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# **Investor and Analyst Meeting**

June 5-6, 2019

1:00-2:00

Lunch

Julie 5 - Clayton, NC		Julie 5 - Glaytoll, NC		Julie 0 - Kaleigii, NC	
7:30	Pick-up from recommended hotels	2:00-2:45	Novel Plasma Therapies Development  T. Willis	7:30	Pick-up from recommended hotels
8:30-9:00	Registration and welcome	2:45-3:30	AMBAR: Grifols' Alzheimer Trial A. Paez	8:30-9:30	R&D Tour
9:00-9:30	Introduction R. Griffols	3:30-4:00	Break	9:30-10:15	China: Facing the Opportunity  A. Martinez
9:30-11:30	Commercial Strategies  L. Morgan/J. Abelson/ C. Schroeder/R. Jagt	4:00-4:30	Q&A	10:15-10:45	Digital Innovation  X. Sueiras
11:30-12:00	Break	4:30-5:00	Tour Introductions	10:45-11:15	Break
12:00-1:00	Industrial Capacity and Plasma Capabilities  P. Allen/E. Herrero/D. Fleta	5:00-6:30	Site Tour: New Fractionation Building and Ebola plant	11:15-12:00	Financials A. Arroyo

Back to recommended hotels

Dinner

7:00

10:00

June 5 - Clayton NC

June 6 - Raleigh NC

**Grifols: A Socially Responsible** 

T. Rione

V. Grífols Deu

Company

Closing

Q&A

Lunch

12:00-12:30

12:30-12:45

12:45-1:15

1:15

June 5 - Clayton NC



Raimon Grífols Roura Co-CEO

**GRIFOLS** 

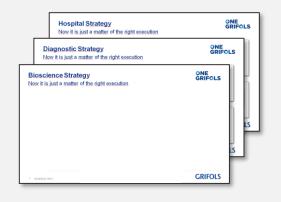
#### A New Chapter of Continued Growth and Success

OVER THE PAST TWO YEARS, EVERY DECISION HAS BEEN A BUILDING BLOCK THAT HAS PAVED OUR WAY INTO THE FUTURE...



...WE BELIEVE EVERYTHING IS NOW IN PLACE, NOW IS A MATTER OF RIGHT EXECUTION...

Key priorities moving forward



...TO ENSURE OUR CONTINUED GROWTH & SUCCESS

Corporate Focus areas





#### A New Chapter of Continued Growth and Success

Past Two Years

OVER THE PAST TWO YEARS, EVERY DECISION HAS BEEN A BUILDING BLOCK THAT HAS PAVED OUR WAY INTO THE FUTURE...





## **Grifols Today**

Over the Past Two Years, Every Decision We Made Has Been a Building Block Paving our Way into the Future



#### ORGANIZATION

- Governance
  - MB/EC
  - · Strategy Board
- Corporate Functions
  - · Strategy Office
  - · Innovation Office
- **Divisions:** Bio Supplies
- Communication Department
- Data Protection Office



#### TALENT

- +7,000 new employees
- Executed a successful succession plan coupled with selected external hiring
- Leadership Day
- Enhanced **compensations & benefits** plans
- Work-life balance measures
- Doubled down training & development



## **Grifols Today**

Over the Past Two Years, Every Decision We Made Has Been a Building Block Paving our Way into the Future





#### **Our Future**

#### Everything Is or Soon Will Be in Place

#### Across

- People & talent
- One Grifols
- China (Shanghai RAAS)

#### Bioscience

- Plasma availability
- AMBAR
- Innovation
- New manufacturing plants

#### Diagnostic

- Wide product portfolio
- Leadership position
- Vertical integration
- New manufacturing plants

#### Hospital

 Expanding product portfolio of compounding control solutions: InclusIV

Now is just a matter of the right execution



#### A New Chapter of Continued Growth and Success

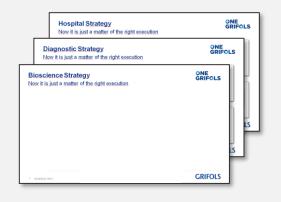
Our Future, a Matter of Having the Right Execution

OVER THE PAST TWO YEARS, EVERY DECISION HAS BEEN A BUILDING BLOCK THAT HAS PAVED OUR WAY INTO THE FUTURE...



...WE BELIEVE EVERYTHING IS NOW IN PLACE, NOW IS A MATTER OF RIGHT EXECUTION...

Key priorities moving forward



...TO ENSURE OUR CONTINUED GROWTH & SUCCESS

Corporate Focus areas





# **Corporate Strategy**

#### There Are a Set of Focus Areas in Which We Need to Focus Moving Forward



#### **One Grifols**

areas

Focus

Operate as one company and leverage on capabilities to unlock synergies



# **Business Optimization**

Identify inefficiencies to improve productivity and optimize value



# **Innovation Performance**

Leverage
technological
advancements to
deliver innovative
solutions and
transformational
breakthroughs



# Customer Centricity

Enhance
organization wide
focus on meeting
and exceeding
customer needs to
build sustainable,
competitive
advantage



#### **Digital**

Build digital
capabilities to
deliver better
outcomes, explore
new areas to play in
and identify new
sources of value



#### **Talent**

Ensure that we have the right talent in the right roles and develop our people to strengthen and expand competencies





Lafmin Morgan
Chief Commercial Officer

**GRIFOLS** 

## **Grifols Today**

Global Presence With a Diversified Revenue Base

# Leading Position in Bioscience with a Growing Position in Diagnostic, Hospital and BioSupplies



A leading producer of essential plasmaderived therapies worldwide



A leader in transfusion medicine from donation to transfusion



Advanced pharmacy specialty products for hospital use



Promotes biological products for non-therapeutic use

Headquartered in Barcelona with more than 22,000 employees in 30 countries

**Grifols Today** 

# **Achieving Lasting Customer Success**



## **Grifols Commercial Is Achieving Lasting Success**

Leadership and Successful Track Record

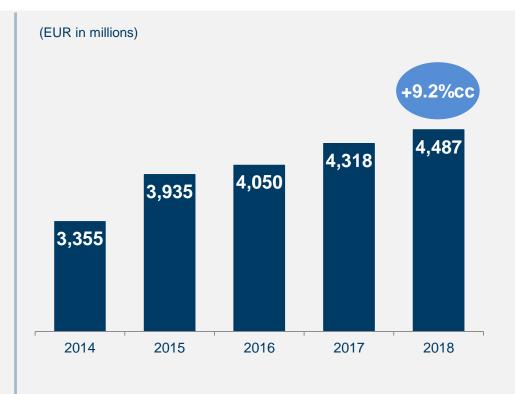
- Talented team with proven ability to execute and overcome obstacles
- Strong market fundamentals across business lines
- Long-term benefit accruing from One Grifols customer focus
- Planned launches building on strong foundation for future growth
- Continued growth through geographical expansion

#### **Successful Track Record**

Demonstrated Ability to Successfully Build on Growth

# Grifols has generated growth by offering timely, relevant solutions to customers

- Expanding our customer base
- Increasing customer trial
- Retaining existing customers
- Introducing new innovative products and solutions



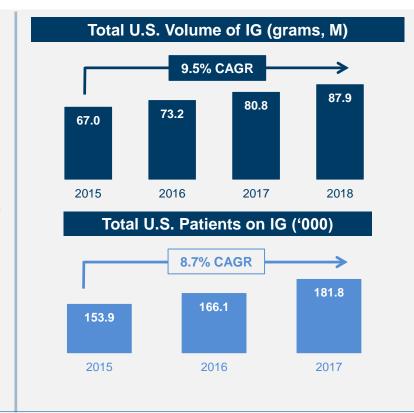


## Immunoglobulins | Patient Growth Fuels Expansion

Growth is Consistent Across Therapeutic Uses and Geographies

# Growing Awareness, Diagnosis & Treatment Therapeutic uses showing the highest U.S. IG volume growth were driven by:

- PIDD: Expanding awareness & discovery of new sub-indications
- CIDP: Increased diagnosis rates & preference for IG as a first-line therapy
- SID: Expanding immune-modulator use in hematology-oncology patients



Source: U.S. PPTA Distribution Data & Data on File PIDD: Primary Immune Deficiency Disorder CIDP: Chronic Inflammatory Demyelinating Polyneuropathy

SID: Secondary Immune Deficiency (not an FDA approved indication in the U.S.)



#### Immunoglobulins | CIDP Indication

Opportunity to Improve Dosing

Results: 72% of CIDP patients received fewer grams than recommended as per the ICE trial. On average, CIDP patients receive 45% of the grams recommended (587.9/1,300 grams)

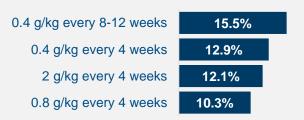


■ Prescribed FEWER g than ICE trial ■ Prescribed per ICE trial ■ Prescribed MORE g than ICE trial

# Actual vs. Ideal IG Prescribing for CIDP Patients

CIDP	Mean IVIG Total Dose Per Patient /Year		
Actual Dosing	587.9		
Ideal Dosing per ICE Trial	1,300		
Proportion of Ideal Dose	45%		

# Most Common Maintenance IVIG Dosing Regimens



Source: ICE trial: https://www.thelancet.com/journals/laneur/article/PIIS1474-4422(07)70329-0/fulltext



#### Alpha-1 | Making a Significant Change in the Course of a Serious Disease

Improving Health Also Represents a Significant Opportunity



**8.3** years

**Average interval** between onset of pulmonary **symptoms and diagnosis** 



2.7
physicians

**Average number of physicians** seen by patients before diagnosis

The longer AAT deficiency remains undiagnosed, the greater the risk for irreparable lung damage

Source: Campos et al., Trends in the Diagnosis of Symptomatic Patients With AATD Between 1968 and 2003, Chest, 2006



#### Alpha-1 | Making a Significant Change in the Course of a Serious Disease

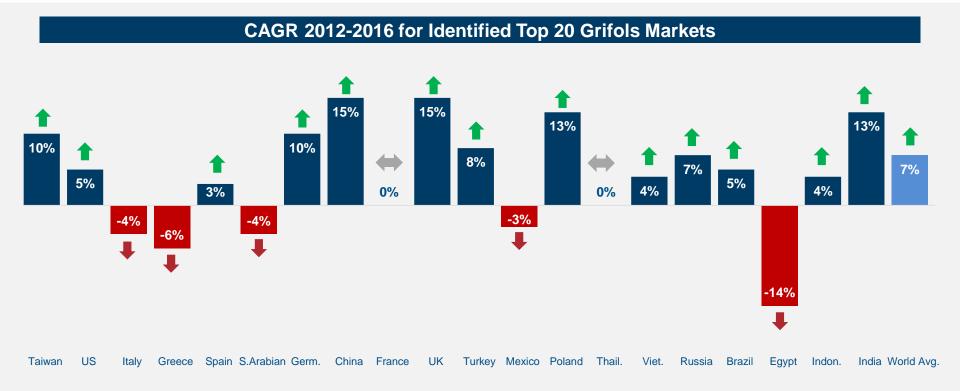
Patient Diagnosis Continues to Drive Growth





# Albumin | Healthcare Access Trends Continue to Support Growth

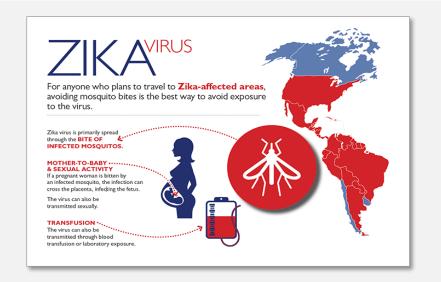
Per Capita Consumption Continues to Grow Consistently Across Grifols Markets

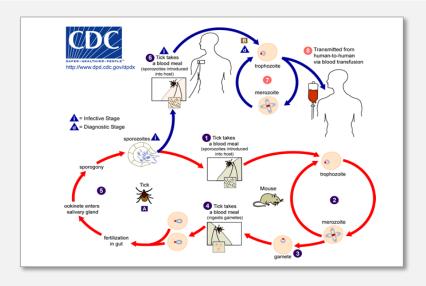


Source: Grifols internal data

#### **Grifols Diagnostic | Ensuring the Safety of the Blood Supply**

Innovating to Address Present and Future Public Health Risks





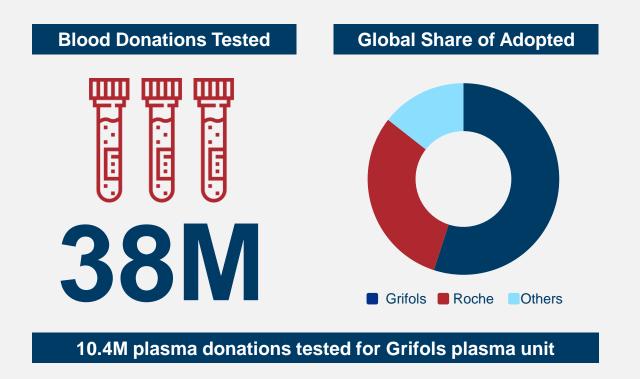
At first opportunity, Grifols R&D team began developing **Zika Virus & Babesiosis** blood screening assays to prevent transmission in the blood supply:

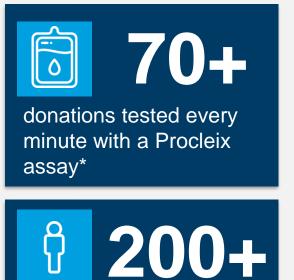
- Grifols **Zika assay** went into use less than 6 months after the epidemic began
- Grifols Babesia assay was approved for use by FDA in February 2019



# Grifols Diagnostic | The Global Leader in Blood Donor Screening

Global Expansion and Plasma Drive Growth





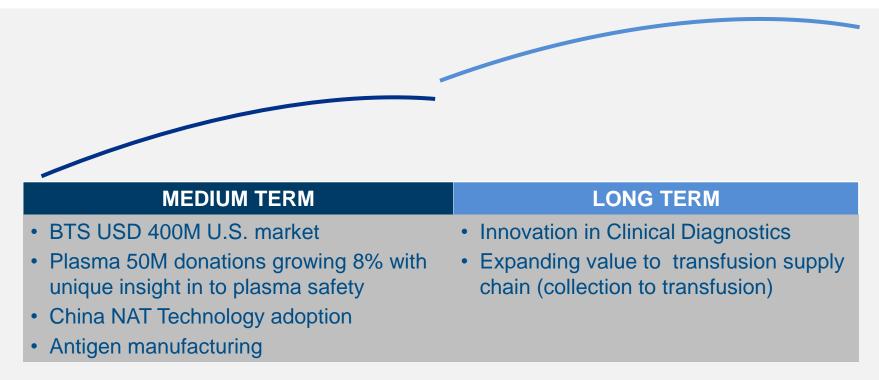
Source: Internal Data. \* Does not include plasma collection

people impacted

every minute\*

# **Grifols Diagnostic | Capturing New Diagnostic Opportunities**

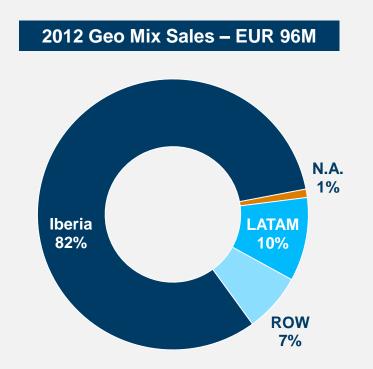
The Stage Is Set for Renewed Growth

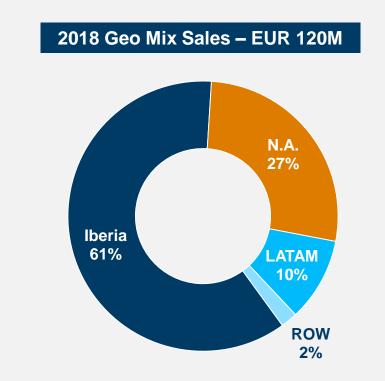




## **Grifols Hospital | Transformed by Strategy Execution**

A Robust Strategy Dynamically Positions the Division

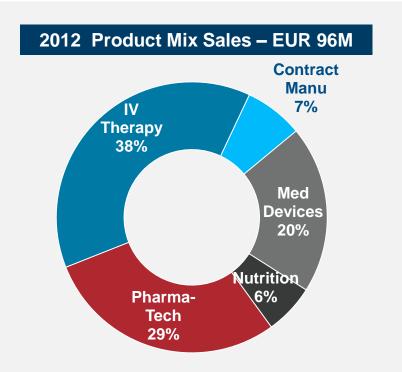


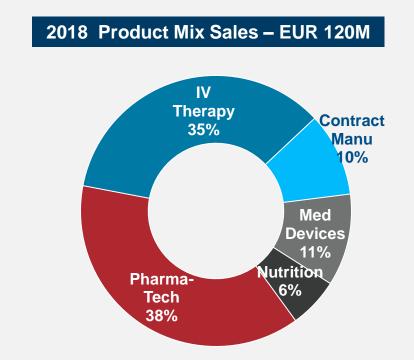




## **Grifols Hospital | Transformed by Strategy Execution**

A Robust Strategy Dynamically Positions the Division







## **Grifols' Commercial Leadership Team**

#### **Experienced Dedicated Leadership**



**Joel Abelson** 

President, Bioscience Commercial Division



**Carsten Schroeder** 

President, Diagnostics Commercial Division



**Rob Jagt** 

President, Hospital Commercial Division

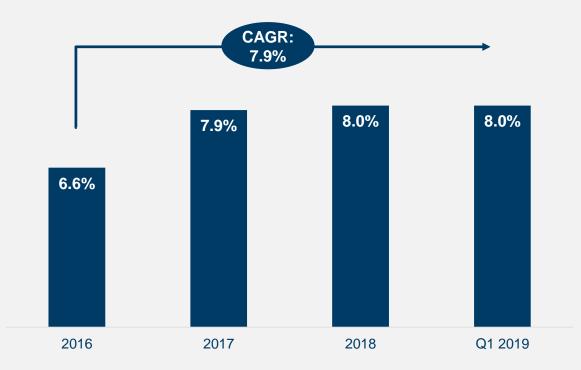
# **Bioscience Division Expanding Grifols' Impact and Delivering Growth**

Joel Abelson
President, Bioscience Commercial Division

**GRIFOLS** 

### **Delivering Continued Sales Growth**

Bioscience Revenue Has Grown at 7.9% CAGR at CC Since 2016



Year-over-year variance as reported in constant currency (CC)

#### **Bioscience Growth Fundamentals Remain Strong**

Leading Position Within Core Business of Plasma-Derived Therapies in 2018

	Global Market Share*	Grifols Global Position	U.S. Market Share*	Grifols U.S. Position
IVIG	25%	#1	35%	#1
Alpha-1	67%	#1	66%	#1
pdFVIII**	17%	#1	45%	#1
Albumin	15%	#2	33%	#2

Per capita utilization and diagnosis are growing for IVIG, albumin and alpha-1

Market growth and expansion strategies continue to deliver results

Grifols continues to invest in the Bioscience Division to sustain growth

\* Market shares in revenue \*\* vWF not included

Source: Grifols Global Plasma Database, Provisional Data 2018



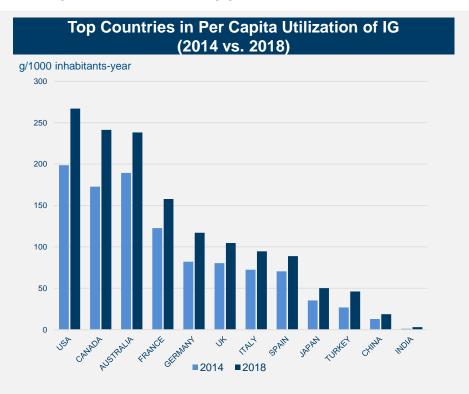
# **Grifols Immunoglobulin**

**Leading Market Growth and Meeting Patient Needs** 



#### Global Demand for IG Continues to Accelerate

#### Multiple Factors Support Continued Growth in Immunoglobulin



#### **Key Factors Driving Growth:**

- Demographic trends: pop. growth, aging populations
- Increased physician diagnosis and treatment of CIDP, PIDD
- Increased need to treat SID due to more aggressive therapies

#### **Top Countries Growth Continues in 2018:**

• U.S.: +9%

Germany: +12%

Spain: +9%

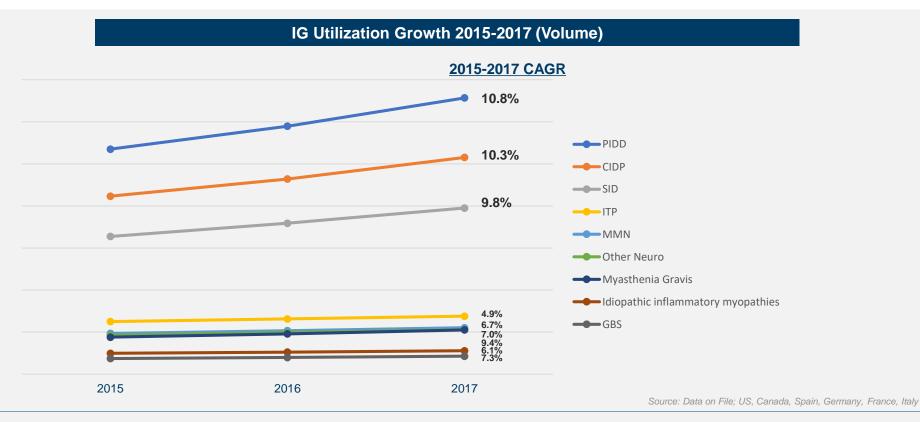
Australia: +10%

Our plasma investments enable us to deliver a consistent supply to our patients

Source: Grifols Global Plasma Database & Marketing Research Bureau Provisional Data, 2018

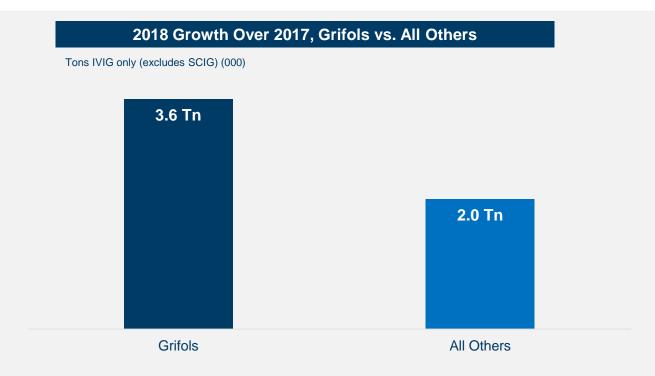
#### **Diagnosis and Treatment Drive Global IG Utilization**

CIDP, PIDD and SID Are Leading Growth



#### Grifols Led Industry Response to U.S. IVIG Demand in 2018

Grifols Accounted for 64% of All YoY U.S. IVIG Growth

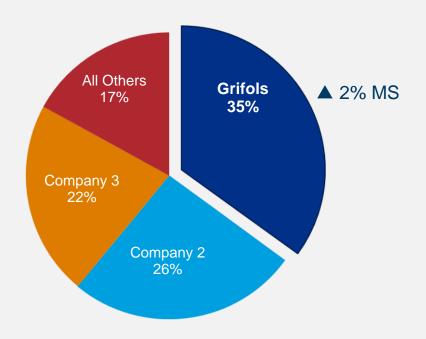


Source: Grifols Global Plasma Database & Marketing Research Bureau Provisional Data, 2018

#### Grifols Increased U.S. IVIG Market Share in 2018

Increased Two Points to 35% in the U.S. in 2018

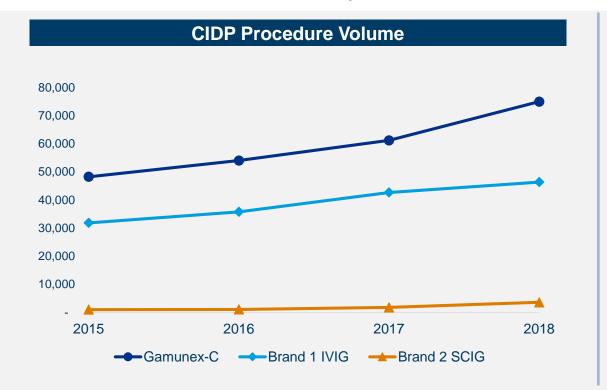
#### 2018 U.S. IVIG Market Share (Volume)



Source: Grifols Global Plasma Database & Marketing Research Bureau Provisional Data, 2018

#### Gamunex®-C Continues to Drive the U.S. CIDP Market

Grifols Grew CIDP Procedures by 23% in the U.S. in 2018, Despite New SCIG Entrant



- 23% Gamunex®-C growth in CIDP procedures (YoY)
- Gamunex®-C grew outpacing market by 6% (market grew at 17%)
- Despite competitors receiving approval for CIDP

Source: Lexis-Nexis, Medical claims data only; Gamunex® -C data includes GammaKed® due to shared J-code

## Gamunex®-C Continues as the Recognized Leader in CIDP

Two-Pronged Approach Focuses on Time to Diagnosis and Our Differentiation

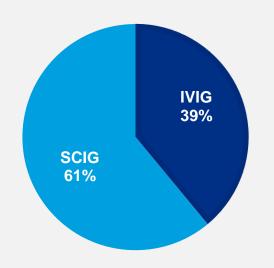




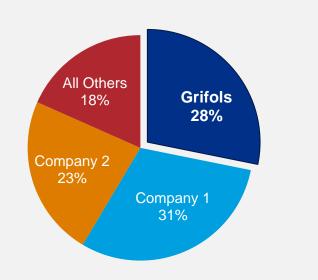
#### Grifols Is Well-Positioned in the U.S. PIDD Market With IVIG

We Are Preparing to Enter the Growing SCIG Market in 2019





# U.S. IVIG Market Share for PIDD Procedures (2018)



Source: Internal Grifols Estimates

Source: Lexis-Nexis, Medical claims data only; Gamunex-C data includes GammaKed® due to shared J-code



## Introducing Xembify™, a New 20% SCIG Option for U.S. Patients

Expanding Our U.S. IG Portfolio to Meet the Large, Unmet Medical Need in PIDD





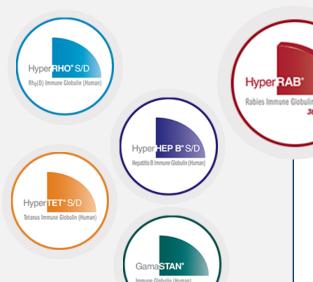




- New 20% SCIG option for patients in search of an alternative treatment
- Continued large unmet need to diagnose and treat patients with PIDD
- Anticipate Q3 2019 FDA approval
- Launch preparation underway

## Grifols Successfully Launched New HyperRAB® in 2018

First Advancement in Rabies IG Administration in 40+ Years





HyperRAB® is the #1 prescribed rabies IG in the U.S. with a market share of 85-90%

#### Grifols launches higher potency formulation (300 IU/mL):

- 2x the rabies antibodies delivered at wound site (compared to existing products)
- Treatment with 50% the volume in a total dose
- Fewer injection sites

#### Feb. 2018 FDA New Formulation Approval. May 2018 Launch:

- 100% conversion to selling new formulation
- 90%+ awareness of new formulation among treaters
- New competitor restricted to minimal market share (6-7%)
- Nearly 80% of previous customers have already transitioned to new formulation

#### Grifols Td vaccine

Tetanus and Diphtheria Toxoids Adsorbed



#### **Key Takeaways**

#### Immunoglobulin: Leading Market Growth and Meeting Patient Needs

- Multiple drivers support continued Immunoglobulin market growth
- In 2018 Grifols increased IVIG share in both the U.S. and EU
- Our recent significant plasma investments have allowed us to ensure a continuous supply of IG to our patients
- Grifols led industry in increasing U.S. IVIG volume in 2018, delivering 64% of IVIG growth
- Gamunex® -C experienced strong growth in CIDP procedures at 23% in 2018 in the U.S.
- Grifols well positioned in U.S. PIDD market with IVIG preparing to enter SCIG market
  - Xembify<sup>™</sup>, our 20% SCIG treatment, license expected Q3 2019
- Grifols retained 90% of the rabies market after new competitor launch in mid-2018



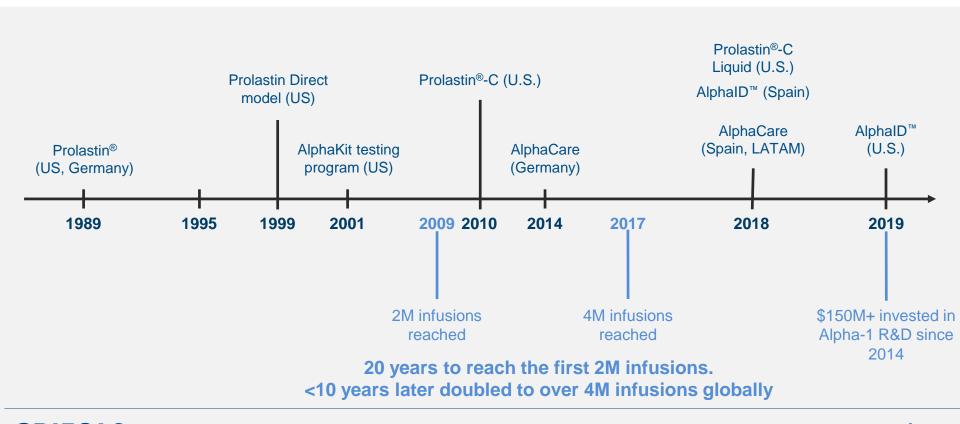
## **Grifols Alpha-1 Antitrypsin**

**Accelerating Growth Through Strategic Investments** 



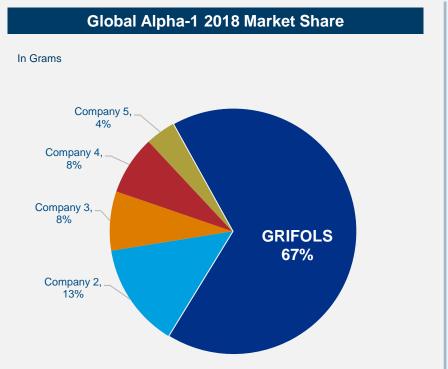
#### **Over 25 Years of Commitment to Alpha-1 Patients**

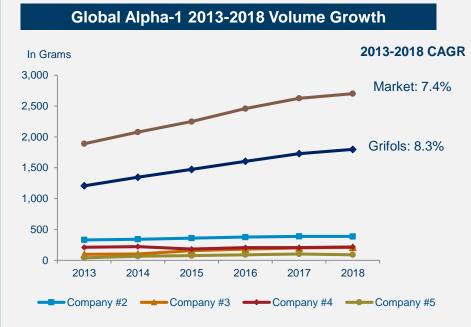
Grifols Has Led the Market With Innovations in Diagnosis and Treatment Since 1988



#### **Grifols Is the Global Market Leader in Alpha-1**

Grifols Alpha-1 Volume Growth Has Consistently Outpaced the Market

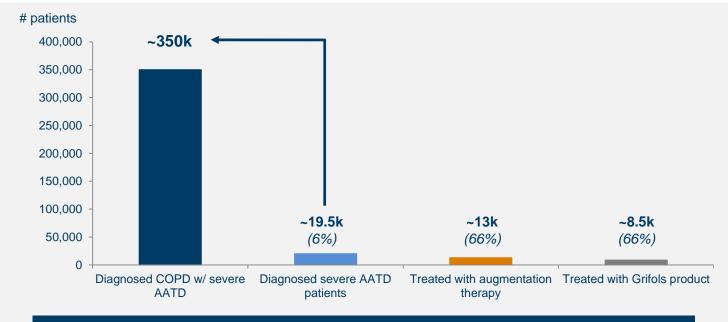




Source: Grifols Global Plasma Database & Marketing Research Bureau Provisional Data, 2018

## **Global Alpha-1 Growth Opportunity Remains Significant**

Less than 10% of WW Patients With Severe Alpha-1 Deficiency Have Been Diagnosed



Significant opportunity still remains to increase diagnosis of AATD patients

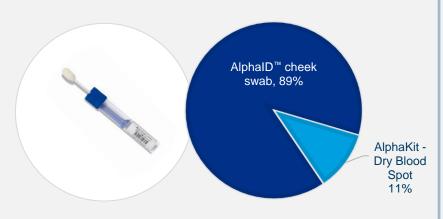
Sources and assumptions: Grifols patients based in 1Q19 patient counts (last update 24APR2019); assume Grifols holds 66% of the total patients based in Grifols BI database (see market overview slide); assume 2/3 of patients diagnosed receive treatment based in market knowledge and affiliate input.



