 mmHg peak-to-peak
Zero-Offset Temp Coefficient	<0.15 mmHg/°C
Gain Temperature Coefficient	<0.1 %/°C
Balance Adjustment Range	±140 mmHg, nominal

Pressure Standby-Calibration Mode	
Specification	Value
Zero Offset	<±1 mmHg
Calibration Steps	0, 25, and 100 mmHg
Calibration Accuracy	<± 0.5 mmHg

Lighted Pressure Output Indicators	
Specification	Value
Range	20 to 200 mmHg in 10 steps
Resolution	20 mmHg
Out of Range	Top light stays on at high pressures
	All lights off at low pressures and no transducer
	connected

Volume Outputs	
Specification	Value
Sensitivity	0.4V/RVU
Frequency Response	460 Hz
Output Resistance	65 ohms
Noise	$\leq 40 \mathrm{mV}_{\mathrm{RMS}}$
Input Connector	Six-pin Redel

Volume Standby-Calibration Mode	
Specification	Value
25RVU Offset (Subject)	$\pm 1.5\%$
Calibration Verification Steps	5, 25, 50 RVU
Calibration Verification Accuracy	± 1.5% @25RVU

Lighted Volume Output Indicators	
Specification	Value
Range	5-50 RVU
Resolution	5 RVU/step

MPVS-400 Data Acquisition	
Specification	Value
Sample rate	200kS/s, max, total across all channels
Frequency response	20kHz max. Software selectable down to 1Hz.
Voltage input range (Ch 4)	$\pm 10V$ (do not exceed $\pm 15V$ max)
Input impedance	>1 mega-ohm

Outputs	
Specification	Value
I ² C Bus	Standard connector
USB	Standard connector
Monitor (Analog) Out	8-pin circular DIN connector

Power	
Specification	Value
Input (Universal)	$90 - 250 V_{AC}$, $50 - 60 Hz$, $40 VA$
Connector type	EN60320/C13
Ground terminal resistance to	<1 ohm
power cord ground pin, 7-ft cord	

Enclosure	
Specification	Value
Ingress protection	IP-20

Warranty Agreement

Extent

This Agreement is between Millar Instruments, Inc. ('Millar') and the purchaser ('the Purchaser') of any Millar product – software, hardware, catheters, cables and/or accessories – and covers all obligations and liabilities on the part of Millar, the Purchaser, and other users of the product. The purchaser (or any user) accepts the terms of this Agreement by using the product. Any changes to this Agreement must be recorded in writing and have Millar's and the Purchaser's consent.

Copyright and Trademarks

Millar develops proprietary sensor technology, catheters, signal conditioning hardware and computer software including Mikro-Tip catheter transducers and PVAN software. All Millar software, hardware, and documentation are protected by copyright, and may not be reproduced or copied in any way, nor may products be derived from or based on it. Millar retains the exclusive ownership of the trademarks represented by its company name, logo, and product names.

Responsibilities

The Purchaser and any others using any Millar product agree to use it in a sensible manner for purposes for which it is suited, and agree to take responsibility for their actions and the results of their actions.

If problems arise with a Millar product, Millar will make all reasonable efforts to fix them. This service may incur a charge, depending on the nature of the problem, and is subject to the other conditions cited in this Agreement.

General Limitations

Millar products are produced to high standards, and should perform substantially as described in the supplied documentation. There is a limited hardware warranty, and technical support is provided for all products.

Nevertheless, since Millar products could be affected by external factors (for instance, an environment filled with electronic noise or the computer system on which the products are run), absolute performance and reliability cannot be guaranteed. No warranty, either express or implied or statutory; other than contained in this Agreement, is made in respect to Millar products. The Purchaser therefore assumes all risks as to the performance and reliability of the products, and the results gained using them. Millar is not responsible for any problems with the computer system not directly related to Millar products.

Millar neither assumes nor authorizes any person to assume on its behalf any liability in connection with the sale, installation, service, or use of its products. Millar shall not be held responsible for special, consequential, or punitive damages of any kind arising out of sale, installation, service, or use of its products.

Millar Limited Hardware Warranty

Millar warrants that at the time of sale to the original Purchaser, the MPVS-300/400 series hardware shall be free from defects in materials and workmanship for a period of one (1) year from its date of shipment to the original purchaser. If there is such a defect, Millar will, at no charge and at its option, either repair or replace the equipment as appropriate. Millar's limited warranty does not cover damage to the product from alterations, misuse, abuse, negligence, or accident.

There are no user-serviceable parts inside the cabinet and the Purchaser should make no attempts to service the hardware. There is no need for the Purchaser to open the cabinet for inspection or maintenance, and doing so within the warranty period will void the warranty.

This warranty applies only to the hardware specified in this document and used under normal operating conditions and within specification. It does not cover hardware modified in anyway, subjected to unusual physical, electrical, or environmental stress, used with incorrectly wired or substandard connectors or cables, or with the original identification marks altered. Since factors beyond Millar's control may directly affect the product and the results obtained from its use, Millar or its agents or employees shall not be liable for any incidental or consequential loss, damage, or expense arising directly or indirectly from the use of this product.

The user shall determine the suitability for use of these devices for research purposes only. Therefore, the user accepts these devices subject to all the terms hereof. Furthermore, Millar does not warrant that equipment is suitable for any specific purpose, other than that explicitly stated by Millar.

Millar hereby excludes all warranties not herein stated, whether express or implied by operation of law or course of dealing or trade usage or otherwise, including but not limited to any implied warranties of fitness or merchantability.

Products Supplied But Not Manufactured by Millar

Millar-supplied products of a third party manufacturer, i.e. data acquisition systems, amplifiers, computers, monitors, and printers including associated cables, probes, accessories, etc. are not warranted by Millar. These items are covered by the original manufacturer's warranties. Please contact the manufacturer directly if you encounter a defect with a third-party product. If a fault arises, Millar will help assist in determining which products may be the source of the problem encountered.

Note: If your problem involves computer hardware, back up data from the hard-disk drives(s) and any other storage device(s) in the product and remove any storage media, such as diskettes or CD's.

Warranty Service

To obtain service under the terms of this warranty, the Purchaser must notify Millar or the nearest authorized Millar distributor of the defect before the warranty period expires. The Purchaser must contact Millar or the nearest authorized Millar Distributor to obtain an RMA (Returned Materials Authorization) number and an address to which the Purchaser must ship the defective product at their own expense. The product should be packed safely (preferably in its original packaging) and have the RMA number prominently displayed on the return packaging, preferably beneath the address or shipping label. Millar will make all reasonable efforts to evaluate the product and provide service as necessary within a reasonable amount of time upon receipt of the returned product. Millar will pay the return shipping costs in the event the product is to be sent back to the Purchaser.

Technical Support

The Purchaser is entitled to free technical support for any Millar product. Millar's technical support staff can provide help and advice concerning installation, operation, particular uses, and problems with Millar products. Millar's technical support staff will not provide experimental protocols or procedural instructions for conducting experiments. However, information of this type may be provided in the supplied product documentation, or on the Millar web site.

Technical support can be provided to the Purchaser via the following options:

Phone	832.667.7000
	8:00AM – 5:00PM, Central Time, M-F
Fax	832.667.7015
	24 hours a day, 7 days a week
Email	info@millarmail.com
	24 hours a day, 7 days a week
Web	www.millarinstruments.com

On-site installation and training for Millar products can be provided for an additional charge. Please contact Millar's customer service department for further details.

Jurisdiction

This Agreement shall be governed by the laws of Texas in the United States of America, and any proceedings concerning it shall be heard and resolved in a Texas court of law.