# HAMILT®N

# Real-Time CO<sub>2</sub>NTROL Solid-State Optical CO<sub>2</sub> Sensors



# **IS YOUR PROCESS IN**



# Increase Yield and Consistency

New research shows that active control of dissolved CO<sub>2</sub> in bioproduction increases product yield and improves consistency of process scale-up and scale-down. Traditional electrochemical sensors indirectly measure DCO<sub>2</sub>, are maintenance intensive, and are prone to measurement drift. CO2NTROL is a Solid State Sensor that directly measures DCO<sub>2</sub> and provides maintenance free, real-time, and in-line control of this new critical process parameter.

#### **AUTOMATED CONTROL OF DCO<sub>2</sub> ENABLES:**



Increased

Product Titer



Better Batch-to-Batch Reproducibility

Consistency More Consistency from R&D to Production Scale Reactors

# Bioreactor Size Impacts CO,

Large and small bioreactors have Mass Transfer Coefficients (KLa) that change the dissolution and stripping characteristics from reactor to reactor. As a result the same control strategy will result in different CO<sub>2</sub> accumulation across R&D, PD, and Production scale reactors. Only real-time control of DCO2 to an optimized profile will result in comparable mass transfer independent of scale.





#### **Optimizing Yield**

While the small surface area to volume ratio of a typical R&D reactor means that CO<sub>2</sub> accumulation is minimal. Real-time control of CO<sub>2</sub> is critical to define the optimal setpoint and optimize product yield.

Scale Up

Mimicking sparging and stripping strategies optimized for R&D will result in different conditions as the surface to volume ratio decreases. Active control of an optimal DCO<sub>2</sub> profile ensures consistency across scales.

#### IMPACT OF CO, ON PROCESS PERFORMANCE:

#### Excessive Accumulation

During a process excess CO<sub>2</sub> accumulation is common as cell concentration increases and more metabolic CO2 is produced. This accumulation reduces intracellular pH resulting in slower enzymatic activity or delayed lactate shift. The outcome is lower production quantity and quality.

#### Excessive Removal

Aggressive aeration and mixing can prevent detrimental accumulation of CO2. However, too little CO2 can slow cell growth, metabolism, and productivity by starving cells of CO<sub>2</sub> needed for the formation of metabolic intermediates. Excessive removal can also reduce buffer capacity in bicarbonate systems.

#### Uncontrolled CO<sub>2</sub> range

Media addition and process adjustments can result in large swings in dissolved CO2 that may go unnoticed with infrequent offline monitoring and adjustment. Continuous inline control at the optimal level yields increased viable cell density, production phase duration, and titer.









#### Scale Down

For existing processes that were developed without DCO<sub>2</sub> control, the efficiency of scale-down studies can be improved by actively controlling DCO<sub>2</sub> to mimic the profile seen in the production reactor.

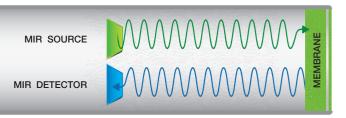


# MAINTENANCE-FREE SOLID-STATE

## Direct vs. Indirect

Traditional electrochemical sensors measure DCO<sub>2</sub> based on the Severinghaus principle. This indirect measurement method combines the challenges of measuring pH and electrochemical DO into one sensor. The result is significant maintenance effort and multiple sources of drift that must be compensated by time-consuming product calibration. Hamilton's CO2NTROL is a maintenance free, solid-state sensor that directly measures DCO<sub>2</sub> resulting in better measurement accuracy and lower cost of ownership.





#### **CO<sub>2</sub> MEASUREMENT PRINCIPLES:**

#### Severinghaus

CO<sub>2</sub> molecules diffuse through a gas permeable membrane into an electrolyte. CO<sub>2</sub> increases the acidity of the electrolyte which is measured by an internal pH sensor.

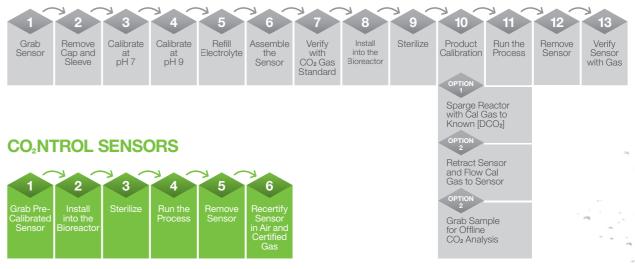
#### **CO<sub>2</sub>NTROL Optical Measurement**

CO<sub>2</sub> molecules diffuse into a gas permeable membrane where the sensor measures the absorption of CO<sub>2</sub>-specific MID IR wavelengths. This absorption correlates to the partial pressure of CO<sub>2</sub> in the media.