#### Grifols Continues to Innovate in the Diagnosis of Alpha-1

New Diagnostic Test Offers More Convenience: Launched in Spain, U.S. to Follow

# Spanish Testing Program (2019) Physician Test Kit Usage When Given a Choice



#### **Grifols Alpha-1 Testing Program\***

# Grifols continues to innovate via the newly launched AlphalD cheek swab

- Improved testing convenience compared to the dried blood spot test
- Presents an opportunity to further expand into the PCP market

# Based on the success in Spain we plan to launch the buccal swab in the U.S. in 2019

Pilot begins mid-year will a full launch by year-end

\*Testing Technology from Progenika Biopharma, a Grifols Company



#### **Grifols Differentiates in Patient Treatment and Support**

Once Diagnosed, We Have Novel Treatment Options and Support Programs for Patients

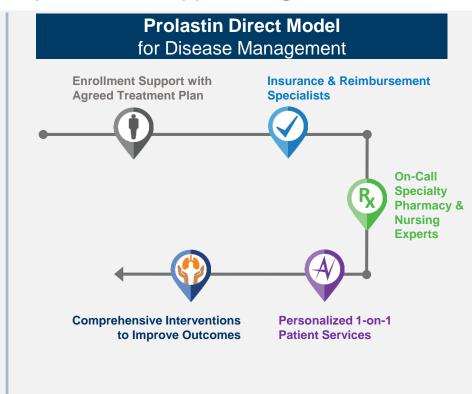
#### U.S. Liquid Launch (July 2018)

 Successful Execution: 85% of patients successfully converted to Prolastin®-C Liquid

#### **Disease Management Programs**

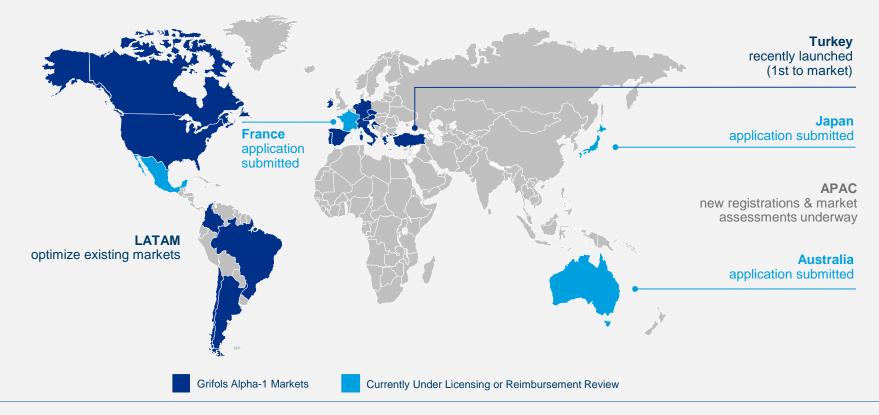
- U.S. Prolastin Direct program has over 95% treatment regimen adherence
- Strong patient participation in global Prolastin disease management programs
  - U.S. (Prolasin Direct)
  - Germany & Spain (AlphaCare)

**EU Nursing Program Launches**: New home infusion programs launched in Germany (H1 2019) and Italy (H2 2019)



#### **Expanding to New Geographies**

#### Untapped Markets Represent Opportunities for Continued Growth





#### **Key Takeaways**

#### Alpha-1: Accelerating Growth Through Strategic Investments

- Prolastin®/Prolastin®-C continues to be the global market leader in Alpha-1
  - Outgrowing the market from 2013-2018 (8.3% CAGR)
- Significant opportunities remain to identify and treat additional patients, with 90%+ of the global market still undiagnosed
- Innovations in product development, diagnosis, education, home infusion and direct-toconsumer initiatives will further differentiate and position Grifols for continued leadership
- The new AlphaID<sup>TM</sup> cheek swab test offers a faster, more convenient diagnostic test for patients and physicians
  - 2018 launched in Spain and 2019 in U.S.

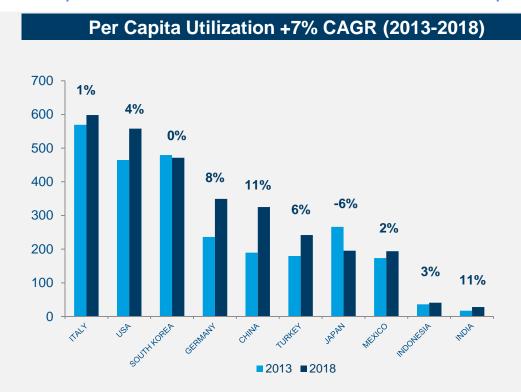
## **Grifols Albumin**

A Key Driver of Bioscience Growth



#### **Global Albumin Market Continues Strong Growth**

Per Capita Utilization of Albumin Grew 7% in Top 10 Countries, Primarily Driven by China

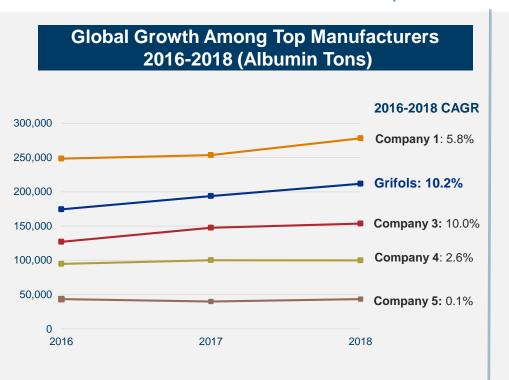


- Global Albumin market grew in volume (grams) at 7% CAGR from 2013-2018
- Per Capita Utilization in the top 10 countries matched global market growth at 7% CAGR over the same period
- Italy and U.S. led in terms of overall utilization, with China and Germany leading in growth
- Consumption continues to increase across nearly all top countries

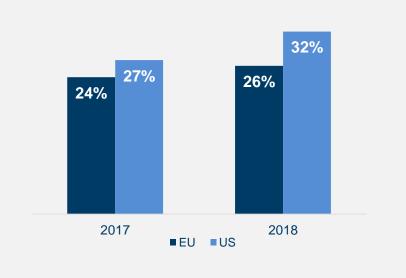
Source: Grifols Global Plasma Database & Marketing Research Bureau Provisional Data, 2018

## **Grifols' Global Albumin Growth Outpaces Competitors**

Grifols Holds #2 Global Position, Captured Market Share in U.S. and EU in 2018



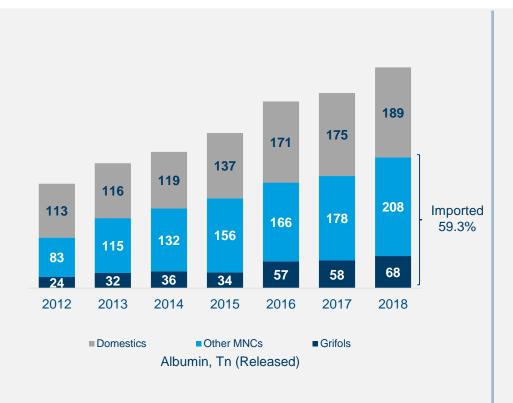




Source: Grifols Global Plasma Database, Provisional Data 2018

#### **Grifols Continues Robust Upward Momentum in China**

Grifols Expansion Strategy in World's Largest Albumin Market



#### 2012-18 Official Release CAGR:

- China Albumin: +13.3%
- Domestic: +8.9%
- Imported (incl. GRF): +17.2%

#### **2012-2018 Grifols CAGR +19.0%:**

- 15% Market Share in 2018 (+1% YoY)
- 24.6% Imported Market Share in 2018

# Grifols expansion strategy in China is delivering positive results

Focus on providing increased supply to Top 11 provinces and driving penetration in the retail sector:

- Top 11 provinces overall sales growth of 32.9% (2016-18)
- Retail channel sales growth of +140% (2016-18)

Source: Institutes of Food and Drug Control Batch Released in China 2012-2018 CAGR



#### **Opportunities in New Indications and Utilization Methods**

Grifols Leads in Albumin Clinical Research Investment (PRECIOSA, APACHE, AMBAR)

	Population, Use	Study Name	Innovation
Liver Cirrhosis	Moderate Cirrhotic Patients Long-Term Albumin Use	ANSWER PRECIOSA*	Increased Utilization New Indication
	Severe Cirrhotic Patients Acute Albumin Use	APACHE*	New Indication
Sepsis	Septic Shock Patients Long-Tem Albumin Use	ARISS ALBIOS 2	Increased Utilization Increased Utilization
Alzheimer's Disease	Mild & Moderate Alz Patients Long-Tem Albumin Use	AMBAR*	Non-Specialty Treatment Protocol

#### **ANSWER STUDY**

April 2017 Published Results

Long-term Albumin use in moderate cirrhotic patients reduces mortality by 38%

At peak, 24 additional tons of albumin would be needed to treat chronic cirrhosis patients with Albumin in EU4\*\* as per Answer Study protocol

\* Grifols-sponsored clinical programs
\*\* EU4 includes Spain, Germany, Italy, UK



## Grifols Albutein® FlexBag 25% Approved in the U.S.

New Flexible Container Designed to Enhance Customer Experience



# Flexible container compliments vials to broaden Grifols offering to customers

- Improved flexible container with overwrap in two sizes
- Launch preparation underway

#### **Key Takeaways**

#### Albumin: A Key Driver of Bioscience Growth

- Grifols holds the #2 global Albumin position, leading growth among top 5 manufacturers (2016-2018)
- Grifols outperformed competitors in 2018 in the U.S., China and Europe
- We believe the Albumin market will continue to grow significant opportunity exists with new indications for long-term/chronic use of Albumin
  - Liver Cirrhosis (including long-term Albumin use) and Sepsis will drive increased consumption and fuel future growth
  - Investing in untapped potential for Albumin that will drive further growth
- Grifols Albutein<sup>®</sup> FlexBag 25% will enhance customer experience



# Grifols pdFVIII/VWF

The Key Role of pdFVIII in Bleeding Disorders



## Paradigm Shift in pdFVIII Continues, Business Impact Absorbed

Grifols Has Weathered the Business Challenge, Growth Opportunities Remain

- A changing market: there is a new treatment paradigm in Hemophilia since the Hemlibra® launch in Q4 2017
- In spite of pdFVIII decline, the Bioscience business continues to grow at ~8% CAGR
- We believe pdFVIII has a pivotal role in mature Hemophilia A markets along with a large, unmet need in emerging markets

#### Patients in Mature Markets Continue to Rely on pdFVIII Therapy

Grifols pdFVIII/VWF Plays a Key Role in the New Therapeutic Environment

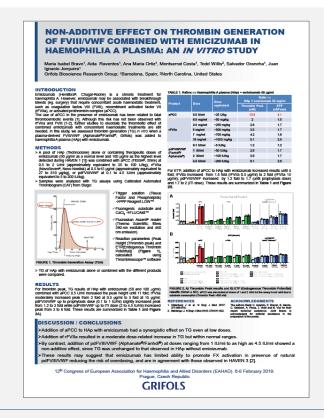
In mature Hemophilia A markets there continues to be an ongoing need for pdFVIII therapy in bleed/surgery management and the prevention/eradication of inhibitors

**Eradication of Inhibitors**: Grifols R&D provided evidence that pdFVIII/VWF can be safely used in combination with new therapies

- Episodic treatment (management of bleeds, in surgery)
- Eradication of inhibitors

#### **Patients Requiring/Preferring Plasma-Derived Therapies:**

- Patients satisfied with traditional prophylaxis or those requiring on-demand treatment
- Treatment of von Willebrand disease (these patients can only be treated with factor therapies)



## **Underlying Demand Is Driving Growth in Emerging Markets**

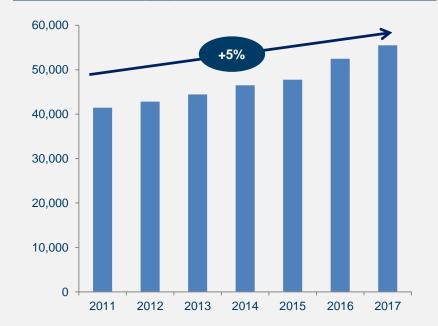
Opportunity to Build on Our Leadership Position to Meet Patient Needs

A large population of underdiagnosed and untreated hemophilia patients exists within emerging markets

Grifols in emerging markets: Focus on improving standards of care and access to treatment together with geographical expansion

- Clinical trial to support low-dose prophylaxis in Indonesia
- Partnerships to accelerate care in Indonesia, India, Philippines via centers of excellence, support in diagnosis, etc.
- Participation in new tenders and further expansion into the Middle East and North Africa

# Evolution of diagnosed HA patients in emerging markets, 2011-2017



Source: World Federation of Hemophilia, 2011-2017



#### **Key Takeaways**

pdFVIII/VWF: The Key Role of pdFVIII in Bleeding Disorders

- Hemophilia experts foresee an ongoing need for the prevention and eradication of inhibitors, bleed/surgery management and VWD treatment with pdFVIII/VWF
- Grifols research shows that pdFVIII/VWF can be safely used in combination with new therapies
- Additional opportunities for pdFVIII/VWF come from emerging markets focusing on access to treatment

# **Key Takeaways**



#### **Bioscience Growth Fundamentals Remain Strong**

#### Product Strategies Will Deliver Continued Growth

**Immunoglobulin Albumin** Alpha-1 pdFVIII

- Grifols is the global & U.S. IVIG market leader, delivering 64% of all U.S. IVIG volume growth in 2018
- Our robust plasma investments support a continuous IG supply to patients
- Xembify<sup>™</sup>, our 20% SCIG treatment for U.S. PIDD patients, is expected to be licensed Q3 2019
- Grifols holds #2 global Albumin position, leading growth among top 5 manufacturers (2016-18)
- Albumin market growth opportunity exists with new indications for long-term/chronic use
- Grifols Albutein® FlexBag 25% to be launched in the U.S.
- Grifols Alpha-1 franchise continues as global market leader, outgrowing the market (2013-18)
- The new AlphaID™ cheek swab test offers more convenience launches in Spain & U.S.
- Hemophilia experts foresee ongoing need for the prevention and eradication of inhibitors, bleed/surgery management and VWD treatment with pdFVIII/VWF
- Additional growth will come from emerging markets focusing on access to treatment

# **Diagnostics Division Global Leader in Transfusion Medicine**

Carsten Schroeder
President, Diagnostic Commercial Division

**GRIFOLS** 

#### A Global Leader in Transfusion Medicine

#### During 2018 We Have Sales in ~100 Countries

The Diagnostic
Division is a global
organization

#### AT-A-GLANCE

- 1,450+ full-time employees supporting Diagnostic success
- Integrated from assay/ instrumentation development through commercialization
- FDA, GMP & CE Licenses



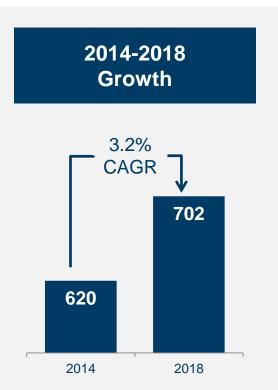


## The Diagnostic Division Reached EUR 702M in Sales in 2018

Sustainable Mid-Single Digit Growth Over the Last 5 Years



**Grifols is the 15<sup>th</sup> largest Diagnostic company** 



Grifols' Largest Customers



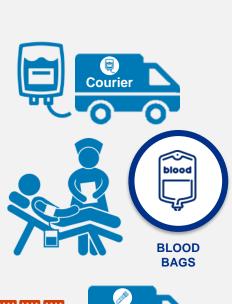




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## The Journey of Blood

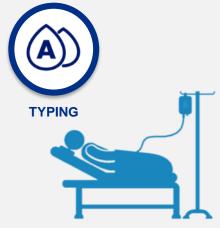
Grifols Plays a Vital Role Ensuring the Safety of the Blood Supply

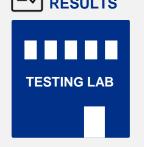












**TEST** 

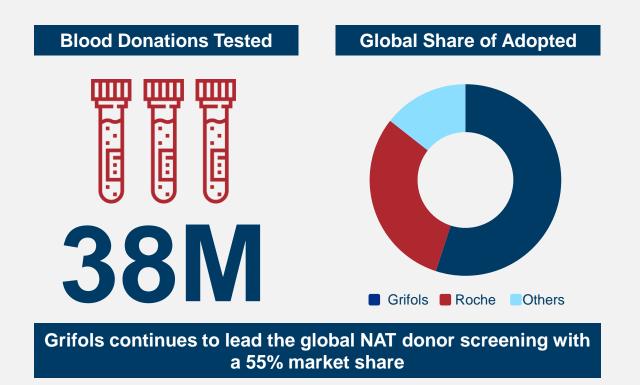


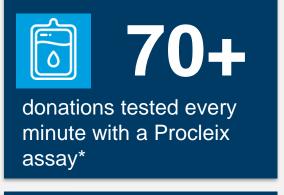




#### **Global Leader in Blood Donor Screening**

Stable NAT Adoption Worldwide Enables Grifols to Grow Around 2% in 2018







Source: Internal Data. \* Does not include plasma collection



## **Top 6 Customers Account for ~21M Donations**

Multi-Year Agreements in Place With All Six Customers

























Creative Testing Solutions, Japanese Red Cross Society and Australian Red Cross Blood Service and logos are registered trademarks of their respective owners.

#### **Responding to Customer Needs**

Successful Product Development Leveraging Vertical Integration







Emerging Pathogens & Panther and Procleix Assays received FDA approval

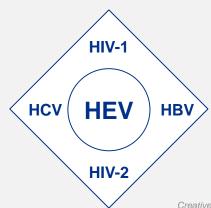












Automation Ready Technology



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#### **Automation Project Is Approaching Market Launch**

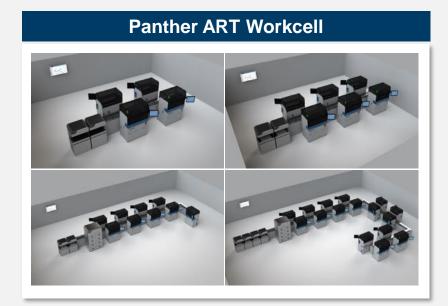
Modular Design to Address Specific Customer Needs and Variable Workflows



# A modified Panther designed to:

- Connect to a network of Panthers
- Connect to track transport system
- Includes waste drain, MTU capacity, other software & hardware modifications

Panther ART is designed to interface with any CLSI-compliant track system



- Track-based sample transport
- Up to 16 Panthers



## **Panther Automation Is Today a Reality**

Functional Track-Based Demo System Installed in Emeryville





### Middleware and Panther Dashboard

Visual Tools That Manage and Prioritize Work in the Laboratory



Focus
operations on
what needs to
be done in the
next minutes,
next hours, and
next shift



## **Above Market Growth Driven by New Products**

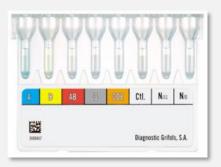
Record Number of Gel Cards Sold

### Blood Typing



9.3% Growth (cc) vs 2017

### **Gel Cards**



**42M** 

Units sold

### **New Distributor**





Runda Medical and logos are registered trademarks of their respective owners.

## Successful New Erytra Eflexis® Launch

250+ Customer Placements Since End Q2 2017

Instrument Placements

Competitive Conversions

250+

Since launch in Q2 2017

+50%

Countries

U.S. Launch

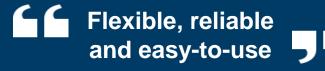


36
Countries

20+

Since FDA approval in Q1 2019





M.Maresca (Gemelli Policlinico, Rome, Italy)



## **U.S. Immunohematology – Over 200 Customer Sites Under Contract**

Launch of New Products Will Continue to Fuel Grifols' Growth

**50%** Growth (cc) vs 2017

## **New FDA Approved**





## Leverage Manufacturing Facilities in Spain and Brazil

Strengthen Our Position in LATAM and Expansion Plans in EMEA

### **Blood Collection**



13.4%

Growth (cc) vs 2017



### Leucored RC T&B, new Asahi Filter

We produce high-quality blood collection bags for collecting and processing whole blood and storing blood components





MURCIA SPAIN



### **Key initiatives**

- Take full advantage of manufacturing facility in Brazil
- Re-launching in EMEA with a soft filter product

## **Leader in Antigen Supply for Immunoassays**

Worldwide Market Leader in Hep/retro Immunoassays Antigens\*



- · CMF manufacturing site
- HCr43 and HBCore antigens

134,759

Antigen shipped (milligrams)

### **Future Growth Drivers**

- Continuous expansion of antigens portfolio
- Exploring contract manufacturing opportunities outside of the agreement



\* In Vitro Diagnostic Market Segment Review 2013-2014 and 2019 Forecast Ad hoc report from Boston Biomedical Consultants, Inc., 2015 and internal estimations

## **Expanding the Possibilities of the New Emeryville Facility**

Center of Excellence for Recombinant Protein Design and Development

### **Protein Expression Platforms**

#### **BACTERIA**



**YEAST** 

 Good for proteins not requiring post-translational modifications

- New vectors in design and development
- Good for complex protein production
- Workhorse for legacy antigen expression

## MAMMALIAN CELLS



 Excellent for expression of glycoproteins

Enables post-translational modifications

### **Robust Immuno-Reagent Pipeline**

### **Donor Screening**

- New or improved antigens and mAbs (i.e. HIV, HCV, HBV)
- New antigens and mAbs for additional pathogens (Zika, Dengue, Babesia, etc.)

### **Hemostasis**

- Novel vWF receptor derivatives
- · Recombinant tissue factor
- Proprietary mAb for improved thrombosis assay

### **Immunohematology**

- Novel rare blood group antigens (stable reagents for extended blood typing menu)
- Fc fusion blocking protein (to resolve interference of daratumumab in antiglobulin testing)

### **Neurodegenerative**

New antigens and mAbs to support Alkahest's drug discovery process in aging related diseases



## The Journey of Plasma

Contributing to the Safety of the Plasma Supply













**TEST** 







## **Plasma Offers Growth Opportunity**

Leveraging on Grifols' Plasma Testing

**Volume Tested** 

**Grifols' Testing** 

Upcoming Transitions

**New Org Chart** 



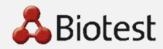
**Donations** 

Tigris to Panther transition EU in P96 FDA approved









Dedicated sales & marketing team



Converting accounts to Procleix and increasing the number of tested donations

## Strategic Alliance in China: Shanghai RAAS

Long-Term Growth Opportunities for Diagnostic

Fast Growing IVD Market

10.9%

CAGR (2016-2021)

New NAT
Blood Donations

+2.5M

(From 2018 to 2020)

Great IH
Opportunity

\$310M

**IH IVD Testing Market** 

NAT Mandate
Plasma Donations

+15.5M

New plasma donations tested



Sources: NIFDC 2018; InterChina survey 2017

### A Global Leader in Transfusion Medicine

Building a Specialty Diagnostics Portfolio

### **Transfusion Medicine**









## **Clinical Diagnostics**



Donor Screening

Global leader in donor screening

**Immunoassays** 

New manufacturing in Emeryville

Blood Typing space

Complete portfolio of instruments, gel cards and reagents Blood Collection

Manufacturing in Spain and Brazil

**Hemostasis** 

Distribution agreement with Beckman

Specialty Diagnostics

Aesku

Promonitor

## **New Organization for Grifols Specialty Diagnostics**

Provide Focus for Future Growth, Following Strategic Plan

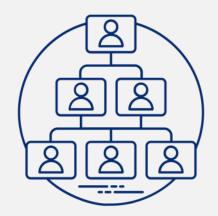
**Specialty Diagnostics** 



3.5%

Growth (cc) vs 2017

**New Org Chart** 



**Promonitor** 



15%

Growth (cc) vs 2017

## We Have a Bright and Exciting Future

Grifols Diagnostic Has Multiple Growth Opportunities



### Global Leader in Transfusion Medicine

Blood & Plasma Donor Screening

Antigen Manufacturing Blood Typing Solutions



# **Growing Loyal Customer Base**

Top 6 NAT Customers
BTS Competitive
Conversions

Sales in 100+ Countries



# Manufacturing Excellence & R&D

Excellent GMP
Facilities
Increasing vertical
integration

Innovative R&D



## Bright & Exciting Future

Emerging Pathogens
Geographical
Expansion, esp. China
Robust pipeline of new
products



# **Hospital Division**

Successful Execution of a Transformational Strategy

**Robert Jagt** 

**President, Hospital Commercial Division** 





## Successful Execution of a Transformational Strategy: Sales Inflection

A Robust Strategy Dynamically Positions the Division

The Hospital Division will continue its strategic evolution to become a comprehensive solutions provider for the Operational Pharmacy - enabling the division to become meaningful contributor to the GRIFOLS group

In 2017 we embarked on a journey to become a comprehensive solutions provider for Operational Pharmacy

Transformational investments in Kiro and MedKeeper to create GRIFOLS COMPLETE have changed the vision and strategic focus for this division

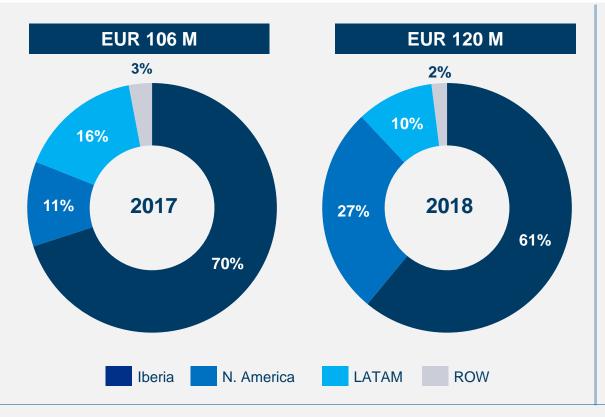
### **Accelerated Performance:**

- Double-digit growth in 2018, following low growth in prior years
- · Growth coming from the U.S.
- Fueled by IV Compounding Portfolio and IV Fluids



## Clear Path to Strengthening Portfolio for Growth

U.S. and Pharmatech Drove 16% cc Growth in 2018



- Growth across all lines
- Dramatic increase in N. America

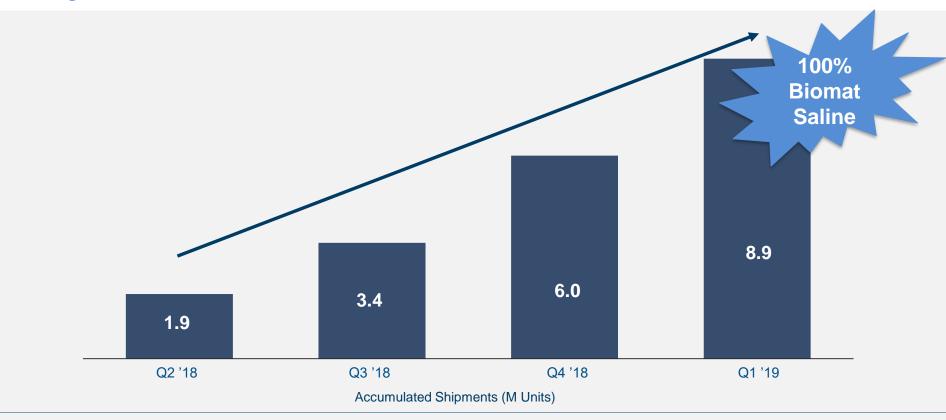
### **Key events:**

- MedKeeper acquisition
- Kiro stake increased to 90%
- Saline launch in U.S.