# Streamlined Workflow

The CO2NTROL development team recognized that for CO2 to become a critical control parameter the sensor had to be easy to use and maintain. To accomplish this we had to reduce the complexity and eliminate liquid solutions and replacement parts. The team developed a Solid State MIR design that is calibrated in certified gas and does not require a product calibration because of drift after sterilization processes.

#### SEVERINGHAUS SENSORS







Simple Calibration In air and gas standard.



Measuring Range Designed for bioproduction (5-1000 mbar).



Maintenance Free

No liquids or replacement parts to change.



Hygienic Design Compatible with Autoclave, SIP, or CIP.





#### Inverted Installation

Not affected by mounting orientation.



#### No Ammonia Fouling

No interference from other dissolved gases.

# GET CCONTROL

**CO₂NTROL** is the newest member to Hamilton's Arc Intelligent Sensor line. Embedded electronics convert the MIR CO₂ measurement into standard digital and analog signals that are easily integrated into your control strategy.





#### ARCAIR

Seamless control of all Arc products from a PC or mobile device. **CO<sub>2</sub>NTROL** pairs with the newest ArcAir software to automatically store all sensor activities, diagnostic data, and health indicators throughout the life of the sensor. ArcAir is ready for GMP with Part 11 and Annex 11 compliance.

#### INTEGRATED INTELLIGENCE

Arc sensors save space and cost with their integrated micro-transmitters. Calibration data is stored in the sensor head enabling high quality lab calibration.

#### **PROCESS DATA**

The **CO<sub>2</sub>NTROL** sensor transmits process data through a hardwired connection. Communication options include modbus RTU and 4-20 mA, with additional compatibility for OPC, Profibus, and Foundation Fieldbus coming soon.

#### COMPLIANT HYGIENIC DESIGN

CO₂NTROL's hygienic design makes it compliant with requirements of biopharma applications. The sensor is EHEDG certified and is ready for GMP compliance.



#### WIRELESS CONNECTIVITY

An optional Bluetooth adapter and ArcAir software enable wireless calibration, configuration, and monitoring of all sensor details.

### **CO2NTROL SPECIFICATIONS**

Measurement Principle	Optical – CO <sub>2</sub> Absorption in Middle Infrared (MIR)		
Measuring Range	5 to 1000 mbar or 0.5 to 100 %-Vol or 7.5 to 1500 mg/L (in liquid phase at 101.3 kPa and 25°C)		
Diameter	12 mm		
Process Connection	PG 13.5		
Wetted Parts	Stainless Steel 1.4435, EPDM (Ethylene propylene elastomer), FDA compliant silicone		
Surface Quality of Steel	Ra < 0.4 µm (N5)		
Steam Sterilizable	Yes		
Autoclavable	Yes		
CIP	Yes		
Operating Temperature Range	From -10 °C to 60 °C		

### **ORDERING INFORMATION**

	A-LENGTH	REF	
CO₂NTROL	120 mm	10087810-11	
	160 mm	10087810-12	
	225* mm	10087810-13	
	325 mm	10087810-14	
	425 mm	10087810-15	

#### ACCESSORIES

Cables – M12:	Cables – VP8:	Accessories:
Arc USB Power Cable M12 - 8 pole,	Arc USB Power Cable VP8 pole,	Arc Wi 2G BT, REF 243470
REF 243490-02	REF 243490-01	Arc View Mobile Advanced (Tablet), REF 10071113
Open end – 3m, REF 355320	Open end – 5m, REF 355265	Arc Wireless Converter BT, Advanced REF 242333 Calibration Station, REF 243575
Open end – 5m, REF 355321	Open end – 10m, REF 355266	
Open end – 10m, REF 355322	Open end – 20m, REF 355268	

\*CO2NTROL 225 have, in reality, a shaft length of 215 mm. This ensures optimal rinsing in replaceable armatures, such as Retractex.

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