## **Strengthening Vertical Integration**

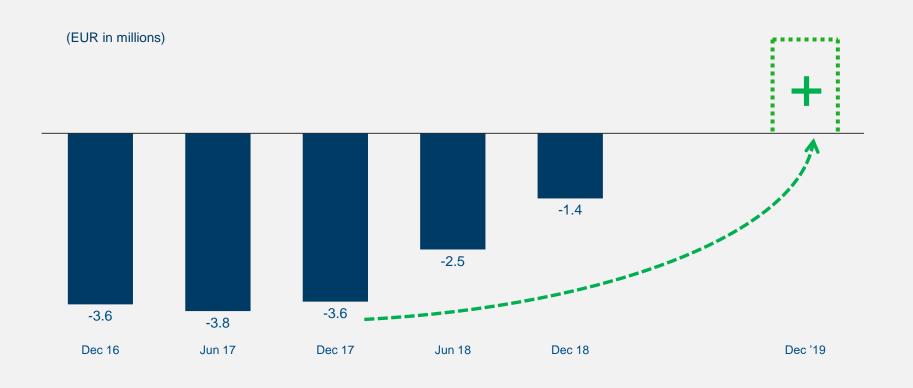
Utilizing Own Manufactured Saline in Grifols' Plasma Centers





## **Hospital on the Path to Profitability**

LTM EBITDA Trend at Constant Currency





# Maintaining a Strong Position and Reputation in Iberia While Accelerating Penetration in the U.S. Market

### By Leveraging Our Strong Position, Reputation and Legacy in IBERIA and LATAM

- Broad portfolio including Pharmatech, IV therapy base, medical devices and clinical nutrition
- Advanced hospital pharmacies
- Established leaders; learning, trialing
- Manufacturing and engineering advantages

### By Accelerating U.S. sales - Offering a Solution Aligned with U.S. Market Drivers & Grifols Strengths





## Current Market Conditions for IV Compounding in the U.S.

Tailwinds Support Market Expansion and Dynamics



Personalized medicine and aging population are driving a growing number of compounded sterile preparations



Regulatory pressures are driving clean room improvements and investments in automation

Tougher FDA, USP & SBoP requirements are leading to new challenges for compounding pharmacies (503a and 503b)



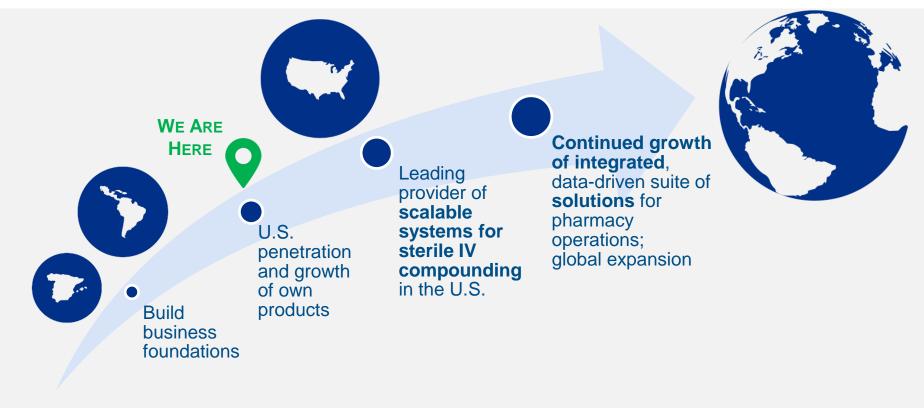
Dynamic conditions exist as many organizations are receiving notification from regulators regarding quality issues

Closures or "cease of operations" occurring in compounding pharmacies within health systems and outsourcing companies



## Pharmatech Ambition for IV Compounding Ecosystem

**Toward Strategy Execution** 





## Global IV Compounding Market Is Set for Sustained Growth

Strategy Poised to Meet Growing Market Needs and Future Demands

### **U.S. Growth:**

- **GRIFOLS Inclusiv** portfolio sales in the U.S. have more than doubled in one year
- Continued strength is expected as ~6k hospitals deal with challenging new requirements
- Current adoption of IV workflow software is estimated at ~20% and experiencing high growth
- Adoption of automation solutions in IV compounding is low and expected to experience sustained growth

### **Global Growth:**

 Other markets have already begun to adopt stricter standards for IV compounding and this trend is expected to continue to expand globally Global IV
Compounding market

\$1.3bn

in 2017

Global Pharmacy Automation market

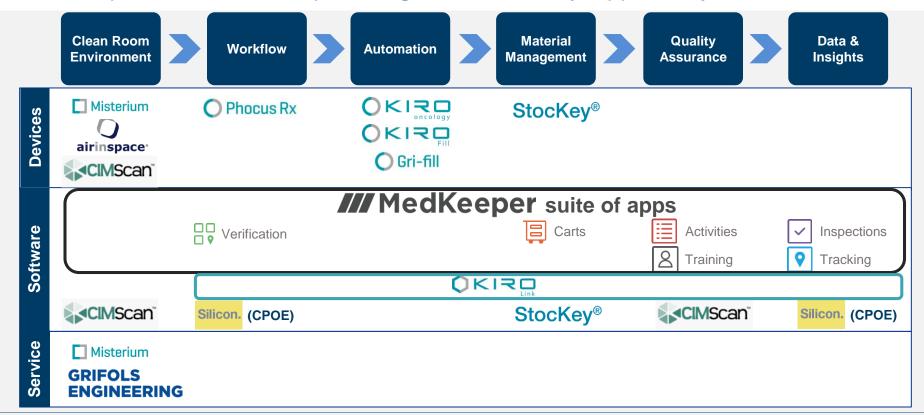
\$6bn

in 2017

Source: Pharmacy Automation: Technologies and Global Markets Report from BCC Research, 2016; IAS026E and also on Global Cleanroom Technology Market Size: Industry Report from Grand View Research, 2017; GVR-2-68038-216-7

## **Complete Solution to Meet IV Compounding Needs**

Most Comprehensive IV Compounding Portfolio: A Key Opportunity





## **Complete Solution to Meet IV Compounding Needs**

Most Comprehensive IV Compounding Portfolio: A Key Opportunity

	Compounding Inventory	Aseptic Procedures Consulting	Cleanroom	IV Workflow Management	Semi- Automated Compounding	Robot Oncology	Robot batch IVs
Grifols	<b>✓</b>	<b>✓</b>	<b>✓</b>	<b>✓</b>	<b>✓</b>	$\checkmark$	<b>√</b> *
Omnicell	<b>✓</b>			<b>✓</b>		<b>✓</b>	<b>✓</b>
Loccioni						<b>✓</b>	<b>√</b> *
ARxium						<b>√</b> **	<b>✓</b>
Baxter				<b>✓</b>	<b>✓</b>		
ICU				<b>✓</b>	<b>✓</b>		
BD	<b>✓</b>			<b>✓</b>			

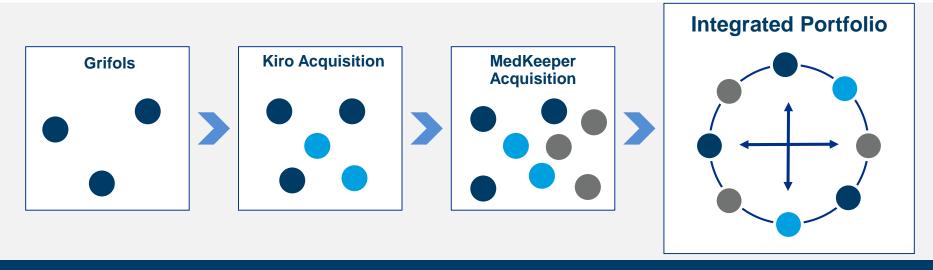
\* Under development: Kiro Fill for Grifols. Little is known about the Loccioni product.

\*\* ARXIUM claims that RIVA could be used for Oncology/HZ drugs, they report one installation but not confirmed results yet Baxter,

ICU (Hospira) and Becton Dickinson (Alaris) have smart infusion pumps (IV Delivery & Administration)

## Positioning for a Comprehensive and Integrated Portfolio

Strengthening Grifols' Portfolio



### **Customer Experience**

Moves from an isolated product offering to an integrated system in which each product plays a key role

Partner vs. Product expert



## **Core Insight**

### It Is All About Safety

- Pain points included the need to meet regulatory requirements, demands for efficiency, and a host of other daily pressures
- However their underlying motivation was safety:



- Safety was far and away their primary motivation —specifically, patient safety
- Most pharmacy directors had chosen to work in hospitals to work more closely with patients
- In their leadership roles they felt they could have a larger impact on safety

### **New Branded Portfolio**

Strengthening Grifols' Portfolio



A single over-arching portfolio of integrated products – customer focused.

Power is in the system vs components

# Protecting patients is your most important responsibility.

inclusiv is an IV compounding portfolio that integrates technology, software and services.

Designed to keep your patients safe through:

- IV Workflow Management Systems
- Modular Cleanroom Systems & Consulting
- $\bullet$  Robotics & Other Machines for Sterile Compounding
- Pharmacy Operations Software

And that's all backed by 75 years of proven experience in sterile manufacturing environments.

Visit www.inclusiv.com to learn more.

HELPS ENSURE USP <797> AND USP <800> COMPLIANCE

## **Portfolio Offers Strong Value Proposition**

## Strengthening Grifols' Portfolio



- Customers felt the range of products in the portfolio presented was comprehensive and few considered it to be missing any components
- They responded very positively to the idea of a single provider that could offer a full portfolio of solutions for the IV compounding area and also found it unique

It seems like a one-stop shop. It's good. I haven't seen anybody actually present it in a complete package like this"

I haven't seen anybody that has breadth of services like this"



## **Inclusiv Brand Communicates Breadth and Depth**



Strengthening Grifols' Portfolio

 The winning brand name and logo successfully communicated the breadth of a connected IV portfolio consistent with the value proposition presented to customers

Is when they have everything that's needed for compounding"

Initially, when I looked at it, it said to me it's everything ...which tells me that it's all things IV"

## **New Campaign Communicates Safety and Strongly Resonates**

## Strengthening Grifols' Portfolio

 The advertising concept successfully communicated patient safety in a way that connected emotionally with customers



 The portfolio was seen as the solution that could help customers protect their patients

We have the ability to either care for the family through the baby or not; it's more than the little one there. It's about the patient, protecting our patients, and that's the most important thing we do, that's why we're pharmacists"

As a healthcare provider it's our responsibility, sort of like a parent to a baby, because they can't protect themselves. It starts with somebody overseeing the protection and the safety process. It's relatable, especially if you have children, as well as some of the history causing the movement in safety in the compounding space"

# Inclusiv Portfolio Is a Good Fit Within Broad GRIFOLS Relationship Strengthening Grifols' Portfolio

"What does **GRIFOLS** signify for me? Stability. Commitment. They are there to develop long-term relationships and that's what I like about them"

## **Key Takeaways**

### Near-Term

### Strengthen

Leverage leadership and expertise in IBAM to accelerate growth and profit in U.S.

Drive value of **MedKeeper & Kiro** acquisitions

Execute core strategies in IV Compounding Control, Contract Manufacturing and IV Solutions

### Mid-Term

### **Expand**

As growth accelerates, resource for sustainability, including expanded Multichannel Marketing capabilities

Consider growth through adjacent strategies

### Longer-Term

### Lead

Leading provider of scalable systems for sterile IV compounding in the U.S.

Continued growth of integrated, data-driven suite of solutions for pharmacy operations through global expansion

Grifols Hospital Division Will Continue Its Strategic Evolution to Become a Comprehensive Solutions Provider for the Operational Pharmacy





## **Evolution in Our One Grifols Approach**

From Opportunistic to Systematic and Strategic



### Research

### **Customer Needs & Expectations**

- Strategic Partnerships:

   Fully understand customer goals
   and priorities offer tailored,
   strategic solutions
- Ease of Doing Business:
   One account manager as singleaccess point for each customer



### Act

### **Strategic Account Management**

- Cross-training: Knowledge of full commercial portfolio
- Cross-pollination: Businesses meet/discuss common customers
- Education: Interpret buying signals/purchasing process across all businesses



### **Measure**

#### **Monitoring Our Impact**

- New Opportunities: As the broader portfolio has been socialized new opportunities for all businesses have emerged
- Awareness: Evolution in customer awareness of our full portfolio & how new opportunities impact our existing baseline business



# IDNs Are Key to Build the U.S. Immunohematology Business

Leverage Existing Relationship to Open Doors for Our BTS Products

IDNs and GPOs Organizations

Grifols Organization

Best Portfolio to Meet Customer Needs

Consolidation is a key trend on the industry

+500

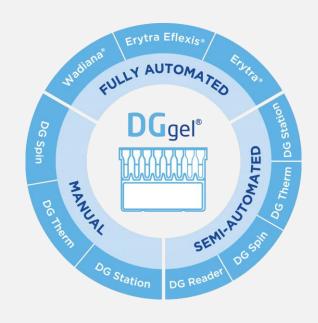
M&A ACTIVITIES (SINCE 2014)

IDNs and GPOs goal is to reduce members' operating costs

# ONE GRIFOLS

We are structured to ensure that each buying experience promotes our corporate values in terms of:

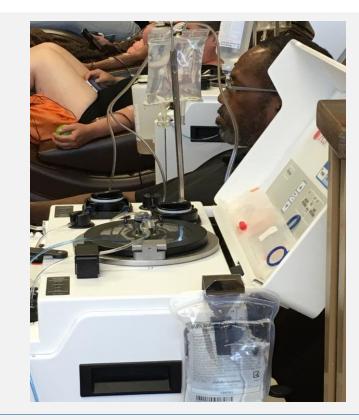
- Patient Safety
- Operational Ease
- Financial Transparency



# **Self-Sustainability with IV Solutions**

Saline Today and Anti-Coagulant in the Future

- 100% of Normal Saline for Biomat is from GRIFOLS
- NDA for anti-coagulant solution (sodium citrate) has been submitted to FDA



# One Grifols Panel Impact on Customer Engagement



#### Grifols, the Provider of the Portfolio

#### A Robust Strategy Dynamically Positions the Division

- Most customers were familiar with Grifols pharmaceutical products and had positive impressions
  - Reliability (consistency of supply) and product quality were mentioned
- Only a minority of customers were aware of the products and services offered by Grifols in the IV compounding area. Those who were familiar with them had positive experiences with clean room design and consultation
- Most found it credible that Grifols could offer the portfolio, citing our ability to meet rigorous cGMP standards required of an FDA-approved pharmaceutical manufacturer

I think they have a good brand name associated with them. Positive connotation"

They are the experts in this.
They are living it through their manufacturing arm. They have the ability to help you out"



# **Improving A1AT Genetic Diagnosis**

Partnerships With Local Laboratory or In-house Testing Services

# Model A local partner laboratory



6,500

SAMPLES TESTED

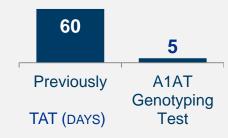
- Reduction on diagnostic times
- Decrease need for sequencing
- Positive user feedback

Model B
Service at Progenika laboratory

# Progenika Biopharma

GRIFOLS

Significant Reduction of Laboratory Turnaround Time



- 3,500 samples processed
- 29 new patients candidate for treatment



Partner with local laboratory or in-house testing services

## **AATD Testing**

#### First Buccal (Cheek) Swab on the Market for Alpha-1 Diagnosis



#### **Benefits to Healthcare Providers and Patients**

Technology, innovation and services improve ease of use and convenience for HCPs and patients



Easy to use



Non-invasive



Detects 99.9% of AATD cases



Rapid results (<1 week)



Reliable results (EU and FDA approval)

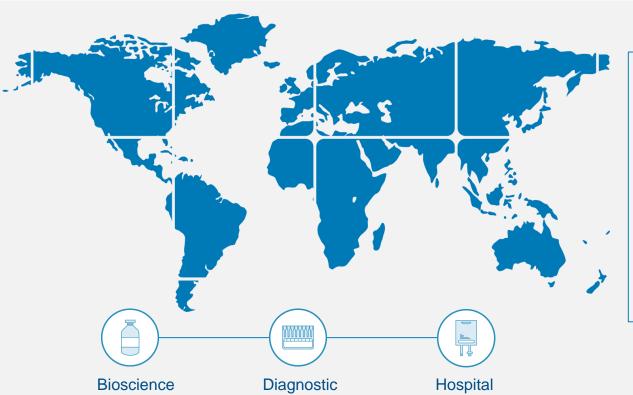
# **One Grifols Panel**

**Internal Impact** 



#### **Creation of Global Commercial Technical Services**

New Global Function Will Support Customers Across All Divisions and Regions

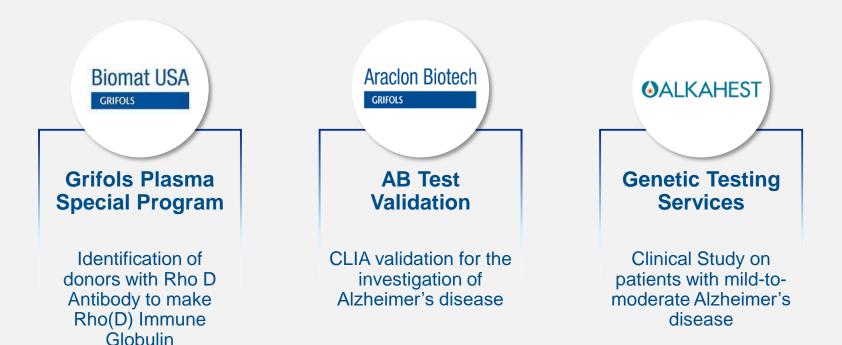


#### **New Ways of Working**

- Identify and use common technologies and approaches
- Strengthen collaboration between industrial and commercial functions by building a customer first framework
- Optimize the customer experience and leverage internal synergies to the benefit of all divisions

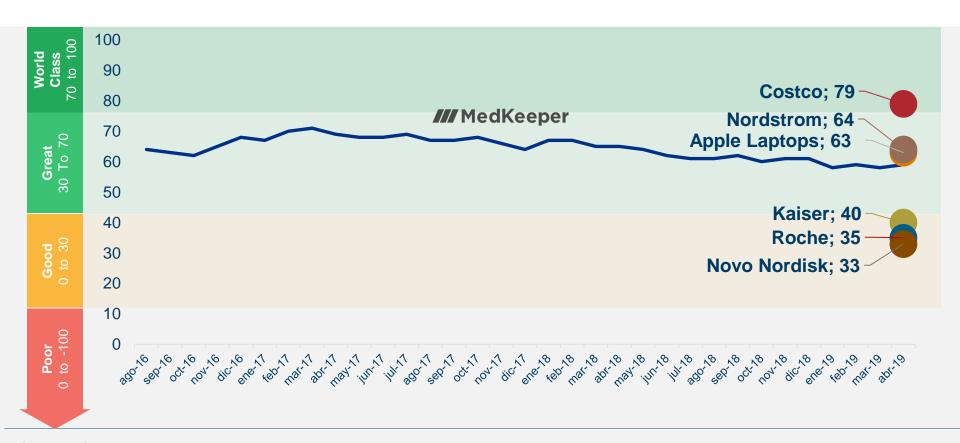
## **CLIA Lab in San Marcos Provides Multi-Divisional Support**

Multiple Collaboration Projects With Bioscience and Other Areas





# **Grifols-MedKeeper NPS Rolling 12-Month Average**





#### **One Grifols Awards**

Enterprise-Wide Incentives Support a One Grifols Culture





# **One Grifols Panel Discussion**



# **Grifols Commercial Is Achieving Lasting Success**

Leadership and Successful Track Record

- Talented team with proven ability to execute and overcome obstacles
- Strong market fundamentals across business lines
- Long-term benefit accruing from One Grifols customer focus
- Planned launches building on strong foundation for future growth
- Continued growth through geographical expansion

# Plasma Procurement Strategy Capacity Leadership in Plasma to Optimize Growth

**Peter Allen** 

President and CEO, Biomat USA

**GRIFOLS** 

# **Grifols Plasma Procurement Is Strong and Well Positioned**

#### Capacity Leadership Drives Growth Opportunities

- 122 additional centers since 2016
- Global network of +290 centers on two continents; pending three
- Collection network will expand to approximately 370 by 2024
- Advance strong self-sufficiency position
- Focus on donor segmentation
- Excellence in logistics and lab testing; accuracy and throughput
- Improving efficiencies through technology and programs

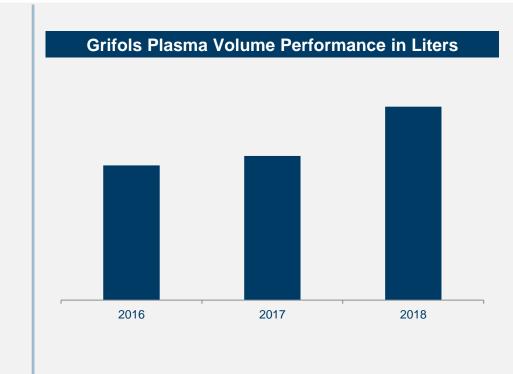




## **Double-Digit Collections Growth With Strong Performance**

#### Capacity Leadership Drives Growth Opportunities

- Plasma obtained have increased significantly, driven by improved processes, extension of hours, facility expansions, and acquisitions
- Benefitting our plasma collection growth is a fully integrated and balanced plasma procurement organization
- Substantial work has been done to focus on both donor and center employee recruitment and loyalty
- · Continued focus on quality performance





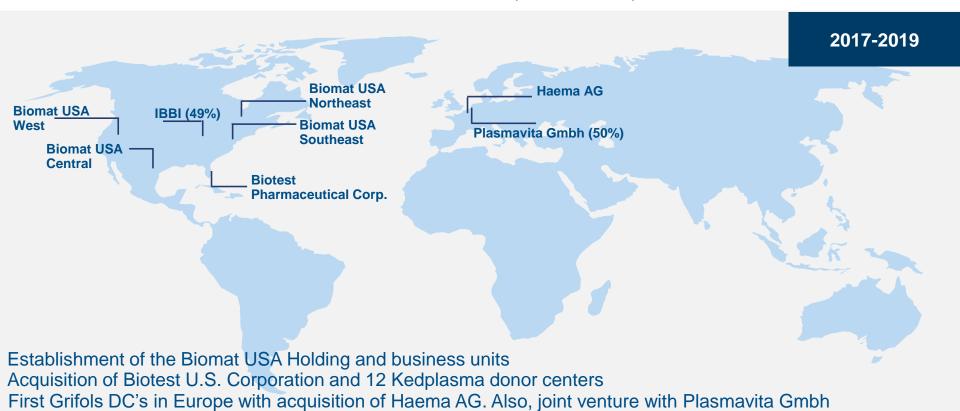
Grifols' Plasma Donor Centers U.S.: 171 (Dec. 2016)



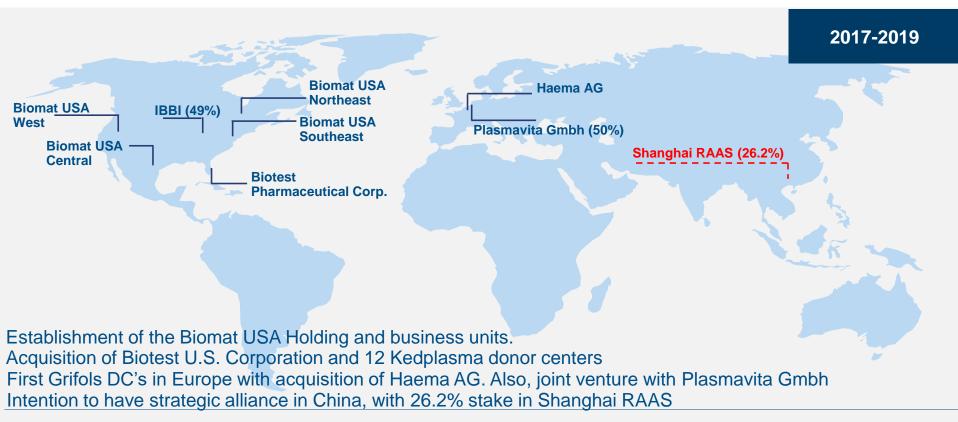










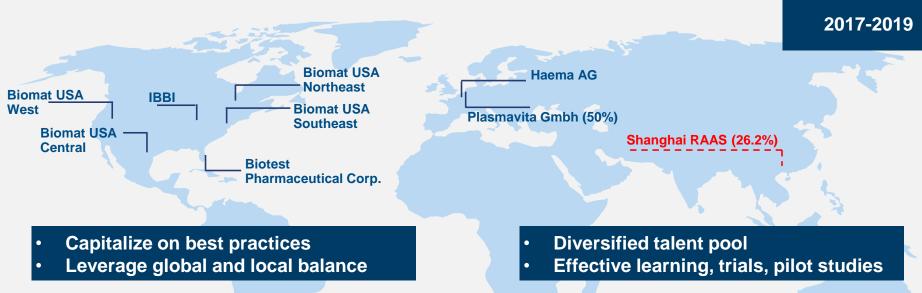








Grifols' Plasma Donor Centers Worldwide: 293 (June 2019)

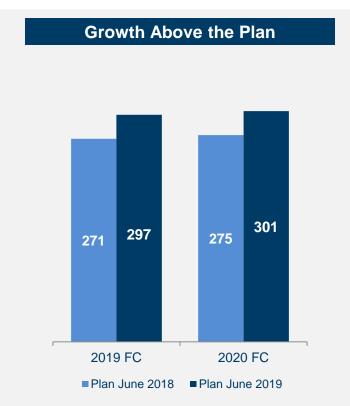


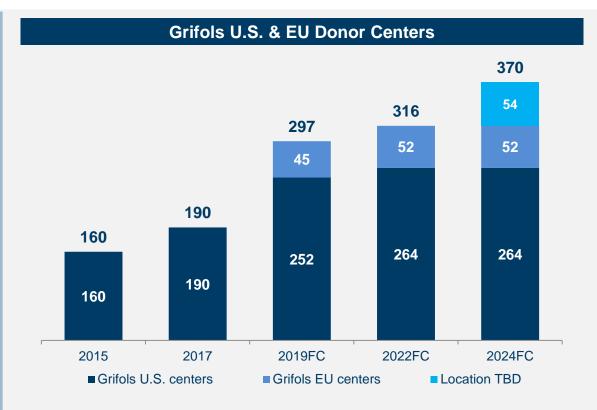
Establishment of the Biomat USA Holding and business units
Acquisition of Biotest U.S. Corporation and 12 Kedplasma donor centers
First Grifols DC's in Europe with acquisition of Haema AG. Also, joint venture with Plasmavita Gmbh Intention to have strategic alliance in China, with 26.2% stake in Shanghai RAAS
Acquisition of the remaining 51% of IBBI



# Plasma Procurement Strategy: Expansion and Diversification

Capacity Leadership in Plasma to Optimize Growth Opportunities

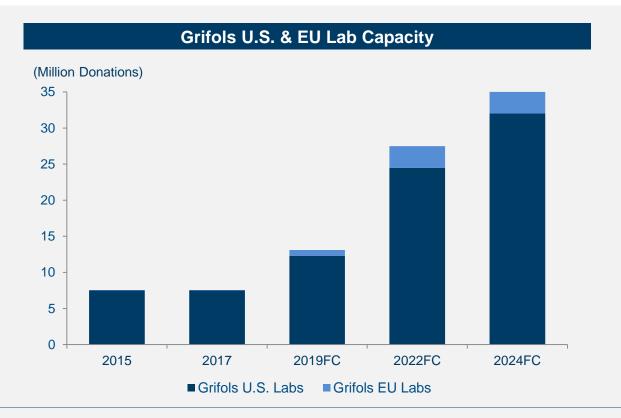






# Plasma Procurement Strategy: Lab Expansion

Capacity Leadership in Plasma to Optimize Growth Opportunities

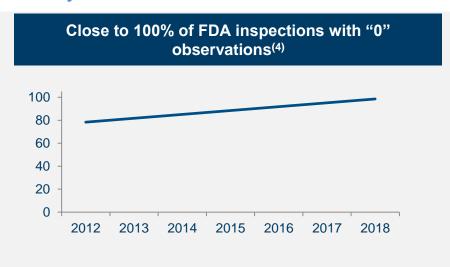




## Regulatory Inspections in 2018

Grifols' High Standards Ensure Operational Efficiency and Sustainable Growth

Agency	Inspection Days	Admin Actions <sup>(2)</sup>
FDA <sup>(1)</sup>	405	0
EU	313	0
COLA/CLIA	86	0
PPTA	82	0
Other <sup>(3)</sup>	116	0
TOTAL	1,002	0



A proven track record: no administrative actions or other regulatory issues promote cost savings across the value chain



<sup>1.</sup> More than 90% of FDA inspections resulted in 0 observations

<sup>2.</sup> Suspension, revocation or loss of any license or certification; warning letter; imposed suspension of any regulated activity, etc.

# **Plasma Procurement Strategy**

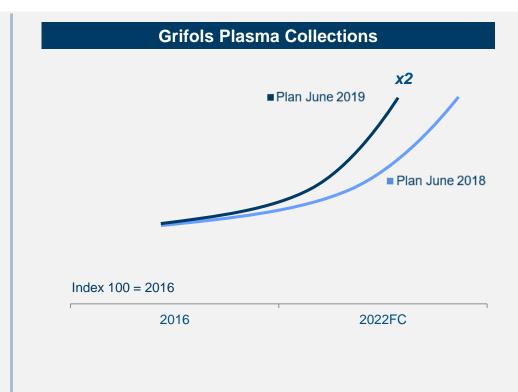
Focus on Grifols' Competitive Advantage

#### **Growth**

- Acceleration in growth of plasma centers
- ↑ Number of donor centers
- x2 Collections 1 year in advance
- ↑ EU plasma supply; diversifying supply
- Expand business development capabilities

#### **Operational efficiencies**

- Standardizing processes
- Continuous quality assurance best practices
- Leverage current and new technologies



# Market and Self-Sufficiency

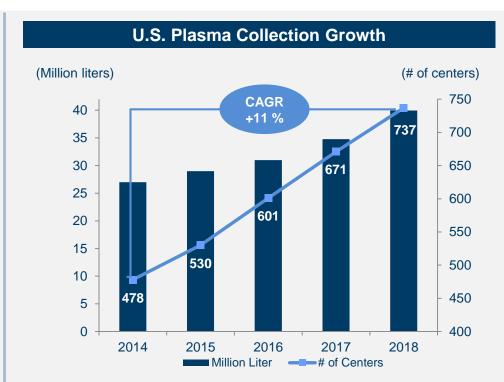


#### **U.S. Plasma Market**

#### Plasma Collection Growth<sup>1</sup>

#### **Consistent Growth**

- Plasma collection has continued to be a large, growing industry year-on-year
- In 2018, the U.S. plasma market collected c.40 million liters
- The number of donor centers reached 737 by the end of 2018
- Increasing collections and recruiting qualified staff are main challenges
- Growth in volume with CAGR +10%
- Growth in <u>centers</u> with CAGR +11%



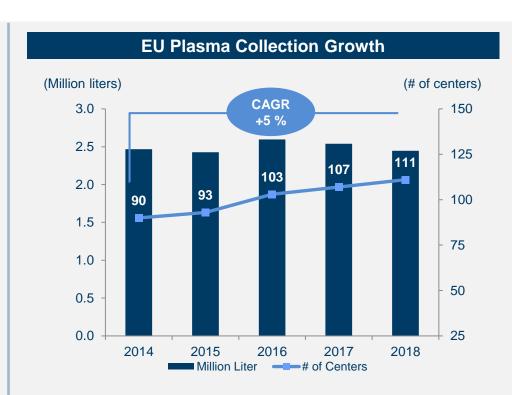
1 Source: PPTA - The Plasma Protein Therapeutics Association data Plasma figures corresponds to plasma from plasmapheresis

#### **EU Plasma Market**

#### Plasma Collection Growth<sup>1</sup>

#### **Restrained Growth**

- In 2018, the EU plasma market collected c.2.4 million liters (Germany 1.6, Austria 0.5 and Czech Rep and Hungary 0.3 million liters)
- The number of donor centers reached 111 by the end of 2018
- Increasing collections and recruiting qualified staff are main challenges, as volume growth is flat in the last years.
- Main players, Grifols (Haema), Octapharma, CSL, Biotest, TMD and KedPlasma
- Growth in centers with CAGR +5%



1 Source: PPTA EPCC - The Plasma Protein Therapeutics Association data Plasma figures corresponds to plasma from plasmapheresis

## **European Collection Dynamics Continue to Evolve**

European Collection Requirements Differ From the U.S.

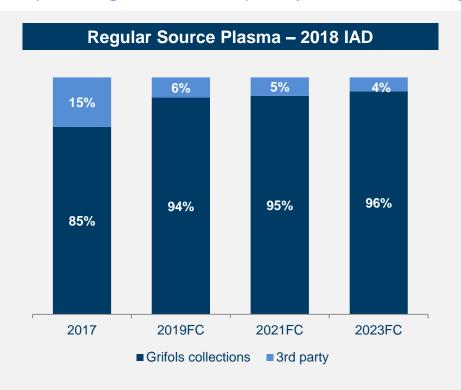
	EU	U.S.
Frequency	Varies; From 2x/7 days (48h Lapse) to 1x/14 days From 24 to 60 donations/year	2x/7 days (1 day btw 2); 104 donations/year
Collection volume	Varies, based on weight or estimation % blood From 600 ml to 850 ml	Std. volume based on weight; 690, 825, 880 ml
Donor compensation	Reimbursement only in 4 countries (CZ, DE, AT & HU)	Not regulated; Compensation for time/effort at market rates
Donor culture	Creating donor cultures: centers smaller/newer	Developed culture in most markets

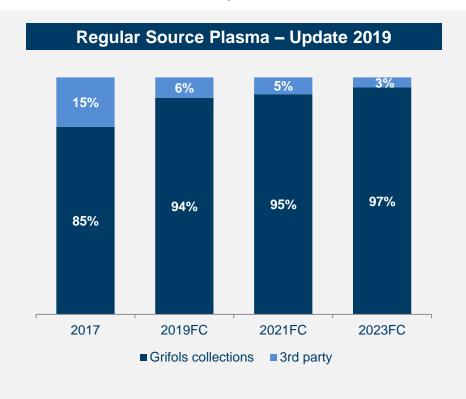
- U.S. utilizing more of its collection capacity; less for export
- EU dependency on U.S. source plasma must be replaced
- EU nations vary in regulatory statutes impacting viability for collectors



# Plasma Procurement Strategy: Update 2019

Expanding Plasma Capacity While Working Toward Self-Sufficiency<sup>1</sup> Ahead of Plan

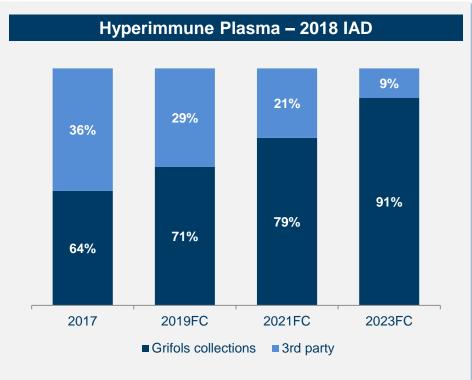


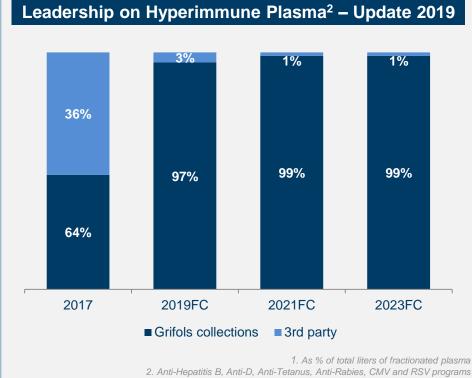


1. As % of total liters of fractionated plasma

# Plasma Procurement Strategy: Update 2019

Expanding Plasma Capacity While Working Toward Self-Sufficiency<sup>1</sup> Ahead of Plan

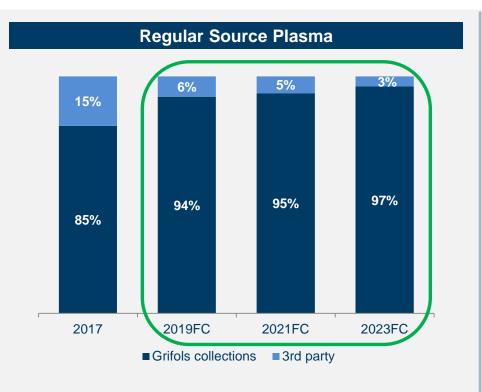


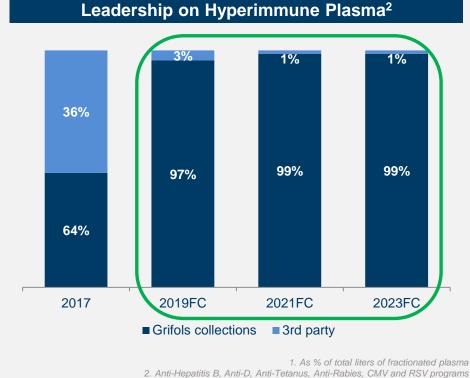




# Plasma Procurement Strategy: Update 2019

Expanding Plasma Capacity While Working Toward Self-Sufficiency<sup>1</sup> Ahead of Plan







# Positioned for Self-Sufficiency in Plasma Collection

One Grifols and Long-Term Contracts Protect Our Market Position and Growth

- Plasma collection centers
- Source plasma
- Hyperimmune plasma
- Testing
- Logistics
- Saline (Biomat USA centers 100%)
- Anti-coagulant (2019 approval)







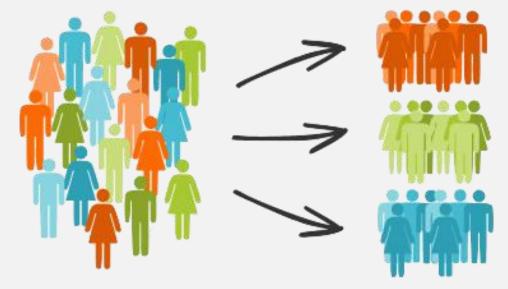


# **Plasma Procurement: Future Growth**



### Plasma Procurement: Future Growth

Delivering Personalized Experience Through Segmentation



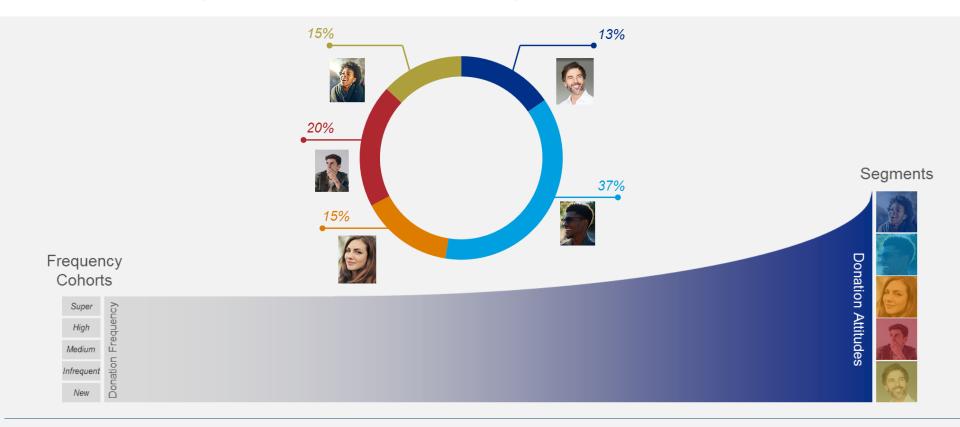
Segmentation analysis seeks to identify natural segments in the market by

MAXIMIZING SIMILARITIES WITHIN SEGMENTS and MAXIMIZING DIFFERENCES BETWEEN SEGMENTS



## Five Segments Identified, Representative Across Fleet

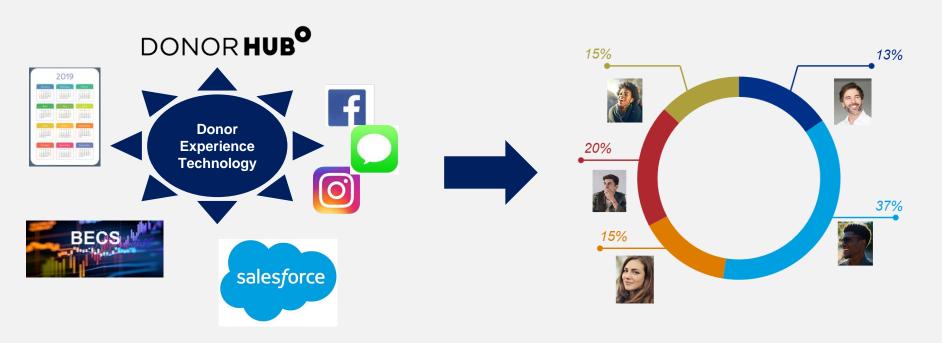
Shift from Differentiating by Donor Frequency to Personalizing by Life Philosophy and Donation Attitudes





### **Technology Ecosystem Enables Engagement**

Uniquely Positioned to Deliver With Internal Development and Partnerships





## **Plasma Productivity Journey**

Journey Status on Track

### Journey achievements

- Plasma technology plan defined in alignment with 4 goals
- Acquisitions providing essential insights
  - Collection technology
  - BECS
- Business Process Management tool







**HAEMONETICS®** 





caring for life

# Plasma Supply Chain Efficiencies



## **Logistics: Integrated Plasma Supply Chain**

Capable Multi-Site System Drives Cost Reductions

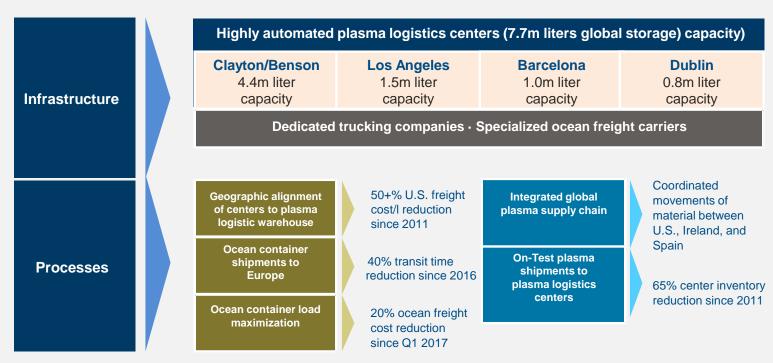
- 125% throughput increase with a modest increase in labor
- One shared database among multiple locations (LA, Clayton and Ireland)
- Grifols U.S. centers and warehouses currently operate with centralized release, enabling efficiencies in inventory management
- Semi-automated plasma clearing lines
- Automated freezer, conveyors and pallet automatic retrieval systems
- Back-up systems to support emergency situations



## **Logistics: Integrated Plasma Supply Chain**

Capable Multi-Site System Drives Cost Reductions

### Alignment across the supply chain drives cost reductions





# **Key Takeaways**



### **Key Takeaways**

### Capacity Leadership to Optimize Growth

#### FOCUS ON SOURCING AND DIVERSIFICATION

- Grifols is committed to maintaining its leadership through a sustainable growth in plasma collection by promoting a fully integrated plasma procurement organization
- Grifols is investing in new centers to continue the plan to reach approximately 370 by 2024
- Plasma procurement is now on three continents, further diversifying plasma supply
- Self-sufficiency positions Grifols for stable performance and cost benefits

#### FOCUS ON DONOR ATTENTION AND EFFICIENCIES

- Donor focus and attention is continuously refined and supported
- · Grifols' commitment to highest quality and safety standards remains top priority
- Operational efficiency improvements include continuous upgrades of plasma centers and customer service to increase donor recruitment and loyalty
- Excellent testing turnaround times and flexibility in testing laboratories



# **Bioscience Manufacturing Operations**

**Excelling at the Fundamentals** 

**Eduardo Herrero** 

**President, Bioscience Industrial Group** 

**GRIFOLS** 

### **Bioscience Manufacturing Operations**

Grifols' Global Footprint Today





4

Manufacturing sites

**USD 1.3+** 

#### billion

in CAPEX investments over the last 5 years

290+

plasma collection centers across the U.S. and Europe

16,000+

Bioscience operations employees

3,500+

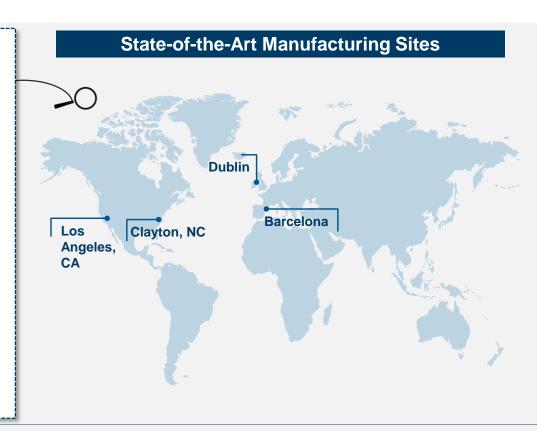
employees in manufacturing sites

450+

R&D employees

12,500+

employees in plasma operations sites



## **Bioscience Manufacturing Operations**

Core Pillars



- **Capacities** 
  - Continuous manufacturing expansion
  - Manufacturing sites operations
  - Fractionation
  - **Purification**
  - Filling
  - Packaging operations
  - Warehouses and QC laboratories
    - **Business model** 
      - L-T growth sustainability
      - Toll plasma fractionation
      - Flexibility and versatility
      - Biosurgery partnership

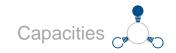
- **Efficiencies** 
  - Manufacturing cost per liter
  - Quality- and safety-driven
  - Digitalization
  - Supply chain
  - Global sourcing
  - Main achievements

# **Bioscience Manufacturing Capacities**

1



## **Continuous Manufacturing Expansion**



Global Enhancement





## **Manufacturing Sites Operations**

Capacities Capacities

A Dynamic and Flexible Approach

	% Manufacturing	(a)	Products	
	activities	Talent*		
Clayton	Fractionation	1,200+	Immunoglobulin Albumin pd Coag. Factors	Immune Globulin (Hyman), 10th
Barcelona	Purification Filling	1,100+	Alpha-1 Antitrypsin Specialty IG, Antithrombin III	CAROLS CAROLS
Los Angeles	Packaging	800+	Immunoglobulin Albumin pd Coag. Factors	Albumin (Human) 20%, USP  reself of V2 Dawn results
Dublin	Packaging	100+	Immunoglobulin Albumin pd Coag. Factors Alpha-1 Antitrypsin Solvents	CRICLS INC.
				* It includes direct and indirect headcount in manufacturing r



## **Fractionation Capacity**

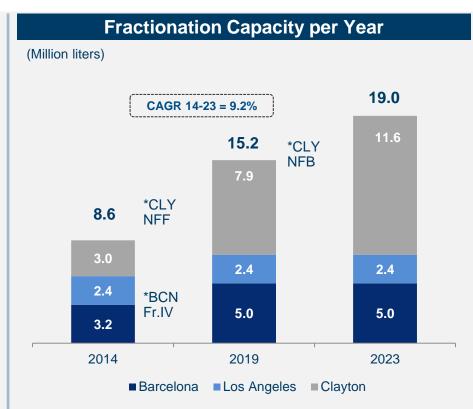
# Capacities \_\_\_\_\_

### Global Capacity Availability at Constant Growth

### **General Remarks**



- Process efficiencies allow continuous incremental capacities: 3% fractionation capacity increase over 2018
- Target fractionation capacity accelerated to be reached in 2021
- Global demand is becoming more complex as new and emerging markets start to play decisive roles
- Grifols Engineering solutions for ABOs in plasma in bags and bottles represent new opportunities (recover and source)





## **Key Proteins Purification Capacity (I)**

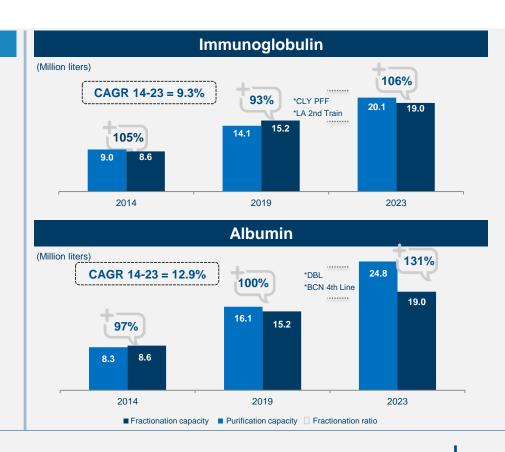


Adapting to Demand Variability

#### **General Remarks**



- Immunoglobulin and albumin are expected to continue driving the hemoderivatives market and Grifols is committed to meeting this demand
- Albumin production increase and additional bag presentation to address the needs of a growing market
- Balanced ratio fractionation and purification





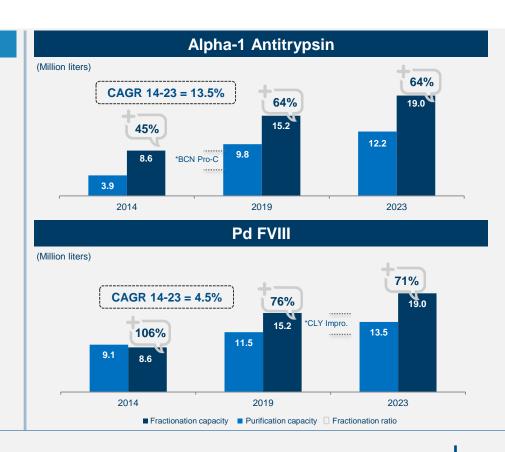
## **Key Proteins Purification Capacity (II)**



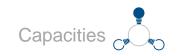
Adapting to Demand Variability

#### **General Remarks**

- Investments and execution of new facilities, along with validation and regulatory submissions ensure Prolastin®-C facilities in the U.S. and Spain will meet global demand
- pd FVIII will continue balancing Grifols' performance. Equipment upgrades and process optimization both support product availability at a competitive cost



## **Evolution of Filling Production**



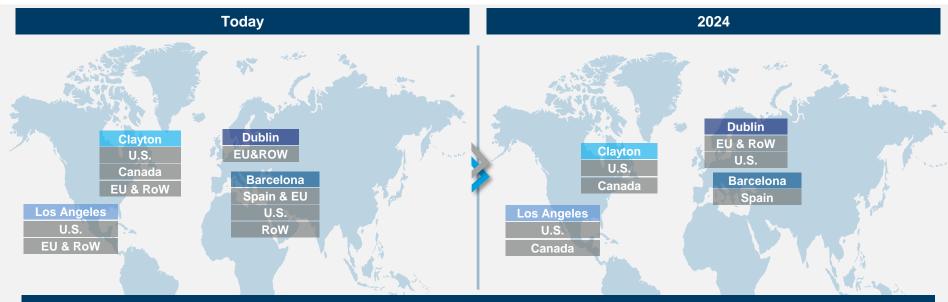
Generating Reliability and Sustainable Growth



## **Packaging Operations**



### Standardized Packaging Process Enhances the Supply Chain



- U.S. plants to cover domestic market
- BCN plant to cover local market and EU toll fractionation
- Increased capacity by site market dedication

- Dublin centralization of packaging activities globally boosts flexibility and efficiency
- Standardized packaging process enables executing on margin recovery at all sites



### Warehouses and QC Labs

# Capacities \_\_\_\_\_

### **Enhancing Manufacturing Chain**



#### **North Carolina Final product Warehouse**

- CAPEX ~ USD 12M
- +6,000 pallet positions → 31,250 ft<sup>2</sup>
- More efficient storage
- Improve physical & IT control over final product
- Risk reduction and no 3<sup>rd</sup> party

#### **Los Angeles Packaging Operations**

- CAPEX ~ USD 17M
- +17,000 pallet positions → +105,000 ft<sup>2</sup>
- · Two new packaging lines
- Future Albumin in Bags packaging line
- +6% of final product storage capacity





#### QC Laboratories in Barcelona

- CAPEX ~ EUR 2.2M
- 6 Kardex carousel → +1.3 M samples
- Clinical archive of samples Library → up to 0.7 M in 2021
- R+D Lab enlargement (+450 m<sup>2</sup>)

#### **QC Laboratories North Carolina & LA**

- CAPEX ~ EUR 5M
- Micro / Sterility Lab
- +107% of area expansion
- Improve performance of assays
- Sterility testing area based on isolators



2



## **Evolution of Manufacturing Cost per Liter**

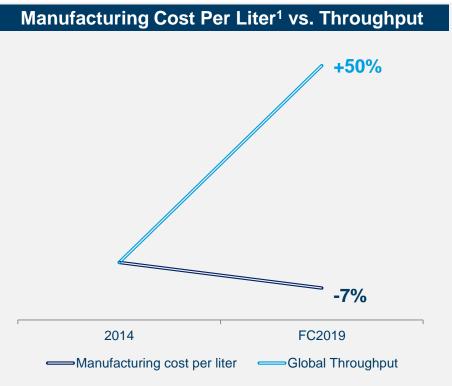


Capacity Leadership in Manufacturing to Optimize Growth Opportunities

### **General remarks**



- New facilities are deployed to work optimally to reduce cost per liter
- Constant upgrades in manufacturing operations lead to continuous yield increases
- Skilled workforce able to produce more per headcount

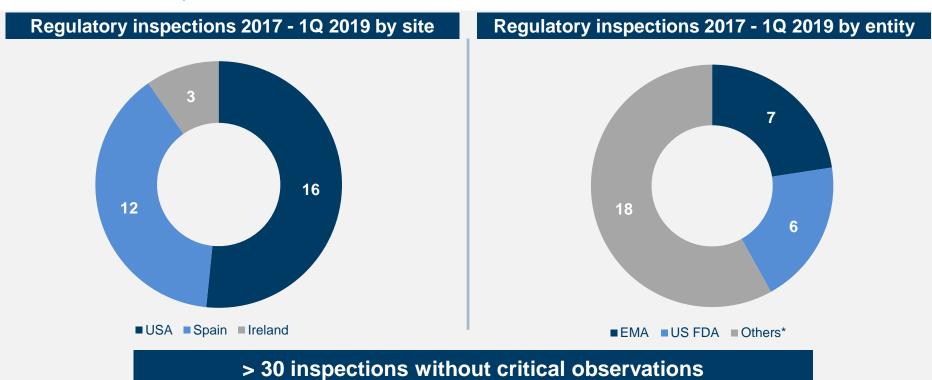


1.- Manufacturing cost per liter does not include amortizations and depreciations

## **Driven by Quality and Safety**



Grifols' Value Proposition



\*Taiwan FDA, Health Canada OSE, CFDI Chinese FDA , MoH of Rep. Kazakhstan, ANVISA Brazil, South Korea MFDS, PPTA



### **Digitalization**

### Preparing for a New Era in Manufacturing

# Efficiencies ••••

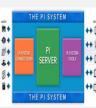
#### **Ecosystem**



Manufacturing plants will collect significant data from operational processes:

- · Big Data
- Predictive analysis
- Artificial Intelligence





Data collected will be leveraged to develop an integral Manufacturing Plant Information System based on KPIs

→ Productivity driven by an efficient ecosystem

### **Ongoing Projects**

#### **Segmentation**

- Automatically guarantee fulfillment of required characteristics for each product license and destination in batches produced
- Production planning based on the capacity of specific lines and manufacturing equipment
- Management of batch allocation according to specific market requirements and prioritization in cases with multiple options
- Cost reduction by ~ EUR 5M for disposal of raw materials and repackaging









#### **Online Notification**

- IT systems in plants connected to an MRP
- Stock optimization
- Minimize documentation errors and information flow up to 85%
- Enhance efficiencies in production processes



## **Supply Chain**



### A New Integrated Planning Process Model: A Roadmap for Grifols' Value Chain

Product
Management
Review

Demand Review Supply Review Integrated Reconciliation

Management Business Review

- ONE Business Plan from strategy to execution, enhancing alignment between production and markets needs
- E2E visibility to improve predictability, anticipation and responsiveness
  - New Tender Management Tool
- Optimize inventory thresholds for plasma, intermediates and finished goods
- Supply chain KPIs
  - Plant attainment
  - · Vials filled, packed and sold

#### **Tender Management Tool**



Vials filled, packed and sold

#### **Plant Attainment**





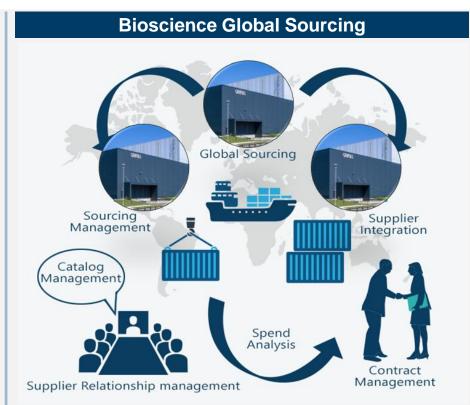


## **Global Sourcing**



### Manufacturing Operations Cost Containment and Risk Mitigation

- Globalization of vendor management is key to ensure both economic efficiency and manufacturing flexibility
- Rationalizing the vendor base allows grouping procurement volumes, leading to better economic outcomes
- Standardizing raw material specifications across manufacturing sites improves flexibility in the supply chain
- Global sourcing mitigates supply chain risks by ensuring double sourcing for critical raw materials
- Nearly 70% of contracts are under global procurement supply and price negotiation instead of local activity by plant, leading to price reductions



### **Main Achievements**



### Manufacturing and Product Approvals - Progress on Track

#### 2018 to Present

- Fr. IV-1/IV-4 co-precipitation in NFF
- Prolastin® from Spanish plasma toll fractionation
- Filling line 3 (Alpha-1 Liquid)
- Approval of Gamunex® 2nd train in LA
- Alternative Plasma storage and packaging in Dublin (FDA)
- Ebola plant and product (CDC and IFER)
- Albumin in bags 25% (FDA)
- Rabies-C and GamaSTAN®-C (FDA)
- IGIV-C (Gamunex®-C) Fraction II+III Paste Weight Increase

### **Ongoing**

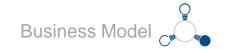
- License submission of Fraction IV<sub>1</sub> from Clayton for Prolastin®-C in BCN
- Submission of 20% SCIG
- Alphanate<sup>®</sup> new method in Clayton
- Albumin in bags 5% and 20%
- Prolastin®-C Liquid 0,5g and 4g
- New packaging lines in Dublin
- Koate<sup>®</sup> Room Temperature
- Modular Thrombin for Ethicon (U.S.)
- 900 IU HyperRAB® vial



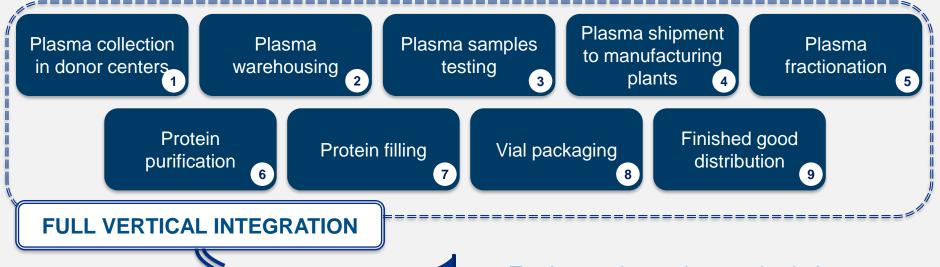
3



## **Long-Term Manufacturing Sustainability**



Prepared to Exploit Market Opportunities



- Total control over the supply chain
- Product availability according demand
- Ability to react to market opportunities

### **Toll Plasma Fractionation**

# Business Model

### Our Track Record Endorses Us

#### **Business Review**



 Promotes countries' selfsufficiency in hemoderivatives



Saving for these countries

- Allows diverting U.S. plasma procurement and manufacturing supply more efficiently
- Develops technology transfer and enhances awareness

#### **Grifols Know-How**



### **Grifols Additional Services**

- Operating in 5 countries
- Projects in new 5 countries (EU and ROW)

Country	Since	Approx. annual volume (L) (2018)	Grifols facility
Spain	1978	375,000	Barcelona
Canada	1988	140,000	Clayton NC
Slovak Rep.	2002	40,000	Barcelona

Reference in Toll Plasma Fractionation with nearly **10 M** of plasma liters collected since 1978

#### **Spain**

- 100% Spanish Plasma under toll fractionation contracts
- 17 different agreements (regions)
- Self-sufficiency: Albumin 56%, F-VIII 47%, IGIV 39%
- 2018 savings in Spanish healthcare system > EUR
   60M

#### **Communication Tools**

Web (Plasma Management Services portal)

PediGri system provides total traceability from donation to the end product

Contract Fractionation Plasma Manual

#### **Quality & Professional Training Services**

Quality program at Academy

Professional Training in GMP

#### **Blood Bank Services Portfolio**

Plasma Transport & Storage

Contingency plans

IPTH & Secure Program



## **Manufacturing Flexibility and Versatility (I)**



Four Manufacturing Plants, One Aligned Approach

#### **Pastes**

Multiple combinations in pastes transfer among manufacturing plants creates back-ups and enables maximizing fractionation facilities utilization

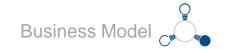


### **Packaging**

Full packaging movement options among manufacturing plants allows meeting potential demand peaks driven by market volatility



## Manufacturing Flexibility and Versatility (II)



Four Manufacturing Plants, One Aligned Approach





## Manufacturing Flexibility and Versatility (III)



Alpha-1 Antitrypsin

### Today

#### Prolastin®

- European market
- Lyophilized product
- Dose = 1g
- Purification & filling capacity > 3 M eqL
- ~70 employees

### Prolastin®-C

- U.S. market
- · Liquid product
- Dose = 1g
- Purification & filling capacity > 8 M eqL
- ~140 employees

## Prolastin®-C facilities versatility enables covering multiple production requirements:

- Liquid and Lyophilized Prolastin
- Factor VIII RV
- Lyophilized Thrombin

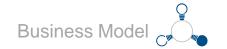
Transition from Lyophilized to Liquid Alpha-1 reports 16% manufacturing cost reduction → solvents, freeze driers and HC

#### **Upcoming years**

Transition from Prolastin® (Lyophilized) to Prolastin® -C (Liquid) for European market



## Manufacturing Flexibility and Versatility (IV)



### Albumin in Bags

### **Performance**

Filling throughput up to 4,000 bags/h

### **Capacity**

Up to 7.5 million of equivalent plasma liters

### Manufacturing presentations

- Bag size of 50 ml,100 ml, 250 ml and 500 ml
- Albumin 5%, 20% and 25%

### **Facilities**

- Pasteurization and quarantine area
- QC laboratory



## **Biosurgery: A New Opportunity (I)**



Leveraging Grifols' Manufacturing Expertise and Industrial Capacity

### **Biosurgery Focus**

- Partner with the global category leader in the treatment of surgical bleeding and leaks, improving and standardizing patient care
- Merging Grifols' track-record in the manufacture of hemoderivatives and biological products and Ethicon's capabilities in medical -device design and robust marketing and commercial structure
- This strategic partnership will lead to a comprehensive biosurgery portfolio, enabling standardizing the use of hemostats and sealants by choosing the appropriate product for each type of bleeding and leak site

# Prevent Leaks / Sealing Stop Bleeding Continuous .... Oozing Problematic Difficult To Access Potential Re-Bleeding Risk

# **Biosurgery: A New Opportunity (II)**



Leveraging Grifols' Manufacturing Expertise and Industrial Capacity

#### Fibrin Sealant VistaSeal®

- New generation of Fibrin Sealant PAS Approval obtained in Q2 2019
- Planned submission to EU in August 2019
- Production in June 2019 → + 300,000 kits by 2020
- Capacity to expand to more than 3M eqL

#### Fibrin Sealant (Packaging & Warehouse)







# **Biosurgery: A New Opportunity (III)**



Leveraging on Grifols' Manufacturing Expertise and Industrial Capacity

#### **LyoThrombin Evithrom**

- FDA submission completed and expected approval in Q3 of current year
- Planned submission in EU in Q2 of current year
- Initial production capacity of 3.8M eqL to be expanded to more than 7M eqL
- Ongoing routine production → +1,000,000 vials in 2020

Grifols Manufactures



Ethicon Distributes



# **Key Takeaways**



#### **Key Takeaways**

#### Manufacturing Expansion Plans to Promote Sustainable Growth

#### **Manufacturing Capacities**

- Global fractionation capacity expansion to be accomplished by 2021, reaching 19 million liters
- Execution of key purification, filling and packaging capacities well balanced as planned
- Addition of facilities for storage, packaging operations and QC laboratories improves supply chain efficiencies in product and inventory management

#### **Manufacturing Efficiencies**

- Continuous improvements in manufacturing performance ensures better manufacturing cost per liter
- Digitalization projects provide overall business optimization in manufacturing and supply chain
- Flexibility in intermediates, purification and packaging operations as alternate manufacturers
- Excellent quality and safety track record

#### **Business Model**

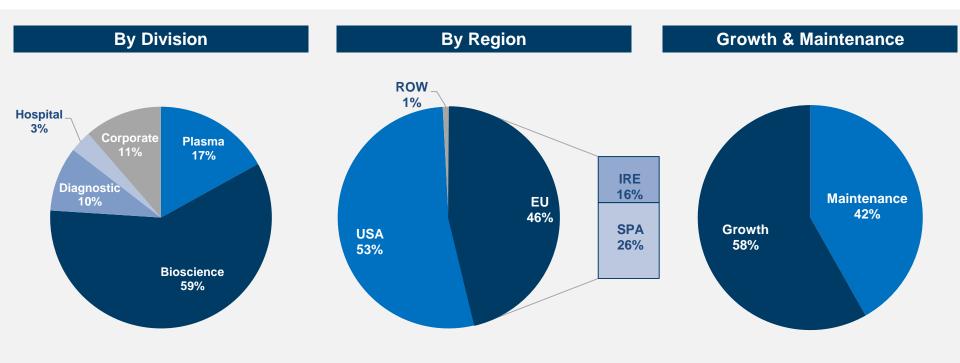
- · Vertical supply-chain integration to promote Grifols' reputation as a reliable and sustainable player in dynamic environments
- Company expertise to expand contract plasma fractionation opportunities
- Partnership in new therapeutics areas using industrial capabilities to promote further growth





# Capital Investments Plan for 2019 – Investing for Growth

**Anticipating Future Market Needs** 

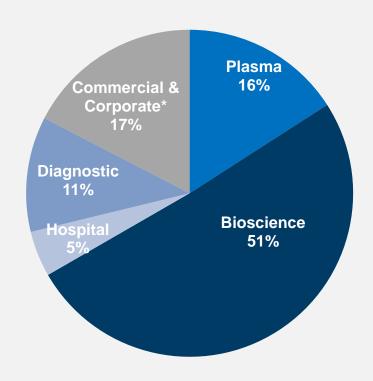




# **Capital Allocation 2018-2022**

1,400 mm

Aimed to Meet Growing Demand



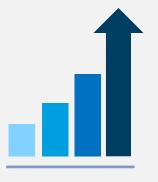
\*Includes land and common infrastructures

## **Capital Investment Program**

Planning for Growth and Leveraging Internal Strengths

ONE GRIFOLS Projects

ONE GRIFOLS **Business Growth Projects** 

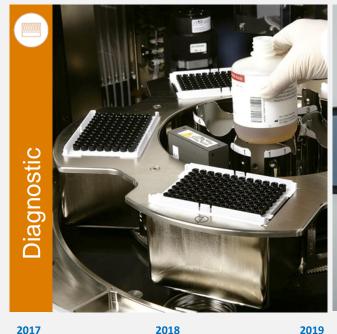


# Diagnostic

# Manufacturing Expansion- San Diego, CA

Consolidation of NAT Reagents Manufacturing Operations









- New facility is located in a nearby 7,000 m<sup>2</sup> standalone building
- This new facility completes the manufacturing operations spin-off from Hologic achieving a higher manufacturing efficiency and improving the product quality
- The new building paves the way for future DX (NAT, IH) growth on the site in further execution phases

2021 2022 2023

Engineering & Approval

Machinery & Construction

Valid.



# Immunoassay Manufacturing Operations, Emeryville, CA







2018



2020

- CMF consolidates all FMV manufacturing operations in a single building, allowing to exit leased properties and reducing site running costs
- The new facility provides enough resources to double the current production capacity for future growth
- New production platform based on mammalian cells culture to develop new antigens and offer specialized GMP CDMO services to third parties

2021 2023

**FDA Antigens** Licensing

Prod. Trans.

2019



# Parets North and Lliçà (Barcelona), Spain

**Current and Future Expansion** 





# Manufacturing Expansion – Lliçà (Barcelona), Spain



Adjacent to Parets Site





- Acquisition of a new land plot of 49,716m<sup>2</sup> in Lliçà, close to the main manufacturing site to expand Diagnostic and Bioscience industrial divisions
- New instruments manufacturing plant will vacate spaces in P4, enabling expansion of reagent- manufacturing capacities. First phase will double the current manufacturing capacity for both reagents and instruments

\*Includes land and common infrastructures

Design

Construction

**Interiors** 

Validation Approval



# Hospital

# Plastic Injection Revamping in Murcia, Spain

Ensuring the Supply of Key Consumables for All Divisions





10<sup>®</sup>

- Cross-divisional project to ensure supply consistency for the business
- Additional manufacturing and assembly capacities for:
  - Bioscience: albumin in bags,
     Fibrinsealant holders, anticoagulant
  - DX: Q Coagulometer cells and racks, BCS
  - Hospital: Fleboflex Luer, Kiro Fill and Oncology disposables

2022

Design

Construction & Interiors

Injectors &Welders

2021



# Anticoagulant and Saline EB3 Line BCN, Spain

Vertical Integration With Plasma Procurement and Hospital





2018







- New fully automated FFS line will provide additional capacity for U.S. IV solutions, as well as production capabilities for new product releases:
  - anticoagulant solutions
  - saline with Luer lock for ○ KIR ☐ and pharmacy compounding

2021 2022 2023

Validation

2019

Approval



# **Bioscience Bag Forming Expansion - Murcia, Spain**

Hospital and Bioscience







- Fully automated robotic forming line for biological bags with multiple port/connector configurations
- This line will further expand Grifols' capacity to produce Albumin in bags and IVIG in the future
- 2,500 BPH capacity (15MMB/year)

2022

Bag formats: 50, 100, 250 and 500 mL

2021

validation

Start-up &

Line design &



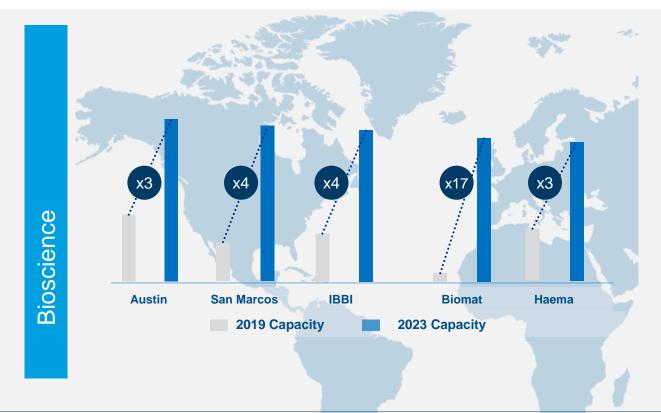
# Bioscience

# **Global Plasma Testing Labs Expansion: Capacities**

Ensuring Access to High-Quality Plasma









# **New Fractionation Building – Clayton, NC**

Expanding Our Fractionation Capacity to 19 million liters



# Purification and Filling Facility (PFF)- Clayton, NC



The World's First Sterile Filling Facility for IgGs in Bags





# Albumin in Bags Purification and Filling Plant - Dublin



Increasing the Production Capacity of Albumin in Bags





## Fibrin Sealant Plant- Parets, Spain

Balancing the Liter of Plasma







- Phase 1 includes the production of fibrin sealant
- Second phase will include topical thrombin and other projects including fibrin sealant from whole-blood plasma and pediatric fibrin sealant
- 3,600 m<sup>2</sup>





## **New Aseptic Filling Facility - Parets, Spain**

Immediate Response to Market Needs





- Leveraging available production areas and infrastructures
- New aseptic filling and freeze-drying areas for new FVIII HC and Lyo **Thrombin**
- Proprietary GSF® technology for aseptic filling
- 1,200 m<sup>2</sup>

2017 2018 2019 2020 2021 2022 2023

**Interiors** 

**GRIFOLS** 

Validation & Start-up

Appro



# **Key Takeaways**



#### **Key Takeaways**

#### Strengthening Competitive Advantage

#### Strategic advantage

Having our own engineering company allows us to build cutting-edge facilities leveraging the market's most competitive investment costs and in the shortest time to market

#### · Technology leadership

Grifols' know-how and experience in developing industrial solutions and facilities positions Grifols as a global technological reference in the sector

#### One Grifols

Grifols' divisional portfolio has been strategically designed and streamlined to leverage capabilities, resources and expertise and vertically integrate critical supplies and operations

#### Consistent execution

Long-range production plan is progressing as expected. Protein-purification and fill-and-finish investments are moving forward, keeping up with the growth in fractionation capacities and in alignment with marketing and commercial demands







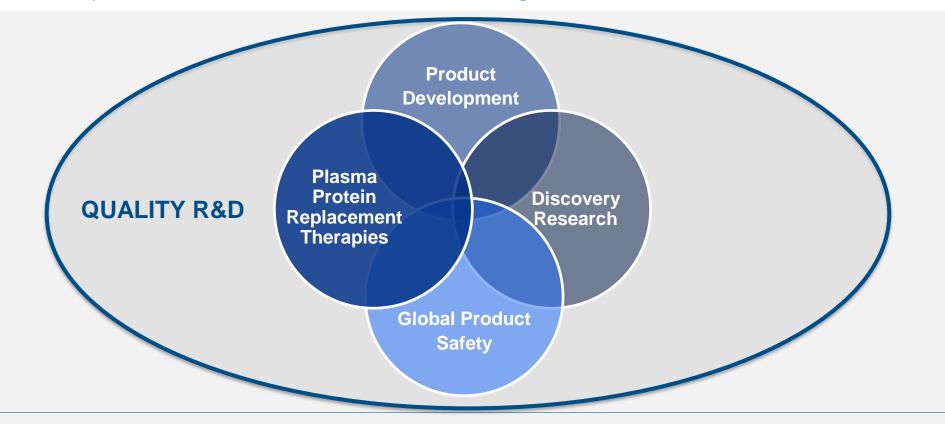
Todd Willis, PhD<br/>VP, Discovery Research

**GRIFOLS** 

# **R&D Bioscience Industrial Group**



R&D Departments Located in Barcelona, Los Angeles and North Carolina





#### Product Development Core Expertise



#### Process Development and Technology

- Process development and formulation
- Process scale-up and transfer to Manufacturing
- Clinical manufacturing (cGMP)

#### **Bioanalytics**

- Process / product characterization
- Assay development, validation, and transfer to QC
- Clinical assay support (immunogenicity)
- Extractable and leachable (E&L) studies

#### **Development Stability**

Intermediate and product stability



#### Plasma Protein Replacement Core Expertise



#### Plasma Protein Replacement Therapies

- Focus on treatment of complex, multifactorial disorders with combination therapies consisting of plasmapheresis and plasma protein replacement
- Collaboration of internal expertise in plasmapheresis and plasma protein therapeutics with key opinion leaders in targeted therapeutic areas
- Movement from treatment of rare disease to management of prevalent diseases

#### Global Product Safety Core Expertise



#### Toxicology

- Board-certified toxicologists
- Design and execute GLP compliant IND-enabling toxicology studies
- Safety evaluation of data from E&L studies

#### Pathogen Safety

- Assess and validate virus/prion clearance capacity of purification processes
- Optimization of pathogen safety steps during process development
- Development of methods to measure viral infectivity, neutralization, antibody content and binding
- BSL 3 containment lab in North Carolina

#### Discovery Research Core Expertise



Deep knowledge of protein biochemistry and purification sufficient to generate innovated project ideas, new IP, and guidance and oversite of external collaborations

#### In vivo Pharmacology

- In-house animal model development
- External CRO oversight of animal studies
- Expertise in PK and PD of plasma and recombinant proteins

#### In vitro Pharmacology

- · Molecular analysis of gene expression
- Cell-based assay formats
- Cellular expression of proteins

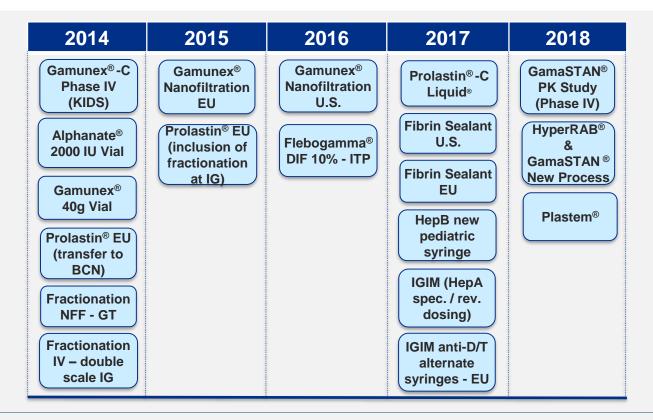


# Bioscience Industrial Group Key Licensures and R&D Development Pipeline



#### **Bioscience R&D**

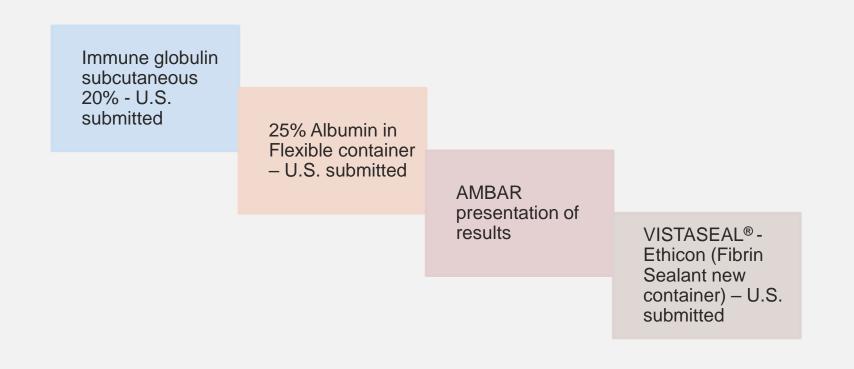
#### Recent Licenses 2014 - 2018





#### **Bioscience R&D**

Key 2H 2018 R&D Highlights





# **Bioscience R&D Development Pipeline – June 2019**

Protein	Preclinical	Clinical Non-pivotal	Clinical Pivotal	Life Cycle Management	Regulatory Review
			Alzheimer's Disease (AMBAR)		
Albumin	New Formulations		Albumin in Liver Failure (APACHE)	Albumin in Bags	
	1 officiations		Albumin in Cirrhosis (PRECIOSA)		
			MG Exacerbations	900 IU HyperRAB®	
_	Anti-Infective	Myasthenia	20% IGSC EU	IGIM-C	20% IGSC US
lg lg	Gamunex® New	Gravis (MG) Maintenance	20% IGSC Flex Dose & Daily Push	Stage 2 (Tet, RhoD, Hep B)	
	Process		Flebogamma PPS	Gamunex in Bags	
Alpha-1	Prolastin® C			Prolastin®-C Liquid New Vials	Project Japan
Alpha-1	Concentrated			Prolastin-C Phase IV	(in preparation)
Factor VIII				FVIII/VWF Reduced Volume	
PPF				Plasmanate® MP	
				Plasmanate® in Bags	
Eibringgen			Fibrin Sealant Pediatric Study		
Fibrinogen			IV Fibrinogen		



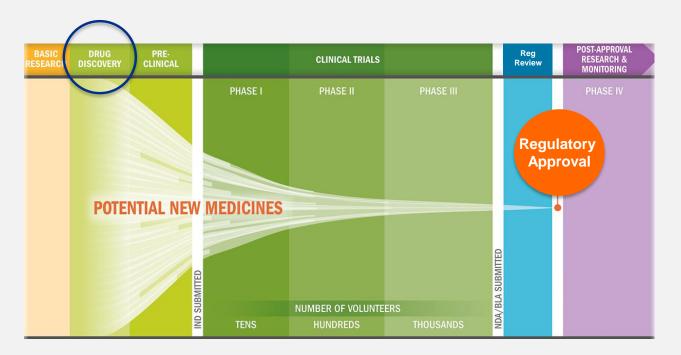
# The Research Pipeline

**Bioscience Discovery Research** 



# **Discovery Research**

#### General R&D Product Development Funnel



Key: IND: Investigational New Drug Application, NDA: New Drug Application, BLA: Biologics License Application

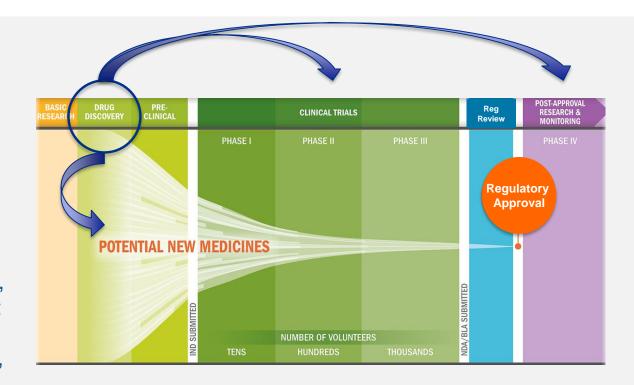
http://www.phrma.org/sites/default/files/pdf/rd\_brochure\_022307.pdf



### **Discovery Research**

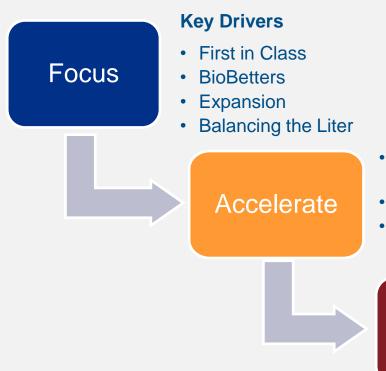
#### How we Operate

- Proof-of-concept studies to demonstrate feasibility (does it work?)
- Mechanism-of-action studies conducted at a molecular level (how does it work?)
- To optimized capabilities, research studies support multiple phases - early research, clinical trials, and post approval



### **Discovery Research**

General Strategy



As long as a medical need exists, no protein therapeutic (**recombinant or plasma-derived**) or therapeutic indication is off limits

- Capitalize on internal research expertise across the organization
- Leverage external research partners / advisors
- Integrate **artificial intelligence** (IBM Watson Drug Discovery)

Execute

- Proof-of-concept studies
- Mechanism-of-action studies
- Quick kills & concentrate resources on "wins"

# **Goals Achieved Through Internal and External Partners**

Sample of Partners

#### **External Experts**





#### **Outsourced Analysis**











#### **External Collaborations**











#### **One Grifols**

- R&D
- Clinical
- Marketing
- Medical Affairs
- Diagnostics
  - **Biomat**
- Hospital



# **Key Strategic Research Partners**

Recombinant polyclonal

platform for

diseases that can

be treated with

neutralizing

antibodies

- Discovery of rare, high-affinity antibody candidates diverse human repertoire
- Identify most efficacious antibodies through massively parallel bioassays that screen millions of antibodies at a time

Plasma proteins for treatment of age-related disorders – "Healthy Aging"

ALKAHEST

 Decoding the plasma proteome during healthy aging and contrasting it to disease states

 Identifying plasma protein fractions to treat complex diseases with multiple mechanism of actions

**GIANT** INNOVATION

Plasma
proteins to
prevent and/or
stop bleeding
during surgical
procedures

ETHICON

a yohmon yohmon company

©GigaGen



#### Four Primary Drivers



#### Innovative protein entities

- Capitalize on unique plasma protein properties, their modifications, and protein combinations
- Utilize latest scientific technologies to reveal connections and relationships between plasma proteins and disease
- Exploit utilization of process waste streams



#### Best in class therapies

- Improve product attributes to enhance safety, efficacy, stability, half-life, convenience, and bioavailability
- Investigate new routes of product administration
  - subcutaneous
  - intradermal
  - aerosol



# New indications and improve position of current products

- Capitalize on emerging technologies (AI) and scientific advancements
- Conduct PoC studies in relevant therapeutic areas
- Partner with key opinion leaders in target therapeutic areas

Balancing the Liter

# Opportunities to develop commercial plasmaderived proteins outside of current portfolio

- Create innovative approaches to
  - reduce development time line and manufacturing cost
  - Identify new potential indications



First in Class



#### **Key Areas of Research**

- Anti-infective with broad microbial recognition and synergistic efficacy with standard of care treatments
  - First anti-infective protein to enter IND-enabling toxicology studies
  - Proof-of-concept studies underway for second microbial target

- Novel neuroprotective protein(s) for treatment of cognitive disorders (Alzheimer's Disease, Parkinson's Disease)
  - Research conducted with internal R&D and in-conjunction with external partners

**BioBetters** 



#### **Key Areas of Research**

- Modulation of autoimmunity
  - Proof-of-concept studies with potentially high efficacy antibodies and comparison to licensed IgG products
- New product formulations
  - Proprietary formulations to maintain desired product attributes and improve clinical outcome in treatment of Alzheimer's Disease and liver disease (cirrhosis)
  - Alternative routes of delivery (subcutaneous and intradermal)
  - New product formulation with an alternative route of delivery to enter IND-enabling toxicology studies

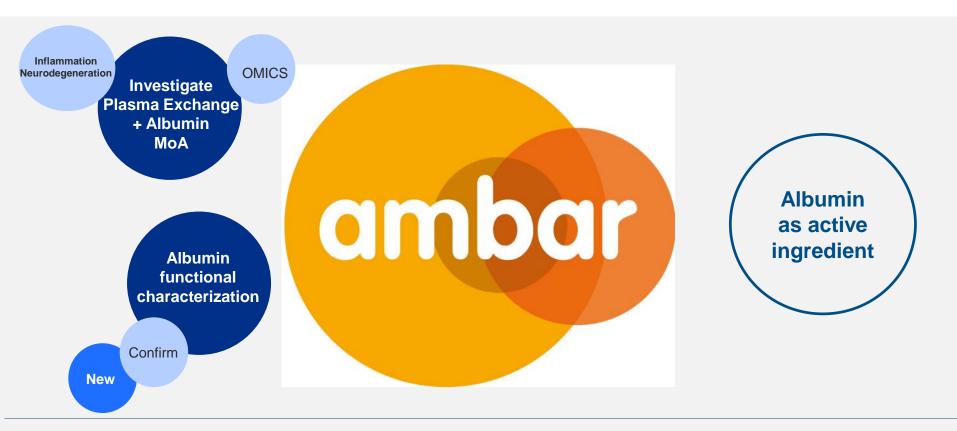
Expansion



#### **Key Areas of Research**

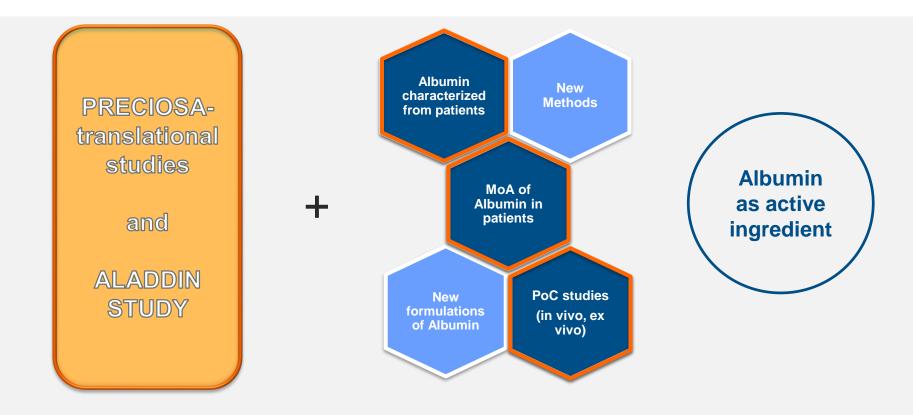
- FVIII
  - Concomitant use of Alphanate® / Koate® with Emicizumab (Hemlibra®), supporting concomitant use of these products
  - Protective effect of VWF towards inhibitor reactivity of FVIII products
- Fibrin Sealant (Fibrinogen and Thrombin) in tissue engineering and surgery
- Albumin as a active protein therapeutic in the treatment of Alzheimer's Disease and liver disease

#### **Albumin Treatment for Alzheimer's Disease**





#### **Albumin Treatment for Cirrhosis**





Balancing the Liter



#### **Key Area of Research**

 Identify key commercial plasma-derived proteins and indications outside of current Grifols product portfolio to Balance the Liter of plasma

#### Create new opportunities

- Introduce new technologies to reduce development and production costs
- Collaborate with internal partners (Regulatory and Clinical) to optimize regulatory path for product licensure
- Collaborate with strategic external partners (Alkahest, Ethicon,...) to identify new potential indications

# **Summary of Research Pipeline by Therapeutic Area**



# Therapeutic Area

Coagulation / Hemostasis	Liver Failure	Tissue Engineering	BioSurgery	Infection	Immunology	Cognitive Disorders	Pulmonology
pdFVIII + Hemlibra	New Formulations	Biolnk	Transplantation	Emerging Pathogens	Autoimmunity	Alzheimer's Disease	New Formulations
New Plasma Protein Opportunities	Cirrhosis		Sealent		New Plasma Protein Opportunity	Parkinson's Disease	
						New Formulations	





# AMBAR: Grifols Alzheimer trial Up-to-Date Clinical and Biomarker Results

Antonio Paez, MD

Alzheimer's Research Group. Medical & Technical Director

**GRIFOLS** 

# **Background**



- 15 years ago we learned that most amyloid-beta (Aβ) circulating in plasma was bound to Albumin.
- A clinical program of Plasma Exchange (PE, Plasmapheresis with Albumin replacement) was initiated in mild-to-moderate AD.
- Pilot and phase II studies with PE showed a decrease in plasma A $\beta$  and an increase in CSF A $\beta$  \*.
- Signals of clinical and functional neuroimaging benefit \*.
- Post-hoc basic research analyses on Albumin have shown an increase of oxidized and glycated forms in plasma and markedly in CSF in AD patients \*\*.

<sup>\*</sup> Boada M et al. Drug News Perspect. 2009; 22(6):325-39 Boada M et al. J Alzheimers Dis. 2017; 56(1):129-143 Cuberas-Borrós G et al. J Alzheimers Dis. 2018; 61(1):321-332

<sup>\*\*</sup>Costa et al. J Alzheimers Dis. 2018;63(4):1395-1404 \*\*Costa et al. submitted to JPAD

# **AMBAR Eligibility Criteria**





Age: 55-85 years



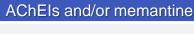
Probable AD (NINCDS-ADRDA criteria)



MMSE score: 18-26



Stable treatment (3 months):





CAT or MRI (12 months)







Moderate: 18-21



#### **AMBAR Treatments**



#### Conventional Therapeutic Plasma Exchange (TPE)

- 1 plasma volume processed
- Albumin replacement (Albutein®): 125-150g

#### Low Volume Plasma Exchange (LVPE)

- New modality of PE developed by Grifols for this trial as maintenance therapy
- Plasma volume processed similar to that of a plasma donation
- Albumin (Albutein®) replacement (less volume infused): 20-40g. IVIG (Flebogamma DIF®) / 4 months: 10-20g

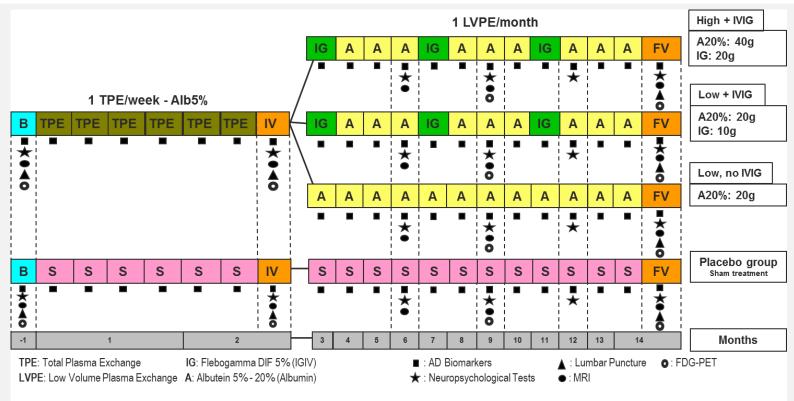
#### Placebo

- Simulated procedure (sham) for both TPE and LVPE. Blind for patient, caregiver and rater
- Devices working in a close circuit manner with colored fluids mimicking plasma and patients falsely connected to devices



#### **AMBAR Schematic**







#### **Outcomes**



#### Co-Primary outcomes

ADAS-Cog: Change from baseline to 14 months

**Presented at CTAD'18** 

ADCS-ADL: Change from baseline to 14 months

#### Secondary endpoints

- MMSE, NPS, NPI, CDR-SB, ADCS-CGIC, CSDD, C-SSRS, QoL-AD, RUD-Lite<sup>®</sup>
- Changes in  $A\beta_{40}$  and  $A\beta_{42}$  in plasma and **CSF**

AD/PD'19 update

- Changes in Tau and P-Tau in CSF
- Changes in brain volume by MRI
- Changes in brain activity by FDG-PET

#### Safety endpoints

AEs and SAEs associated with plasma exchange

# Presented at CTAD'18 + AD/PD'19 update

# **Primary Efficacy Analysis**



#### Main population

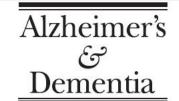
mITT: patients undergoing at least 1 TPE

#### Pre-specified primary analyses (MMRM)

- mITT: 3 treatment arms vs. placebo
- mITT: all patients vs. placebo (all patients share the same plasma removal component of the treatment)
- "Mild AD" (baseline MMSE 22-26) and "Moderate AD" (baseline MMSE 18-21) vs. placebo







Alzheimer's & Dementia: Translational Research & Clinical Interventions 5 (2019) 61-69

#### Featured Article

# Plasma exchange for Alzheimer's disease Management by Albumin Replacement (AMBAR) trial: Study design and progress

Mercè Boada<sup>a,b</sup>, Oscar López<sup>c</sup>, Laura Núñez<sup>d</sup>, Zbigniew M. Szczepiorkowski<sup>e</sup>, Mireia Torres<sup>d</sup>, Carlota Grifols<sup>d</sup>, Antonio Páez<sup>d,\*</sup>

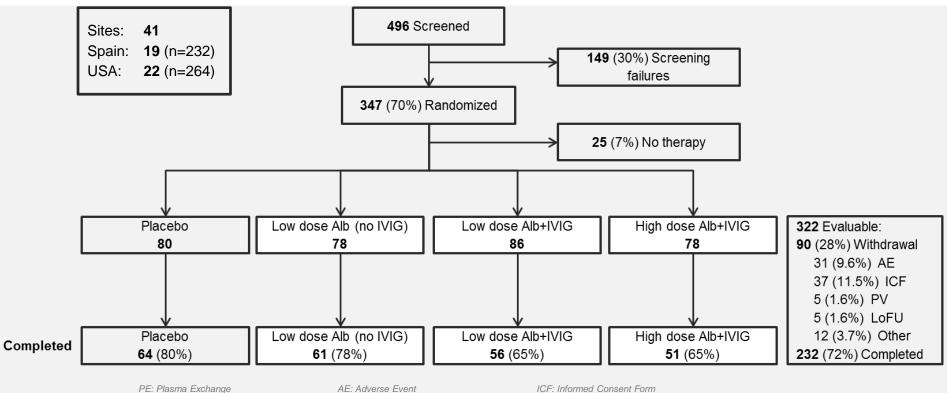
<sup>a</sup>Research Center and Memory Clinic, Fundació ACE, Institut Català de Neurociències Aplicades, Barcelona, Spain
 <sup>b</sup>Facultat de Medicina i Ciències de la Salut, Universitat Internacional de Catalunya, Barcelona, Spain
 <sup>c</sup>Departments of Neurology and Psychiatry, University of Pittsburgh School of Medicine, Pittsburgh, PA, USA
 <sup>d</sup>Bioscience Research Group. Grifols S.A., Barcelona, Spain
 <sup>e</sup>Department of Pathology and Laboratory Medicine, Dartmouth Hitchcock Medical Center, Lebanon, NH, USA



# **Patient and Site Disposition**

PV: Protocol violation





LoFU: Lost of Follow-up

# **Demographics: All Patients**



	Placebo (N=80)	Low albumin, no IVIG (N=78)	Low albumin + IVIG (N=86)	High albumin + IVIG (N=78)	Total (N=322)
Age (yrs.) Mean (SD)	68.44 (8.38)	68.47 (7.48)	69.47 (6.92)	69.54 (7.90)	68.99 (7.66)
Age Group (n, %) <65 65-75 >75	29 (36.3) 33 (41.3) 18 (22.5)	26 (33.3) 37 (47.4) 15 (19.2)	17 (19.8) 52 (60.5) 17 (19.8)	22 (28.2) 35 (44.9) 21 (26.9)	94 (29.2) 157 (48.8) 71 (22.0)
Sex (n, %) Male Female	44 (55.0) 36 (45.0)	35 (44.9) 43 (55.1)	38 (44.2) 48 (55.8)	31 (39.7) 47 (60.3)	148 (46.0) 174 (54.0)



# **Demographics: Mild (MMSE 22-26)**



	Placebo (N=44)	Low albumin, no IVIG (N=32)	Low albumin + IVIG (N=49)	High albumin + IVIG (N=36)	Total (N=161)
Age (yrs.) Mean (SD)	68.11 (7.89)	69.97 (6.36)	70.16 (6.39)	68.76 (7.91)	69.27 (7.16)
Age Group (n, %) <65 65-75 >75	15 (34.1) 19 (43.2) 10 (22.7)	8 (25.0) 17 (53.1) 7 (21.9)	8 (16.3) 31 (63.3) 10 (20.4)	11 (30.6) 17 (47.2) 8 (22.2)	42 (26.1) 84 (52.2) 35 (21.7)
Sex (n, %) Male Female	24 (54.5) 20 (45.5)	16 (50.0) 16 (50.0)	24 (49.0) 25 (51.0)	20 (55.6) 16 (44.4)	84 (52.2) 77 (47.8)



# **Demographics: Moderate (MMSE 18-21)**



	Placebo (N=35)	Low Dose, no IVIG (N=46)	Low Dose + IVIG (N=37)	High Dose + IVIG (N=42)	Total (N=160)
Age (yrs.) Mean (SD)	69.1 (1.52)	67.43 (1.19)	68.54 (1.25)	70.12 (1.22)	68.76 (0.64)
Age Group (n, %) <65 65-75 >75	13 (37.1) 14 (40.0) 8 (22.9)	18 (39.1) 20 (43.5) 8 (17.4)	9 (24.3) 21 (56.8) 7 (18.9)	11 (26.2) 18 (42.9) 13 (31.0)	51 (31.9) 73 (45.6) 36 (22.5)
Sex (n, %) Male Female	19 (54.3) 16 (45.7)	19 (41.3) 27 (58.7)	14 (37.8) 23 (62.2)	11 (26.2) 31 (73.8)	63 (39.4) 97 (60.6)





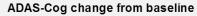
# **Primary Results: Global Cognition and Function**

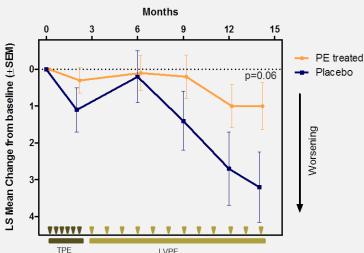


# Global Cognition (ADAS-Cog): All patients



#### **Combined arm**





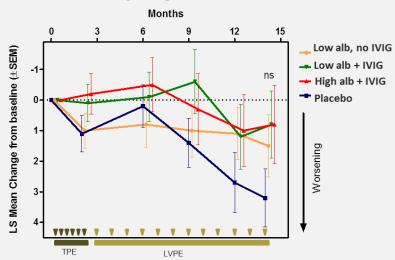
	PE treated	Placebo
Change from baseline	1.0	3.2
Diff. vs. Placebo	-2.1	-
p-value	§ 0.06	-
Less decline	66 %	-

LVPE

#### N = 322

#### **Treatment arms**

#### ADAS-Cog change from baseline



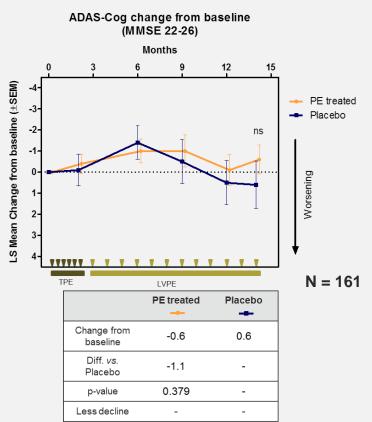
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		-	-	-
Change from baseline	1.5	0.8	0.8	3.2
Diff. <i>vs.</i> Placebo	-1.6	-2.4	-2.4	-
p-value (adj. mult. testing)	0.247	0.215	0.215	-
Less decline	50 %	75 %	75 %	-

N = 322



# Global Cognition (ADAS-Cog): Mild

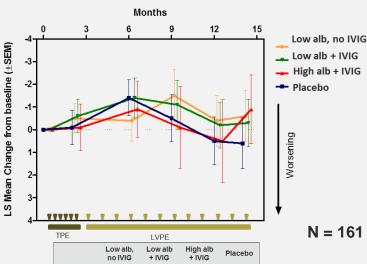
#### **Combined arm**





#### **Treatment arms**

ADAS-Cog change from baseline (MMSE 22-26)



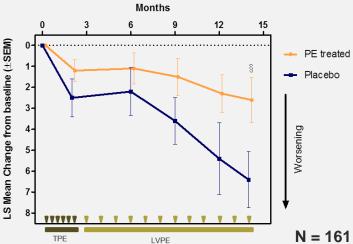
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	-0.6	-0.3	-0.9	0.6
Diff. vs. Placebo	-1.2	-0.9	-1.4	-
p-value (adj. mult. testing)	0.558	0.558	0.558	-
Less decline	-	-	-	-

# **Global Cognition (ADAS-Cog): Moderate**



#### **Combined arm**

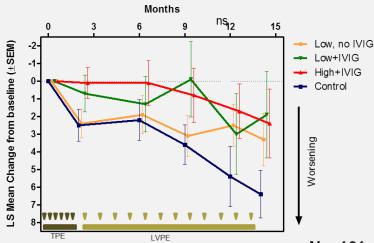




	PE treated	Placebo
Change from baseline	2.6	6.4
Diff. <i>vs.</i> Placebo	-3.9	-
p-value	§ 0.05	-
Less decline	61 %	-

#### **Treatment arms**

## ADAS-Cog change from baseline (MMSE 18-21)



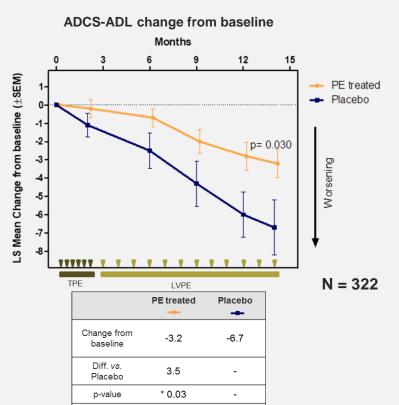
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		-	-	-
Change from baseline	3.3	1.9	2.4	6.4
Diff. vs. Placebo	-3.1	-4.5	-4.0	-
p-value (adj. mult. testing)	0.177	0.177	0.177	-
Less decline	48 %	70 %	63 %	-

N = 161

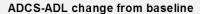
# Global Function (ADCS-ADL): All Patients

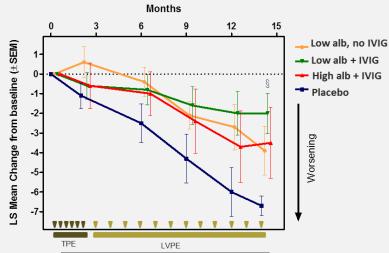


#### **Combined arm**



#### **Treatment arms**





	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	-3.9	-2.0	-3.5	-6.7
Diff. vs. Placebo	2.8	4.7	3.1	-
p-value (adj. mult. testing)	0.163	§ 0.056	0.163	-
Less decline	42 %	70 %	46 %	-



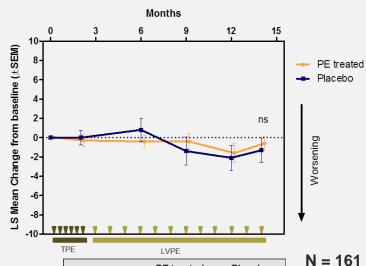
Less decline

52 %

# Global Function (ADCS-ADL): Mild

#### **Combined arm**

ADCS-ADL change from baseline (MMSE 22-26)

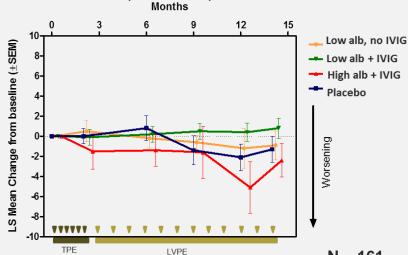


	PE treated	Placebo
Change from baseline	-0.6	-1.3
Diff. <i>vs.</i> Placebo	0.7	-
p-value	0.664	-
Less decline	-	-



#### **Treatment arms**

ADCS-ADL change from baseline (MMSE 22-26)



	Low alb,	Low alb	High alb	Placebo
	-	-	-	
Change from baseline	-0.9	0.8	-2.4	-1.3
Diff. vs. Placebo	0.3	2.1	-1.1	-
p-value (adj. mult. testing)	0.869	0.792	0.869	-
Less	-	-	-	-

N = 161

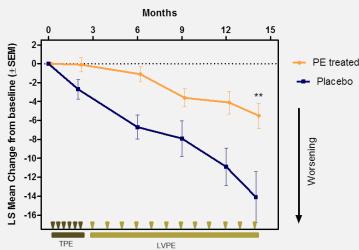


# Global Function (ADCS-ADL): Moderate



#### **Combined arm**

ADCS-ADL change from baseline (MMSE 18-21)

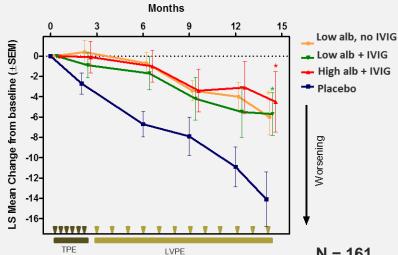


	PE treated	Placebo
Change from baseline	-5.5	-14.1
Diff. vs. Placebo	8.6	-
p-value	** 0.002	-
Less decline	61 %	-

N = 161

#### **Treatment arms**

ADCS-ADL change from baseline (MMSE 18-21)



	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		-	-	
Change from baseline	-6.0	-5.7	-4.5	-14.1
Diff. <i>vs.</i> Placebo	8.0	8.3	9.5	-
p-value (adj. mult. testing)	* 0.02	* 0.02	* 0.01	-
Less decline	57 %	59 %	67%	-

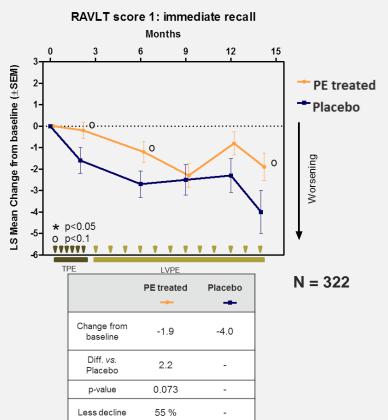
N = 161

# Secondary Clinical Endpoints



# Memory (Verbal learning): All patients

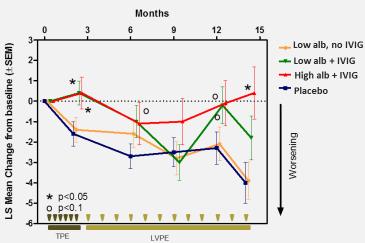
## **Combined arm**



#### alzheimer management by albumin replacement

#### **Treatment arms**

RAVLT score 1: immediate recall



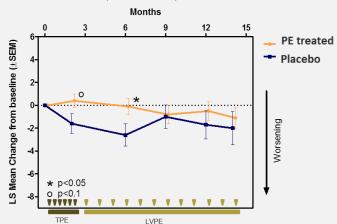
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		<del></del>		
Change from baseline	-3.9	-1.8	0.4	-4.0
Diff. <i>vs.</i> Placebo	0.2	2.2	4.4	-
p-value	0.915	0.131	0.004	-
Improv.	0 X	0.6 X	1.1 X	-



# Memory (Verbal learning): Mild

## **Combined arm**

RAVLT score 1: immediate recall (MMSE: 22-26)



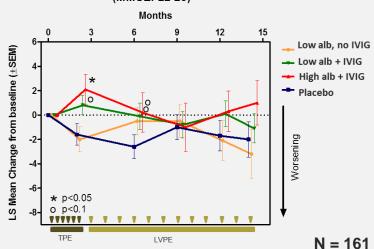
N = 161

	PE treated	Placebo
Change from baseline	-1.1	-2.0
Diff. vs. Placebo	0.9	-
p-value	0.600	-
Less decline	45 %	-



#### **Treatment arms**

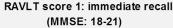
RAVLT score 1: immediate recall (MMSE: 22-26)

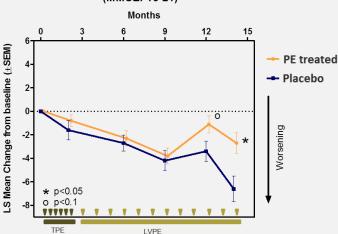


	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	-3.2	-1.1	1.0	-2.0
Diff. vs. Placebo	-1.2	0.9	3.0	-
p-value	0.596	0.663	0.194	-
Improv.	-	0.5 X	1.5 X	-

# Memory (Verbal learning): Moderate

#### **Combined arm**





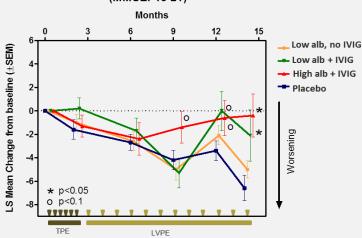
	PE treated	Placebo
Change from baseline	-2.7	-6.6
Diff. <i>vs.</i> Placebo	3.9	-
p-value	0.024	-
Less decline	59 %	-

N = 161

#### alzheimer management by albumin replacement

#### **Treatment arms**

RAVLT score 1: immediate recall (MMSE: 18-21)



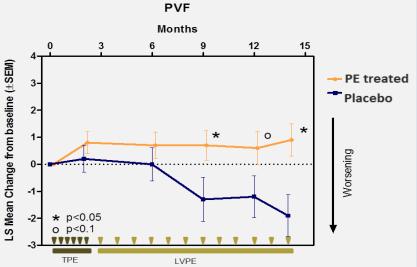
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	-5.0	-2.1	-0.4	-6.6
Diff. vs. Placebo	1.7	4.6	6.2	-
p-value	0.384	0.041	0.003	-
Less decline	26 %	70 %	94 %	-



# Language (Verbal fluency): All patients



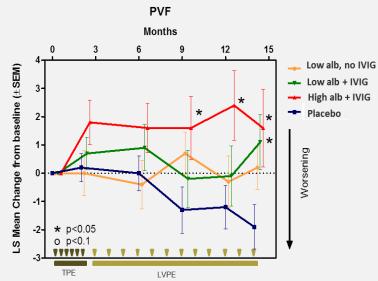
## **Combined arm**



	PE treated	Placebo
Change from baseline	0.9	-1.9
Diff. vs. Placebo	2.8	-
p-value	0.007	-
Improv.	1.5 X	-

#### N = 322

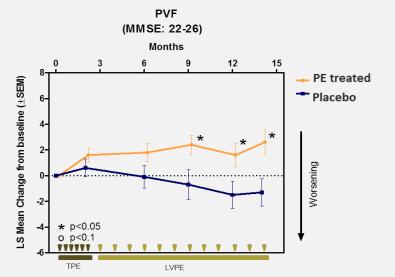
#### **Treatment arms**



	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
	-	-	-	-
Change from baseline	0.2	1.1	1.6	-1.9
Diff. vs. Placebo	2.0	3.0	3.5	-
p-value	0.106	0.022	0.008	-
Improv.	1.1 X	1.6 X	1.8 X	-

# Language (Verbal Fluency): Mild

# **Combined arm**



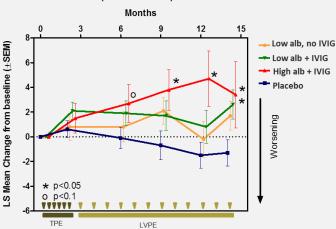
	PE treated	Placebo
Change from baseline	2.6	-1.3
Diff. vs. Placebo	3.8	-
p-value	0.013	-
Improv.	3 X	-



#### alzheimer management by albumin replacement

#### **Treatment arms**

PVF (MMSE: 22-26)

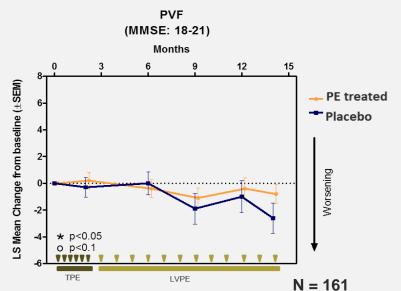


	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		-	-	
Change from baseline	1.7	2.6	3.4	-1.3
Diff. vs. Placebo	3.0	3.9	4.7	-
p-value	0.141	0.036	0.025	-
Improv.	2.3 X	3 X	3.6 X	-



# Language (Verbal Fluency): Moderate

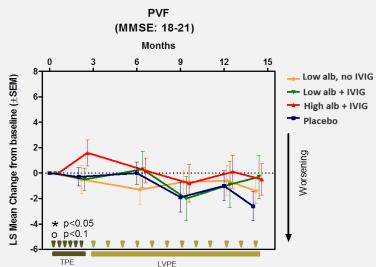
## **Combined arm**



		PE treated	Placebo
	Change from baseline	-0.8	-2.6
	Diff. <i>vs.</i> Placebo	1.7	-
	p-value	0.194	-
GRIFOLS	Less decline	65 %	-



#### **Treatment arms**

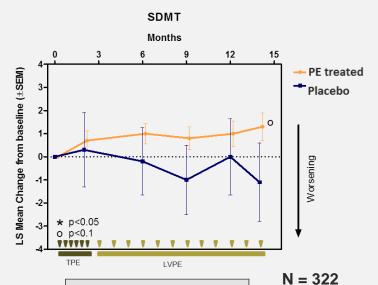


	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	-1.4	-0.3	-0.5	-2.6
Diff. vs. Placebo	1.2	2.2	2.1	-
p-value	0.441	0.213	0.200	-
Less decline	46 %	85 %	81 %	-

# **Executive Function (Processing Speed): All Patients**

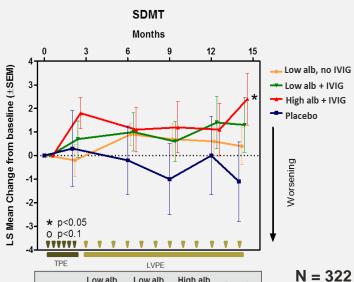


#### **Combined arm**



	PE treated	Placebo
Change from baseline	1.3	-1.1
Diff. vs. Placebo	2.4	-
p-value	0.054	-
Improv.	2.2 X	-

#### **Treatment arms**



	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
				-
Change from baseline	0.4	1.3	2.4	-1.1
Diff. vs. Placebo	1.5	2.4	3.5	-
p-value	0.313	0.119	0.030	-
Improv.	1.4 X	2.2 X	3.2 X	-

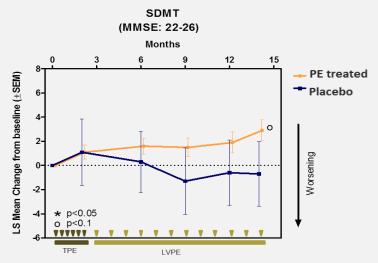




# **Executive Function (Processing Speed): Mild**



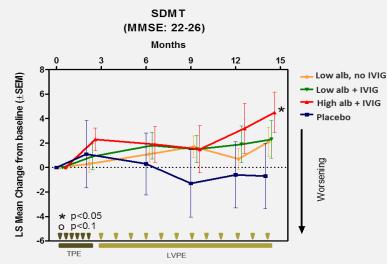
#### **Combined arm**



	PE treated	Placebo
Change from baseline	2.9	-0.7
Diff. <i>vs.</i> Placebo	3.6	-
p-value	0.055	-
Improv.	5.1 X	-

N = 160

#### **Treatment arms**



	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
		<del></del>		
Change from baseline	2.1	2.3	4.5	-0.7
Diff. <i>vs.</i> Placebo	2.9	3.0	5.2	-
p-value	0.244	0.173	0.039	-
Improv.	4.1 X	4.3 X	7.4 X	-

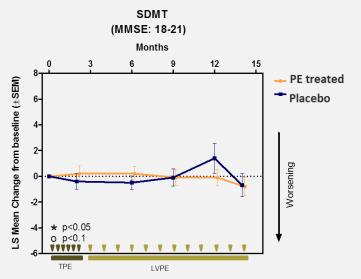




# **Executive Function (Processing Speed): Moderate**



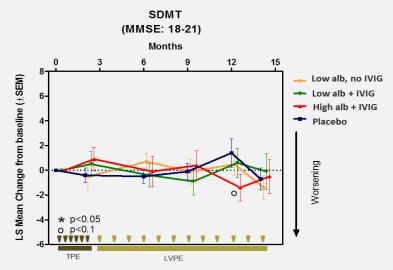
#### **Combined arm**



	PE treated	Placebo
Change from baseline	-0.8	-0.7
Diff. vs. Placebo	-0.1	-
p-value	0.931	-
Less decline	-	-

#### N = 151

#### **Treatment arms**



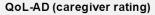
	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
	-			
Change from baseline	-1.4	-0.1	-0.5	-0.7
Diff. <i>vs.</i> Placebo	-0.7	0.6	0.2	-
p-value	0.658	0.757	0.910	-
Less decline	-	86 %	29 %	-

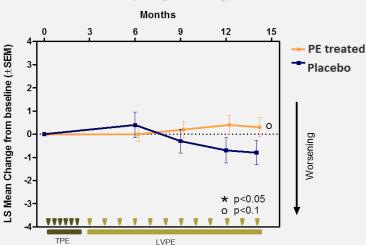


# **Quality of Life (Caregiver Rating): All Patients**



#### **Combined arm**



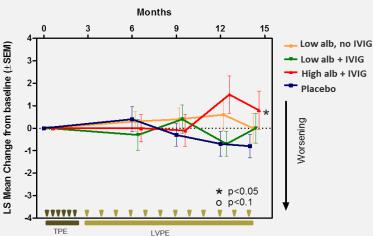


	PE treated	Placebo
Change from baseline	0.3	-0.8
Diff. <i>vs.</i> Placebo	1.1	-
p-value	0.07	-
Improv.	1.4 X	-

N = 322

#### **Treatment arms**

#### QoL-AD (caregiver rating)

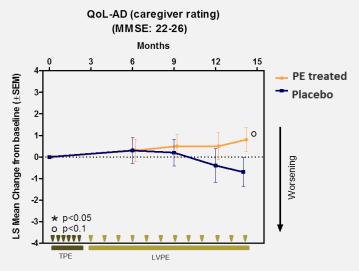


	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	0.0	0.0	0.8	-0.8
Diff. <i>vs.</i> Placebo	0.8	0.9	1.6	-
p-value	0.280	0.288	0.050	-
Improv.	-	-	2 X	-



# **Quality of Life (Caregiver Rating): Mild**

#### **Combined arm**



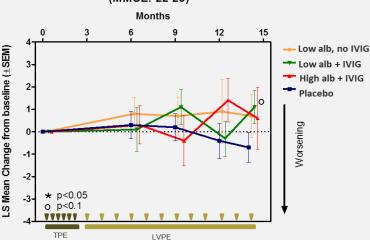
	PE treated	Placebo
Change from baseline	0.8	-0.7
Diff. <i>vs.</i> Placebo	1.5	-
p-value	0.073	-
Improv.	2.1 X	-

#### N = 156

#### alzheimer management by albumin replacement

#### **Treatment arms**

QoL-AD (caregiver rating) (MMSE: 22-26)



	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
Change from baseline	0.7	1.1	0.6	-0.7
Diff. <i>vs.</i> Placebo	1.4	1.8	1.3	-
p-value	0.204	0.085	0.264	-
Improv.	2 X	2.6 X	1.9 X	-

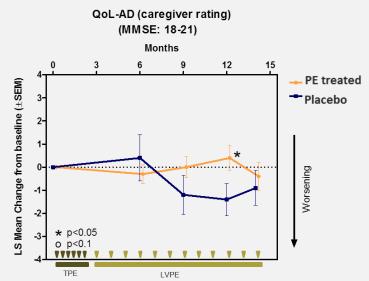


# **Quality of Life (Caregiver Rating): Moderate**

N = 156

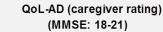


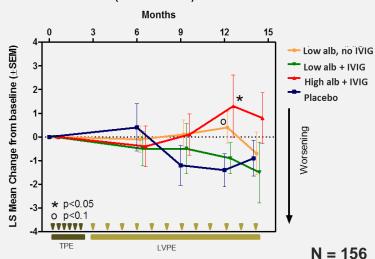
### **Combined arm**



	PE treated	Placebo
Change from baseline	-0.4	-0.9
Diff. vs. Placebo	0.5	-
p-value	0.639	-
Less decline	56 %	-

# Treatment arms





	Low alb, no IVIG	Low alb + IVIG	High alb + IVIG	Placebo
	-	-	-	-
Change from baseline	-0.7	-1.5	0.8	-0.9
Diff. <i>vs.</i> Placebo	0.2	-0.6	1.6	-
p-value	0.857	0.644	0.181	-
Improv.	0.2 X	-	1.8 X	-



# **Other Clinical Endpoints**



#### Analyses ongoing:

- NAB (NAB Naming Test)
- CSDD (Cornell Scale for Depression in Dementia)
- NPI (Neuropsychiatric Inventory)
- CDR-Sb (Clinical Dementia Rating, Sum of boxes)
- ADCS-CGIC (Clinical Global Impression of Change)
- C-SSRS (Columbia Suicide Severity Rating Scale)
- Neuroimaging analyses (MRI and PET)

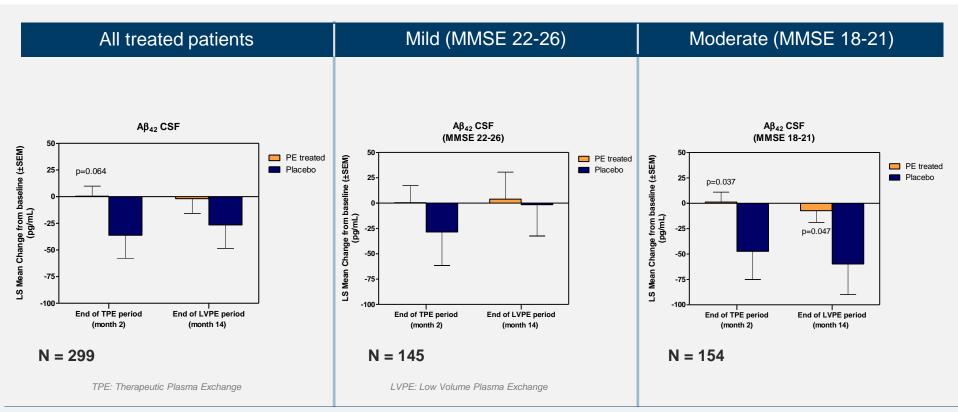


# **CSF Biomarkers** Aβ<sub>42</sub>, Tau and P-Tau



# $CSFA\beta_{42}$

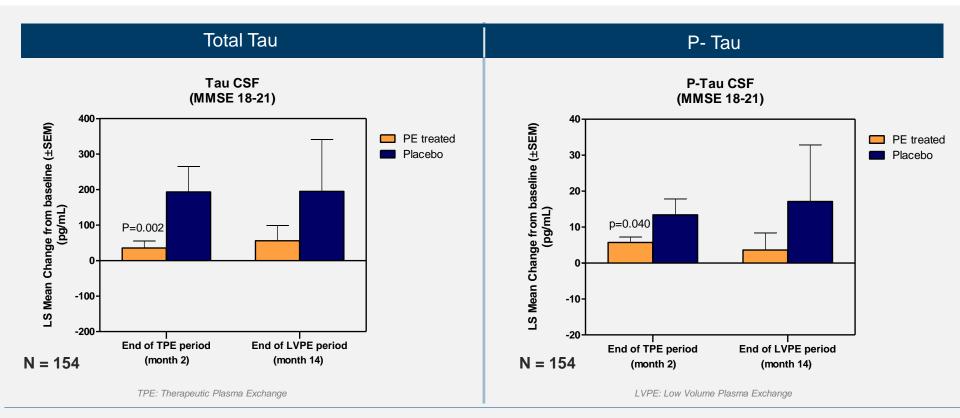






# **CSF Tau: Moderate (MMSE 18-21)**









# Safety Adverse Events (AEs)



# **Most Frequent AEs Related with PE**



	Total	Placebo	Low, no IVIG	Low + IVIG	High + IVIG
Total PE	4709	1223	1207	1180	1099
TPE LVPE	1718 2991	435 788	430 777	448 732	405 694
	PEs with AE, n (% of PE)	PEs with AE, n (% of PE)			
Anaemia	43 (0.9)	2 (0.2)	13 (1.1)	16 (1.4)	12 (1.1)
Catheter local reactions	119 (2.5)	0	43 (3.6)	31 (2.6)	45 (4.1)
Catheter / Device infection	15 (0.3)	0	7 (0.6)	5 (0.4)	3 (0.3)
Muscle spasms	49 (1.0)	0	16 (1.3)	4 (0.3)	29 (2.6)
Dizziness	30 (0.6)	1 (0.1)	8 (0.7)	13 (1.1)	8 (0.7)
Headache	11 (0.2)	3 (0.2)	4 (0.3)	2 (0.2)	2 (0.2)
Paraesthesia	28 (0.6)	0	16 (1.3)	1 (0.1)	11 (1.0)
Presyncope	28 (0.6)	1 (0.1)	8 (0.7)	14 (1.2)	5 (0.5)
Syncope	11 (0.2)	0	4 (0.3)	3 (0.3)	4 (0.4)
Hypotension	102 (2.2)	0	37 (3.1)	37 (3.1)	28 (2.5)



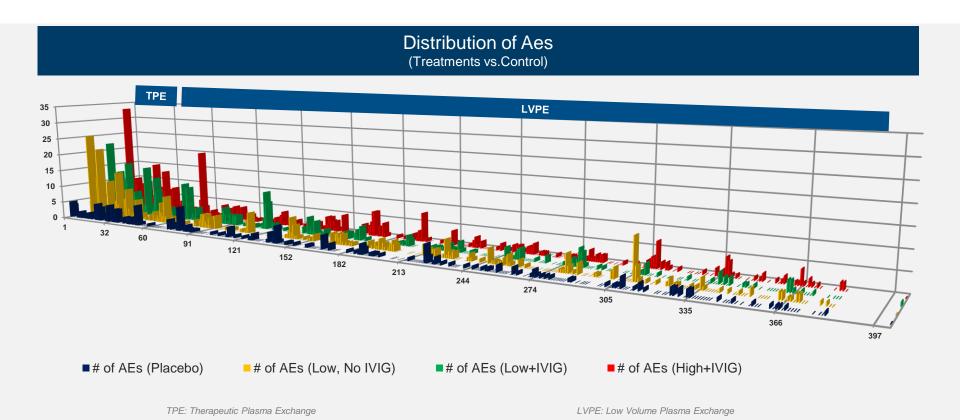
# **Infections**



	Total (N=322)	Placebo (N=79)	Low, no IVIG (N=78)	Low + IVIG (N=86)	High + IVIG (N=79)
Total PE TPE LVPE	4709 1718 2991	1223 435 788	1207 430 777	1180 448 732	1099 405 694
Total of Infections	159	38	52	37	32
Patients with infections	147	33	49	34	31
% of patients with infections	N/A	41.8	62.8	39.5	39.2
Infections per 100 patients	N/A	48.1	66.7	43.0	40.5



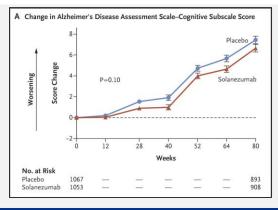
# **AEs: "Manhattan" Chart**

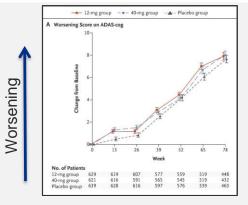


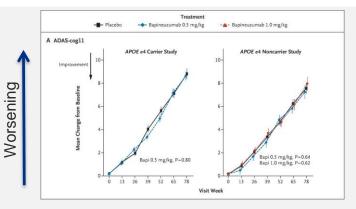


# Recent BACE and Mab Programs Discontinued due to Lack of Efficacy vs. AMBAR: ADAS-Cog

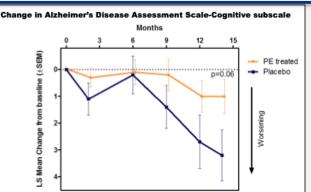






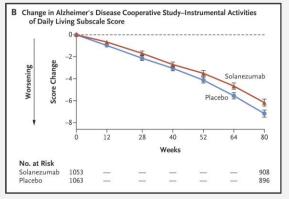


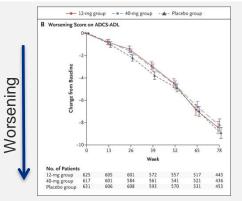
Morsening Workening

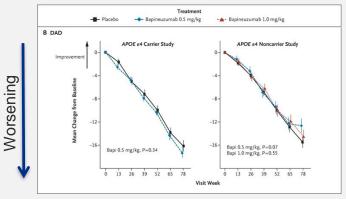


# Recent BACE and Mab Programs Discontinued due to Lack of Efficacy vs. AMBAR: ADCS-ADL



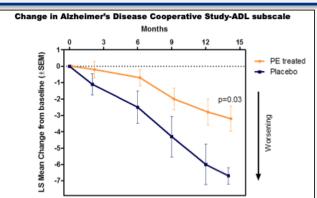






**AMBAR** 





# **Key Messages: Efficacy**



#### Primary endpoints:

- Mild AD: No decline, neither active nor placebo
- Moderate AD: 61% statistically significant less decline in both ADAS-Cog and ADCS-ADL of treated patients as compared with placebo at 14 months All three treatment arms statistically different from placebo (ADCS-ADL)

#### Secondary endpoints:

- All-patient analysis: statistically significant <u>improvement</u> in memory, language, processing speed and QoL (caregiver) of High-Albumin+IVIG arm compared with placebo at 14 months
- Mild AD: statistically significant <u>improvement</u> in language and processing speed and of High-Albumin+IVIG arm Borderline significance in QoL (caregiver)
- Moderate AD: statistically significant <u>improvement</u> in memory and QoL (caregiver) of High-Albumin+IVIG arm



# **Key Messages: Safety and Feasibility**



- Feasible: 4,709 procedures
  - 1,223 sham; 3,486 real
  - 1,718 TPE; 2,991 LVPE
- 72 % of patients completed the study
- · AE rate seems to depend on volume infused and IVIG dose, as expected
- Low rate of plasma exchange procedures related with AEs
- **Safety conclusion**: PE, both TPE and LVPE, was safe, well tolerated and feasible in mild-to-moderate AD patients, with a 72% of patients completing the study

# **Key Messages: Biomarkers and Infections**



#### Biomarkers:

- CSF Aβ<sub>42</sub>: stabilization in the treated patients and decline in placebo arm particularly for moderate AD
- <u>CSF Tau and P-Tau</u>: less increase in the treated patients compared to placebo particularly for moderate AD

#### Infections:

- Patients treated with plasmapheresis without IVIG had more infections than those treated with IVIG and also than those in the placebo arm
- The rate of infections not related with catheter was lower in patients receiving IVIG



# We are Profoundly Grateful to the Patients and Families that Have Kindly Participated in the AMBAR Clinical Trial



# Thanks to the Pioneers...





IV CONGRESSO INTERNACIONAL DE TRANSFUSÕES DE SANGUE

III CURSO INTERNACIONAL DE MEDICINA E CIRURGIA DE URGÊNCIA

I EXPOSIÇÃO MUNDIAL DO SANGUE

I COLÓQUIO DE HEMATOLOGIA AFRICANA



#### Dr. E. DE MORAIS & T. DE SOUSA SOBRINHO (Porto)

Deux cas de purpura thrombopénique guéris par transfusion sanguine intrasternal

Two cases of thrombocytopenic purpura cured by intra sternal blood transfusion

Dois casos de purpura trombopénica curados por hemotrans-

#### Drs. A. LESSA & A. DE SOUSA (Lisboa)

Demonstration clinique et radiologique des possibilités de transfusion par voie osseuse

Clinical and roentgenological demonstration of the possibilities

of transfusion in the bone-marrow Demonstração clínica e radiológica das possibilidades da via

Dr. I. MASSONS & W. OPPENHEIMER (Barcelona)

La plasmatherapie heterologue en Pédiatrie Heterologous plasma transfusion, in Pedriatrics A plasmoterapia heterologa em Pediatria

#### Dr. A. P. FRANCO (Lisboa)

Étude hemathologique d'un cas de maladie hemolytique du nouveau-né, à la première grossesse, du à l'incompatibilité Rh Haematological study of a case of Haemolytic Disease of the new-born in the first pregnancy, due to the Rh incompatibility Estudo hematológico dum caso de doença hemolítica de recém-nascido, devido à incompatibilidade Rh em primeira

#### Dr. J. A. GRIFFOLS-LUCAS (Barcelona)

La plasmaphorèse chez l'homme, avec un rapport sur la pre-

Plasmophoresis in man. Report of the first series of cases Plasmoforese no homem, Notícia da primeira série de casos

#### Dr. J. WILLENEGGER (Basiléia)

Nouvelles recherches sur le donneur universel dit dangereux New researches on the universal blood donor known as dan-

Novas pesquizas sobre o dador universal considerado perigoso

- 18 -





# Bioscience R&D & Clayton Facilities Tours **Todd Willis & Daniel Fleta GRIFOLS**

# **Bioscience R&D Facilities Tour**



# Bioscience R&D - RTP

# Facilities and Organization



#### 30,000 ft<sup>2</sup> state-of-the art R&D facility

- Open lab concept to maximize space
- Specialized areas for bench-scale and macrobench processing, filling suite, and cell culture rooms
- BSL-3 facility and containment practices for handling Risk Group 3 pathogens

- 90 scientists and support staff
  - Product Development
    - Bioanalytics
    - Process Development
  - Product Safety
    - Toxicology
    - Pathogen Safety
  - Discovery Research
  - Quality R&D





# Safety Guidelines

We will split into 4 groups to facilitate the tour. Please stay with your group

When walking on an area with metal grading, please stay on the designated pathway Watch your step,
especially on
transitions
between
different
surfaces

Please do not touch anything as many of the systems are energized

# Please feel free to ask questions



#### **Tour Route**

# **New Fractionation Building:**

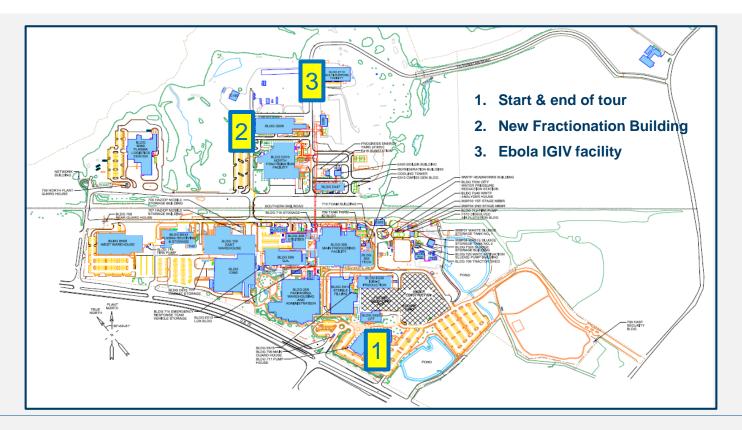
- 6MM Liter plasma / year capacity
- Construction complete during 1Q 2019, start up in progress
- Features latest version of Automatic Bottle Opener ABO<sub>6</sub>

# Ebola IGIV facility:

- Totally isolated facility for processing plasma to final container
- Batch size is 75L of plasma



# **Tour Map**





# **New Fractionation Building General Information**



- Surface: 85,000 sq.ft. building area
- Capacity: 6MM Liters plasma/ yr capacity
- Execution: Two production trains with 16 vessels per train and eight buffer vessels
- Floors: Two production floors with an interstitial areas above each floor

#### New Fractionation Building General Information











- Most recent version of Grifols
   Engineering Automatic Bottle Opener
  - Automated de-palletizing, loading and repalletizing
  - Includes RFID tracking capability
  - Throughput rate of 1,250 bottles/hour
- Reactors integration in the clean room.
   Grifols patented method for installation of vessels in the clean room
- Process rooms with daylight for operators working environment convenience
- Filter presses with CIP system built-in
- Central buffer preparation area for both trains



Convalescent Plasma Immunoglobulin Facility





#### Convalescent Plasma Immunoglobulin Facility

- Design and construction of a Plasma Fractionation and IGIV purification facility + filling and conditioning segregated
- Batch Size 75 Liters in bags of aprox.500ml plasma from donors with anti-ebola antibodies
- 300gr Gamunex<sup>®</sup> will be manufactured by batch
- Production capacity of up to 3 batched per week
- IGIV filling at 10% using Gri-Fill® technology in 100 ml bags (30 bags)
- The facility will also have labelling, packaging and warehouse previous to sending

#### Convalescent Plasma Immunoglobulin Facility

#### **Plant**

Total Structure area: 1,300m<sup>2</sup>

Clean Rooms, QC Lab & Lockers: 427m<sup>2</sup>

Total Clean Rooms 241m<sup>2</sup>

52 tons structural steel

#### **Equipment**

- 4 Units HVAC + 1 Steam generator
- 2 Chillers(5 C and -25 C°)
- 1 Plant water treatment & Osmosis
- 1 Autoclave + 1 Liquid sterilizer HWFI loop/tank

The construction started in January 2015 and finished in October 2015





## **Investor and Analyst Meeting**

June 5-6, 2019

1:00-2:00

Lunch

7	une 5 – Clayton, NC	Julie 5 - Glaytoll, NC		Julie 0 - Kaleigii, NC	
7:30	Pick-up from recommended hotels	2:00-2:45	Novel Plasma Therapies Development  T. Willis	7:30	Pick-up from recommended hotels
8:30-9:00	Registration and welcome	2:45-3:30	AMBAR: Grifols' Alzheimer Trial A. Paez	8:30-9:30	R&D Tour
9:00-9:30	Introduction R. Griffols	3:30-4:00	Break	9:30-10:15	China: Facing the Opportunity A. Martinez
9:30-11:30	Commercial Strategies  L. Morgan/J. Abelson/ C. Schroeder/R. Jagt	4:00-4:30	Q&A	10:15-10:45	Digital Innovation  X. Sueiras
11:30-12:00	Break	4:30-5:00	Tour Introductions	10:45-11:15	Break
12:00-1:00	Industrial Capacity and Plasma Capabilities  P. Allen/E. Herrero/D. Fleta	5:00-6:30	Site Tour: New Fractionation Building and Ebola plant	11:15-12:00	Financials A. Arroyo

Back to recommended hotels

Dinner

7:00

10:00

June 5 - Clayton NC

June 6 - Raleigh NC

**Grifols: A Socially Responsible** 

T. Rione

V. Grífols Deu

Company

Closing

Q&A

Lunch

12:00-12:30

12:30-12:45

12:45-1:15

1:15

June 5 - Clayton NC

## **China: Facing the Opportunity**

**Grifols Next Growth Engine** 

Amarant Martinez 马敏伟 VP, China Affairs Office 中国事务办公室副总裁

**GRIFOLS** 

# 1. China and its Healthcare Market: Key Figures

#### China is a Strategic Market for Grifols

By 2030, China is Expected to Become #1 Economy in the World<sup>5</sup>

*In* 2018...

**1,415** Million people<sup>1</sup>

**15.8%** of World Economy<sup>1</sup>

**18.5%** of World population<sup>1</sup>

**+6.6%** GDP growth 2018<sup>1</sup>

In the World...

55% of ALBUMIN market<sup>2</sup>10% of IVIG market<sup>2</sup>5% of pdFVIII market<sup>2</sup>

**12.0M** of blood donations NAT-tested<sup>3</sup> USD310M IH IVD testing market size<sup>4</sup>

Sources: 1Fitch Solutions;

2 Global Plasma Industry Database 2017 (values); 3 NIFDC 2018; 4 InterChina survey 2017; 5 HSBC

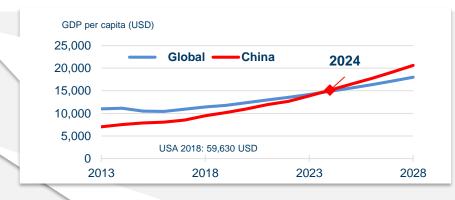


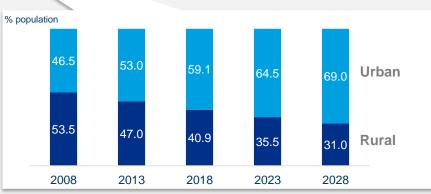
### China's Healthcare Market Fundamentals are Strong

#### Long-Term Sustained Growth and Healthcare Demand

- China's economic development
- Continued expansion of China's healthcare system and medical insurance: Health China 2030
- Raising urbanization levels: 994M in 2028
- Aging population & chronic disease burden



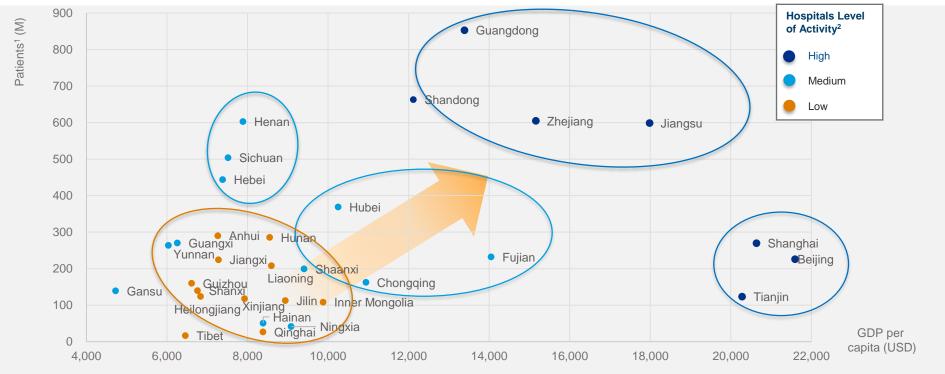




Source: Fitch Solutions

#### **China's Provincial Healthcare Landscape**

Clusters: Provinces at Growth Stage Offer Opportunities to Expand into

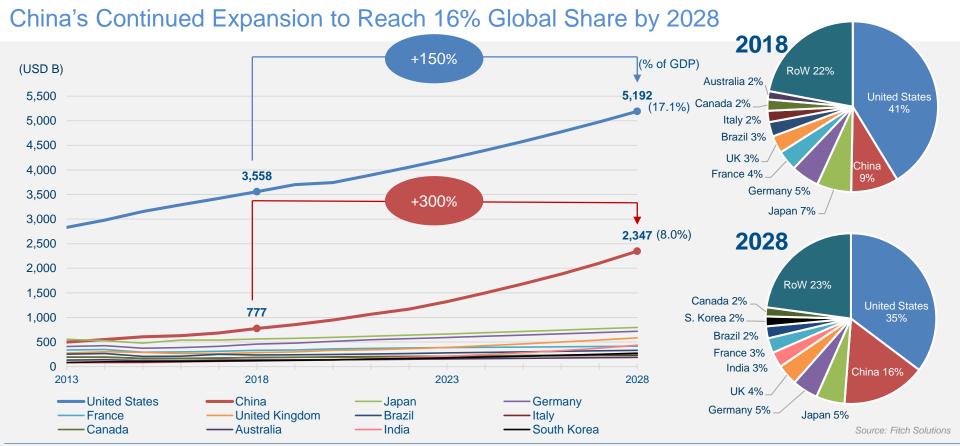


Sources: National Health Commission Year Book 2018; Fitch Solutions

1 Patients includes out-patient visits & in-patient admissions; 2 Hospitals Level of Activity = Patients / Population



## **Global Health Expenditure**





### China as a Leader in Healthcare Digital Transformation

Digital Transformation is a Key Pillar of Grifols Strategy

- ~40% physicians have used online consultation
- 20 M active users and >700 k registered physicians on top online consultation platforms
- 1.5 M physicians active on top 3 online platforms
- 63% vs 37%: Innovative channels\* have overtaken traditional channels when obtaining professional information

- 3.0 M patients making appointment on top 3 platforms
- 656 k patients buying drugs on top 3 B2C platforms
- 15.6 M patients using largest online consultation platform



## 2. Grifols Long-Term Commitment to China

#### **China is Going West and Beyond**

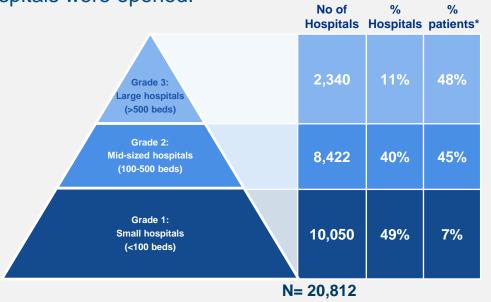
#### The Belt and Road Initiative (BRI) was Launched in 2013

- Foreign Policy
- 60 countries
- 65% world's population
- 40% Global GDP (2017)
- USD 80 B investment
- USD 6.47 Trillion goods' trade
- Trade routes
- Infrastructure



## **Continued Reform of the Healthcare System to Improve Access to Care**

 Only from Aug 2013 to Nov 2018, 783 new Grade III Hospitals were opened!



Source: National Health Commission Year Book, 2018 \*consolidated figure in-patient and out-patient



National Reimbursement Drug List (NRDL)

National Supplementary Drug List (NSDL)

**Volume Based Procurement** (4+7; 31 drugs)

**GPOs** (Guangzhou pilot)

**DRGs (Diagnosed Related Groups)** 

**Tiered Care System** 

Rational Drug Use / Medicine Proportion

**Prescription Outflow** 

**Direct to Patients (DTP) Pharmacies** 

Zero mark-up

**Two-Invoice System** 



#### Major Advancements in the 2017 NRDL vs. 2009

Dynamic RDL Adjustment is Next, from 2020



Drug	Class		Indication			
	2009	2017	2009	2017		
Albumin	В	В	Emergency treatment; Industrial insurance	Emergency treatment; Critical care; hypoalbuminemia caused by cirrhosis, cancer or hydrothorax and ascites, albumin level lower than 30g/L		
IVIG	В	В	Children's severe virus infection; Industrial insurance	Primary Immune Globulin Deficiency; Septicemia of Newborn; Severe Primary Immune Thrombocytopenia; Kawasaki Disease; Generalized Myasthenia Gravis; Acute Guillain-Barre Syndrome		
IMIG	В	В	Not indicated	Measles; Preventive treatment of infectious hepatitis		
pdFVIII	A (EDL*)	A (EDL)	Prevention and treatment Hemophilia A	Prevention and treatment Hemophilia A		
rFVIII	В	В	When hemophiliac presents severe bleeding, and pdFVIII is not available	Paediatric haemophilia A; for adult haemophilia A has bleeding		
rFIX	Not included	В	Not indicated	Paediatric haemophilia B; for adult haemophilia B has bleeding		
PTC	В	В	Surgical bleeding; bleeding caused by cirrhosis or liver necrosis	Surgical bleeding; or bleeding caused by liver diseases; Haemophilia B; Haemophiliac has FVIII inhibitor		
Fibrinogen	В	В	Emergency treatment of hypofibrinogenemia	Active bleeding caused by hypofibrinogenemia		
Rabies IG	В	В				
Tetanus IG	В	В		In 2018, Tetanus IG was included in the Essential Drug List		

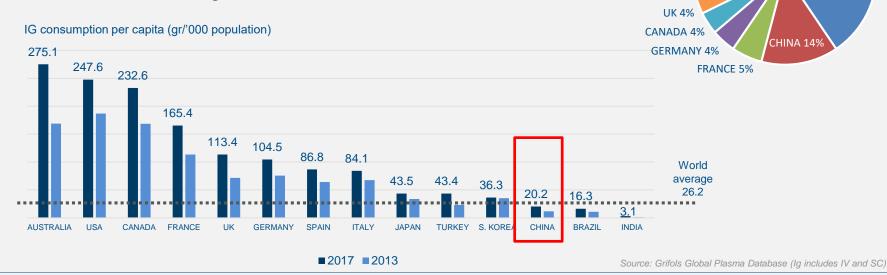
\*EDL: Essential Drug List, fully reimbursed



#### **Development of IG Therapies in China**

China Stands as the 2nd Worldwide IG Market but Use per Capita Remains Low

- Access to therapies
- Demographic development and clinical use per capita
- Diagnosis and awareness of treatment options and indications
- Chronic conditions regular treatment





**RoW 18%** 

USA 40%

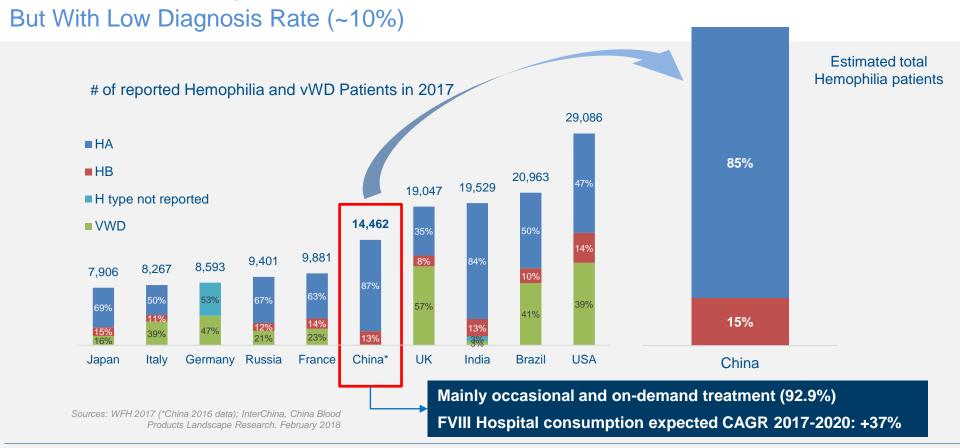
SPAIN 2%

**ITALY 3%** 

JAPAN 3%

**AUSTRALIA 3%** 

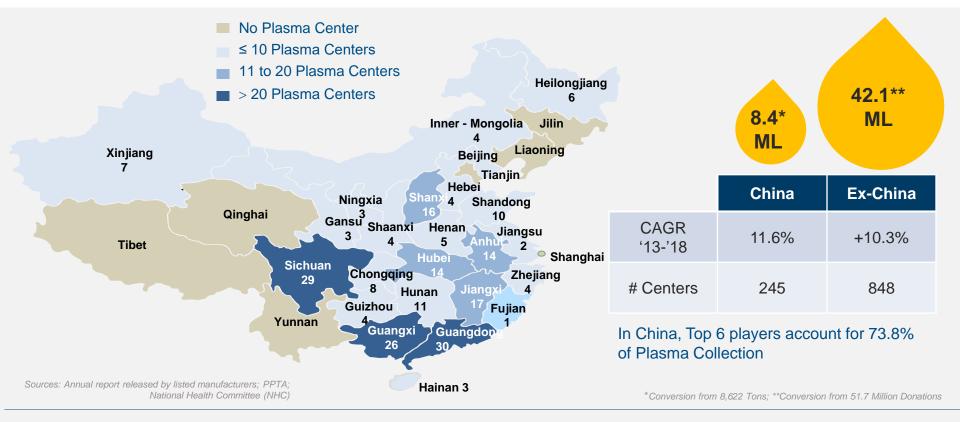
## China has a Large Hemophilia Population





### Plasma Procurement Landscape in 2018

#### We Expect Plasma Collection in China to Continue Expanding



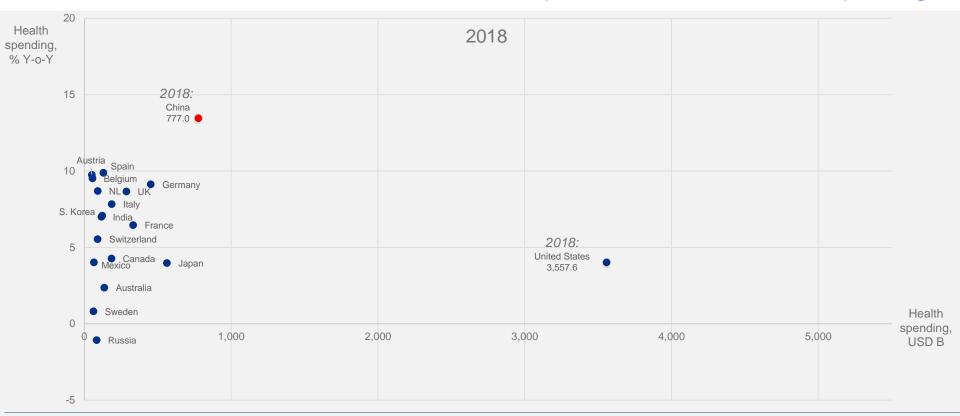


# 3. A Unique Opportunity for Grifols Across Bioscience and Diagnostic



### **Unparalleled Opportunity In an Untapped Market**

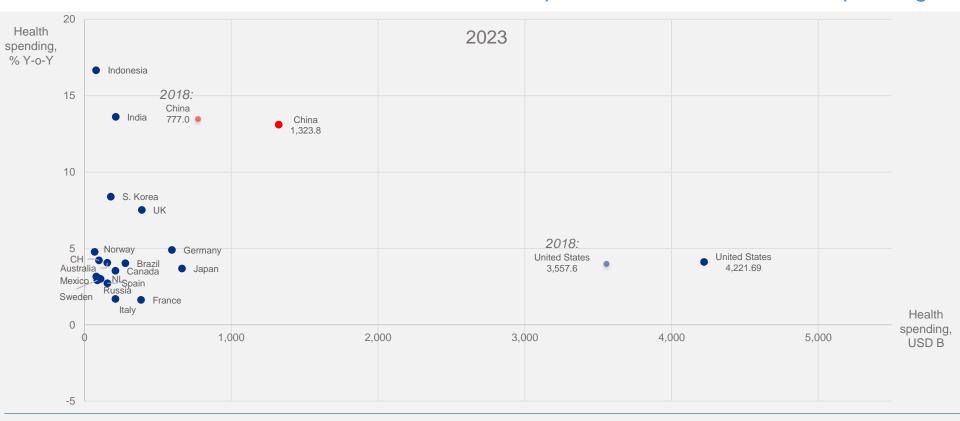
2018-2028 China and U.S. at the Forefront with Equal Contribution to Health Spending





## **Unparalleled Opportunity In an Untapped Market**

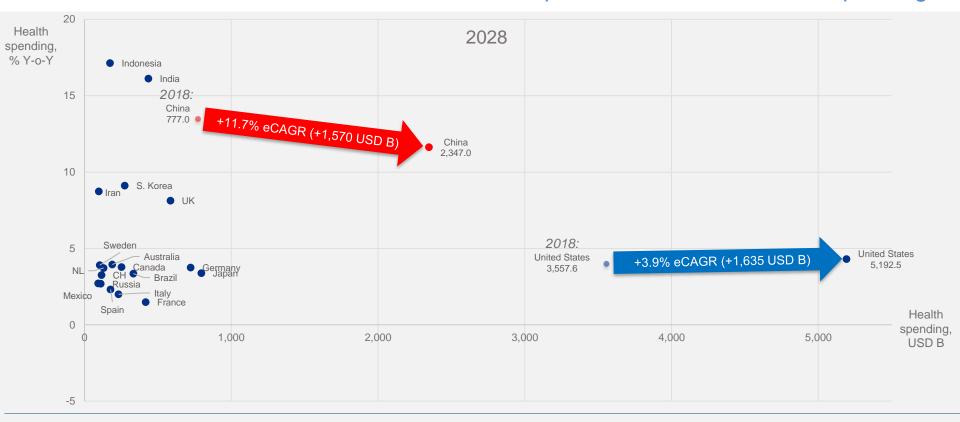
2018-2028 China and U.S. at the Forefront with Equal Contribution to Health Spending





### **Unparalleled Opportunity In an Untapped Market**

2018-2028 China and U.S. at the Forefront with Equal Contribution to Health Spending





## The Plasma Market in 2017, Top 30 Countries

Grifols Currently Can Only Participate in China in the Albumin Business

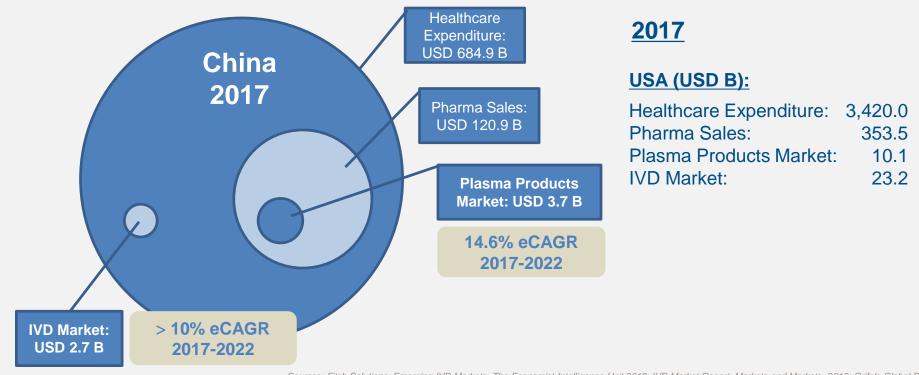


\*USA, China and Germany capped at USD 1.0 B Source: Grifols Global Plasma Database



#### **China's Healthcare Market**

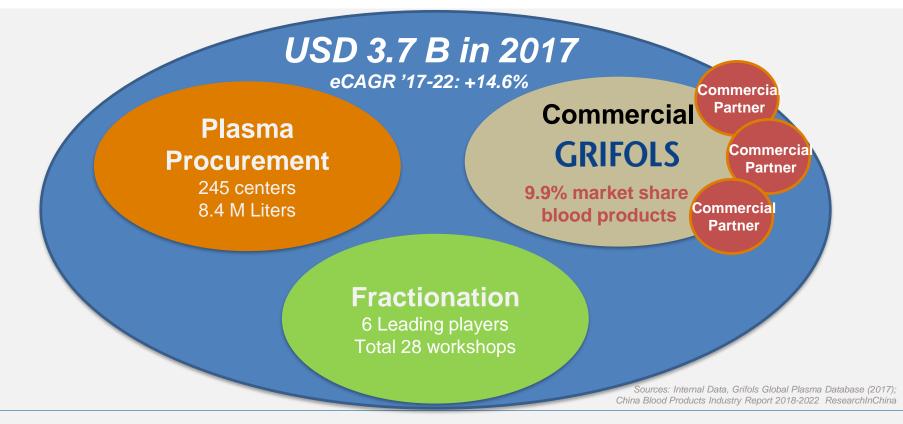
Double Digit Growth in Both Plasma and IVD (2017-2022)



Sources: Fitch Solutions; Emerging IVD Markets, The Economist Intelligence Unit 2018; IVD Market Report, Markets and Markets, 2016; Grifols Global Plasma Database (2017); China Blood Products Industry Report 2018-2022, ResearchInChina.

## **China Plasma Space**

Grifols to Contribute to the Whole Plasma Value Chain

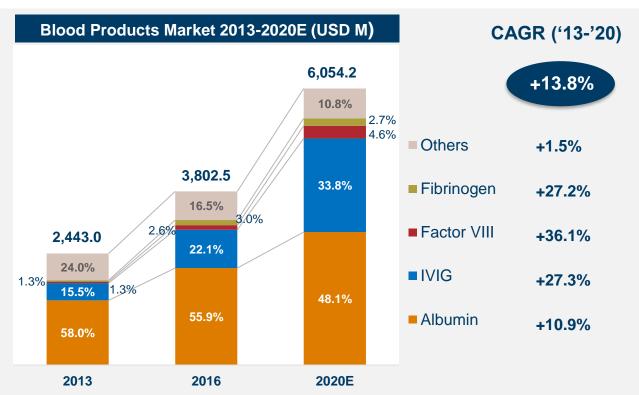




## **Commercial Opportunity: Bioscience**

High Potential of Coagulation and IVIG Products

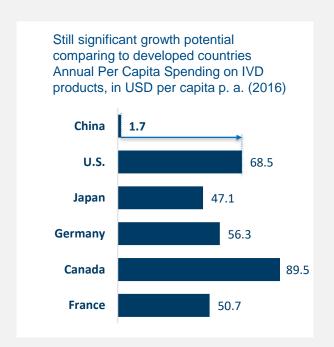
- Increased awareness and access to care
- Expanded NRDL implementation
- Hemophilia diagnosis and prophylaxis treatment
- IVIG: Neurology & Rheumatology areas

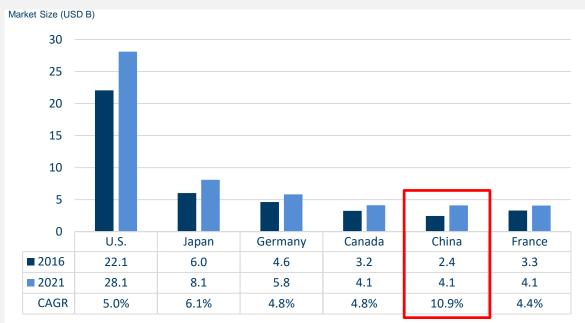


Source: InterChina, China Blood Products Landscape Research. February 2018

#### **China IVD Market**

#### China is the Fastest Growing IVD Market and Still with Great Potential





Sources: IVD Market Report, Markets and Markets, 2016; Fitch Solutions; Internal analysis



#### The Global Leader in NAT Blood Donor Screening

Significant Business Opportunity from 2020



- China is the fastest growing region in the plasma industry
- Full NAT mandate on plasma donations from 2020 will add +15.5 M plasma donations to be tested
- Viral testing (HAV, Parvo B19) and Emerging pathogens (HEV, Zika and Babesia)

Source: Q1 2017 Internal Data. It does not include plasma collection



## 4. Shanghai RAAS: The Right Partner

#### **Shanghai RAAS**

#### The Right Partner

- 2nd company in plasma collection volume in China
- 41 plasma centers across 11 provinces
- 1993, Shanghai RAAS becomes licensee of New York Blood center's S/D virus inactivation technology
- First company in China to adopt NAT testing, since 1995. NAT performed at three different times from the collection to the final product
- Shanghai RAAS and Grifols operations are truly complementary in China
- High potential for value creation







Source: Company's Annual Reports; Internal Analysis



 $\checkmark$ 

8

 $\checkmark$ 

 $\checkmark$ 

 $\checkmark$ 

 $\checkmark$ 

9

6

 $\checkmark$ 

6

 $\checkmark$ 

7

\*CBPO: China Biologic Products, including Shandong Taibang and Guizhou Taibang, CNBG: China National Biotechnology Group, including Shandani, Wuhan, Lanzhou, Guizhou (former Guizhou Zhongtai) Institutes and Rongsheng, RAAS: Shanghai RAAS, Tonrol, Zhengzhou RAAS and Zhejiang Haikang.

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✓

14



**Human Albumin** 

Freeze-dried IVIG

Lyophilized Hepatitis B IG Intravenous Hepatitis B IG

**Human Coagulation FVIII** 

**Lyophilized Thrombin** 

**Human Prothrombin Complex** 

**Total N Products** 

**Hepatitis B IG** 

Rabies IG

Tetanus IG

**Fibrinogen** 

Fibrin Sealant

Histamine IG

IVIG

IMIG

## **5. Execution is Key:** Creation of China Affairs Office

CFDA approval

for Human

1988

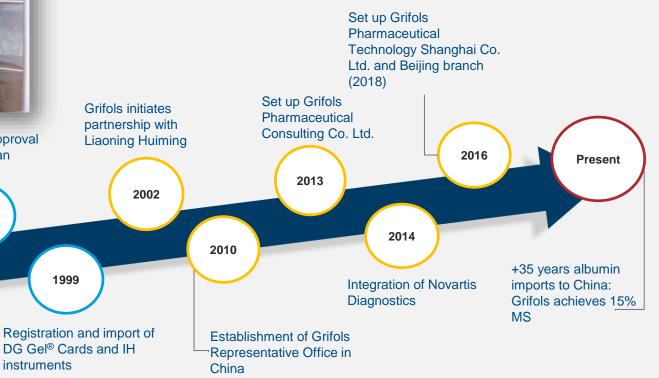
1985

China issues

Article 49

**Albumin** 

## Grifols in China: 35+ Years of History



Lucas on a commercial visit to China 1984

First import of

Intramuscular

Immunoglobulin

Dr. Victor Grífols

Start Human

in China

1984

Albumin sales

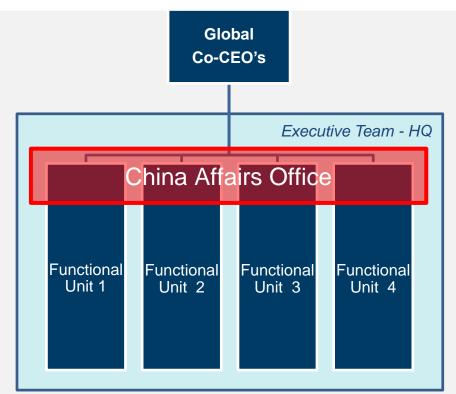


1983

#### **China Affairs Office**

Grifols is Ready and Has Its Own Model to Ensure Success

- · Creation of the China Affairs Office
- Reporting to the Company's CEO's
- Global alignment and execution under ONE Grifols concept
- Collaborative and agile
- The right people with the right experience in China since 1983
- Long-term focus





#### **Grifols in China**

#### With the Right Partner

- Bring innovative therapies to the Chinese patients
- World-class medical education in chronic and rare diseases
- Grifols Engineering expertise in biopharmaceutical engineering and consultancy to develop best-in-class industrial facilities
- Vertical integration with our Grifols NAT systems and broader virus detection platform
- With Grifols' partner, creation of a leading Industrial and Commercial platform, realizing the full potential of the Alliance and setting the ground for future expansion



## Key Takeaways China: Facing the Opportunity



#### **Key Takeaways**

China: Facing the Opportunity

- China as a driver of the worldwide healthcare expenditure with equal contribution to growth as the U.S. Strong healthcare fundamentals and continuous expansion of access to care
- China offers unparalleled opportunities across Bioscience and Diagnostic to become the next growth engine for Grifols
- Grifols has a long-term commitment to China
- Shanghai RAAS: the right partner
- Execution is the Key: Grifols is ready and has its own model to ensure success and capture the significant opportunity



#### **Core Pillars**

- Digital Innovation is not new for Grifols
- With our new organization, Grifols is moving to the next level
- Digital optimization and transformation to boost business growth
- Our culture will drive our future

#### Solid Foundation

- Strong technology team
- · Core skills and business knowledge
- High expertise in develop internal and external/commercial solutions
- Experts on service transformation/organizations integration
- · Reliable background on digitalization projects execution
- Powerful partnership ecosystem





Boosting efficient, agile, and innovative services and technology solutions for... 22,000+ users in 30 countries; 290+ donor centers; 13 manufacturing facilities; and more...

**Digital Background** 



provisible en de magnine o esperaba, el mueva Lines o actuel, compe como Staff o dovo Garcia co attas liness de consultor.

# March'82 New business line introducing Computers and Software

forets (luga fotografias

momento les ofrecessos y lodos Viley.

Fero... tenemos mas comes que infrecer: En primer lugar, el ordenador ya no es unicasents el Commundore, es tambien un nanva Sistema Computador de aspecto y forma profesional que nos permitira aborder empresablema ambienta. Ambien sos superiores esta apolicionas, dades sos superiores







GRI-CEL informática American Hospital Supply Corp.

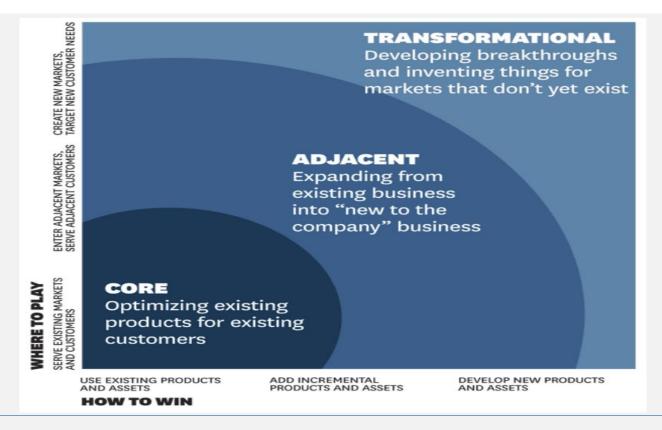
What Is the Difference Today?

#### Multiple technologies that once combined allows exponential change





#### **Innovation Models**





#### **Digital Business Transformation Approach**

Mission

Build digital capabilities to deliver better outcomes, explore new areas to play in and capture new sources of value

**Ambition** 

- Improve customer/patient/donor/employee experience
- Optimize operations efficiency/productivity
- Add value to our products/services changing the go-to-market model
- Unlock new value sources

Transformation

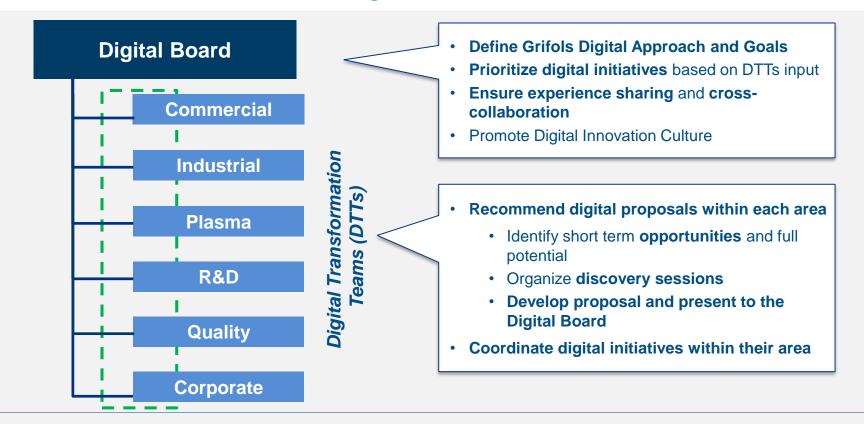
Optimization

Digitalization

Roadmap



Digital Board will Orchestrate the Grifols Digital Transformation





Digital Board will Orchestrate the Grifols Digital Transformation





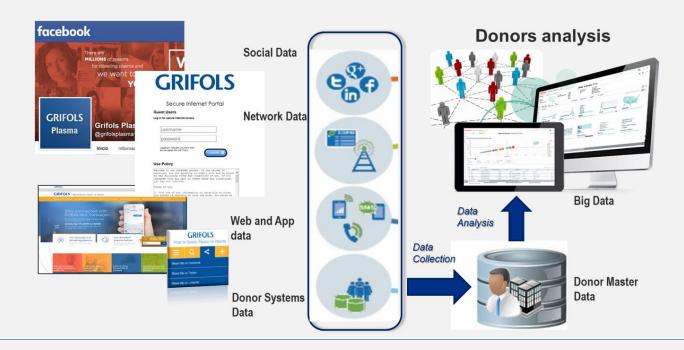
**Key Initiatives** 

#### **Currently 40 initiatives on-going**



Key Initiatives: Improve Customer/Patient/Donor/Employee Experience

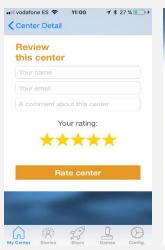
#### Improve Knowledge of Donors Digitalization to Improve Donor Experience





#### Key Initiatives: Improve Customer/Patient/Donor/Employee Experience

#### **Personalized Interactions**









Key Initiatives: Optimize Operations Efficiency/Productivity



## Digitalization of Donor Centre Operations with Grifols Donation System

- Donor Experience
- Operations Efficiencies
- Enterprise Digital Twin



Key Initiatives: Optimize Operations Efficiency/Productivity





#### **Data Analytics and Machine Learning**

- Supply Chain Optimization
- Manufacturing Process Optimization
- Connected Factory
- Predictive Maintenance

#### **Virtual and Augmented Reality Technology for:**

- Technical service remote assistance
- Employee Training
- Engineering Design

Key Initiatives: Optimize Operations Efficiency/Productivity



## Robotic Process Automation, Smart Workflows, Character Recognition and Natural Language Tools

- Automate routine tasks through existing interfaces (e.g., data extraction and cleaning)
- Integrate groups of tasks performed by humans & machines (e.g., calculating and applying allocations)
- Conversion of analog inputs into digital data (e.g., performing 2/3 way invoice matches)
- Create seamless interactions between humans & machines (e.g., chatbots for customer service)
- Digital Workplace for employees

**Blockchain technology discovery initiative with SAP Consortium** 



Key Initiatives: Add Value to our Products/Services Changing the Go-to-Market Model





## Robust ecosystem including software, services and devices focused on Hospital needs

- Improving security and quality
- Creating efficiencies (automation and data value)
- Integration and connectivity







Key Initiatives: Add Value to our Products/Services Changing the Go-to-Market Model



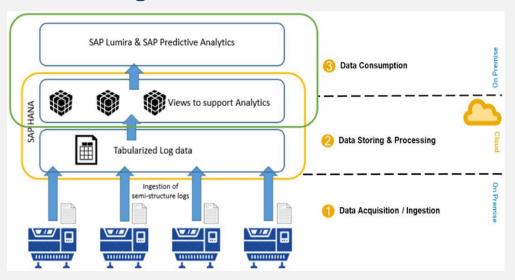
## Middleware Platform for Blood and Plasma Screening in Laboratories

- Optimize Labs Operations
- Increase Quality Processes
  - Increased Traceability
  - Streamlined Results Management
- User Satisfaction (easy to use easy to train)



Key Initiatives: Add Value to our Products/Services Changing the Go-to-Market Model

#### **Diagnostics Instruments: Connected devices and data analytics**



- Customer satisfaction and reliability
- Reduce ownership cost
- Field crew efficiency

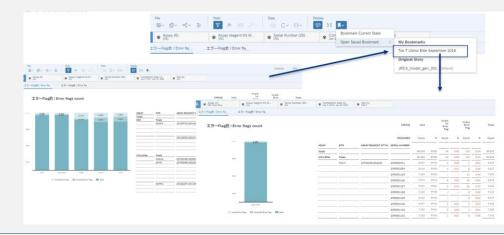


Key Initiatives: Add Value to our Products/Services Changing the Go-to-Market Model



#### **Japan Red Cross Analytics Services**

- Convert data into knowledge
- Improve testing outcomes and instruments performance
- Add value which help Grifols to differentiate and enhance relationships



Key Initiatives: Unlock New Value Sources

#### R&D drug and treatment discovery

- Identification of new protein entities
- Identification of new indications for our current products



Artificial Intelligence to evaluate both structured (clinical data) and non-structured information (manuscripts, patents, etc.)

It is Not Only About Technology

- Grifols Digital Day & Grifols Digital Talks
- Collaboration space
- Innovation space
- Co-innovation programs









## **Key Takeaways**



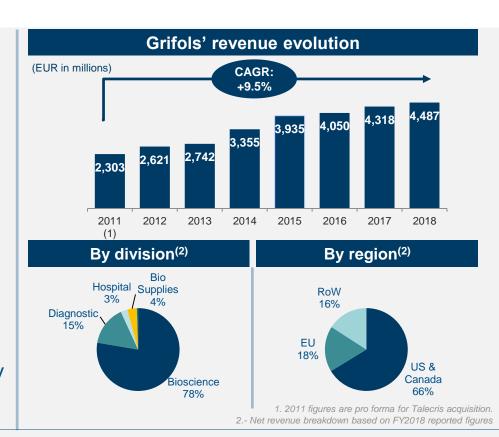
#### Key Takeaways

- Digital Innovation is not new for Grifols
- · With our new organization, Grifols is moving to the next level
- Digital optimization and transformation to boost business growth
- Our culture will drive our future



#### Positioned to Support Long-Term Growth

- Global presence with a diversified revenue base and solid upward growth
- Demonstrated ability to successfully grow businesses both organically and through acquisitions
- Significant value creation through acquisitions
- Poised to increase its exposure to the fastgrowing Chinese market
- Leading player in plasma-derivatives industry
- Vertically integrated business model



#### Positioned to Support Long-Term Growth through Innovation

- Grifols has been firmly committed to innovation since its foundation
- In 2018, Grifols intensified its net R+D+i investments by 15% at cc to EUR 291M¹. This investment represents 6.5% of revenues
- More than EUR 1,350M invested in R+D+i over the last 5 years
- Grifols advocates an integrated R+D+i strategy that comprises both in-house initiatives and external projects in investee companies whose research complements its core business
- Grifols earned the distinction as one of the top 1,000 global firms that dedicate the most resources to R+D in "2018 Global Innovation 1000" by Strategy&, the consulting arm of PwC

1. Taking into account net investments for both internal and external research initiatives

Positioned to Support Long-Term Growth through Global Expansion

- Strategically, Grifols seeks to strengthening its presence in China as a key global market
- Grifols' first commercial operations in China began back in the 1980's, and today it represents one of the major markets for Albumin
- The strategic alliance with Shanghai RAAS pursues to boost growth of its plasma-derived products and diagnostic solutions in China
- This agreement is an important step forward in Grifols' sustainable growth and long-term strategy, generating value for all of its divisions

## **Financial Highlights for 2018**



#### **Results Performance in 2018**

#### Financial Highlights

Sales EUR 4.5 billion Operating Growth +9.2%

EBITDA EUR 1.2 billion Adjusted Net
Profit
EUR 681 million

Net Operating Cash Flow EUR 962 million

- Strong operating sales growth driven by robust performance of Bioscience
- Reported growth in all divisions and geographic regions
- Substantial FX headwinds impacting top line (EUR 227 million)
- Margins impacted by higher plasma costs as a result of both organic and inorganic efforts, to fulfill the continued demand for its plasma-derived therapies
- Record-high dividend payout of EUR 279 million
- Leverage management remains among the company's top priorities

#### **Results Performance in 2018 - Bioscience**

Strong Sales Performance Driven by Volume and Price



- Robust sales of the main plasma proteins
- Solid demand for immunoglobulins in the U.S. and some EU markets, and alpha-1 antitrypsin sales remain strong in core markets
- Higher sales volume and positive pricing environment
- The renewal processes of certain licenses in China suffered delays in the last quarter of 2018, impacting sales growth and inventory

#### **Results Performance in 2018 - Diagnostic**

Performance Driven by NAT Technology and Blood Typing



- Higher NAT solutions sales were primarily fueled by an increase in plasma donations and the growing use of the Zika-virus screening test (Procleix® Zika Virus)
- Strong sales for NAT solutions in Latin America, Poland and Indonesia, in addition to the U.S.
- Continued efforts in the Middle East
- The blood-typing line notably contributed to the overall performance, particularly in the U.S. and in core markets in Latin America, Europe, and Saudi Arabia

#### **Results Performance in 2018 - Hospital**

Double-digit Growth Driven by Strong Performance in the U.S.



- Sales of all business lines grew in 2018, especially the Pharmatech line in the U.S. market. A key strategic area for future growth including MedKeeper and Kiro Oncology products
- The division also reported higher IV solutions sales, especially the physiological saline solution manufactured in the Murcia (Spain) plant

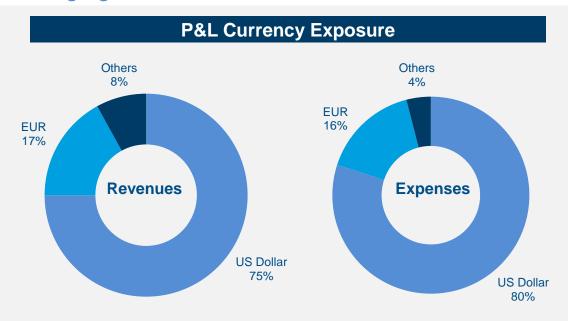
### Delivering on our ambitions in 2019

- Strong organic growth in all geographic regions
- Bioscience: Robust sales growth; cost per liter of plasma stable driven by higher plasma collections; new product launches; and margins improvements
- Inventory build-up as a key pillar of strategic growth
- Diagnostic: Extended contracts with 6 top accounts. Maintaining market share
- Hospital: Focus on execution leveraging on the U.S. market
- Successful closing of strategic acquisitions, integration well on track
- Substantial progress in implementation of strategic initiatives
- Expected FX tailwind



### **FX Tailwind**

### **Backed on Natural Hedging**



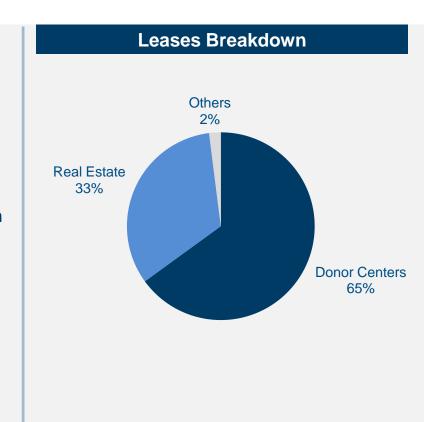


# **Applying IFRS 16: Major Impacts**

### IFRS 16: Major Impacts on Balance Sheet and P&L

### Lessee Accounting

- Single model for lease accounting by lessees
- Recognition a right-of-use asset (ROU) representing its right of use the underlying asset throughout the lease term and a lease liability representing its obligation to make future lease payments
- The ROU is measured at cost less accumulated depreciation
- Impact in P/L due to change in the nature of expenses.
   Lease cost replaced by: depreciation of right of use assets and interest on lease liabilities
- No impact on net cash flow. Shift form cash flows from operating activities to cash flows from financing activities
- · As per the financial covenant, no impact on leverage ratio





# IFRS 16: Major Impacts on Balance Sheet and P&L

**Lessee Accounting** 

Balance Sheet	
March 31, 2019	
ASSETS (right-of-use)	
EUR 690M	
DEBT (lease liability)	
· ·	
EUR 711M	

P&L

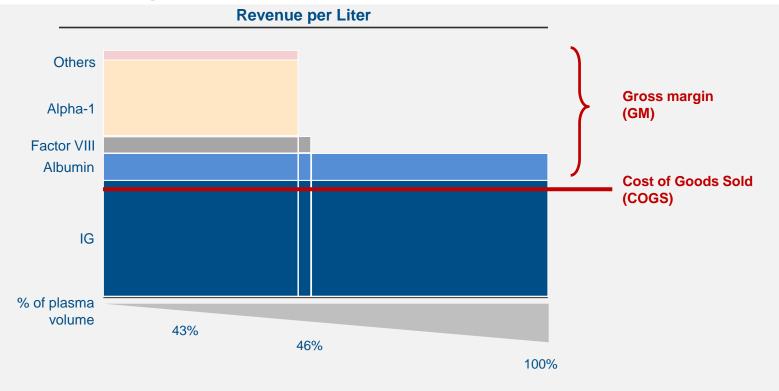
Estimates for FY20	19¹ (In EUR million)
+ Lease expense	56
= EBITDA	56
- Higher Depreciation	-58
= EBIT	+2
- Interest Expenses	-30
<b>Profit Before Taxes</b>	-28

1. FY2019 estimated based on 1Q 2019A. Ex-rate USD-EUR 1,15

# Plasma Economics



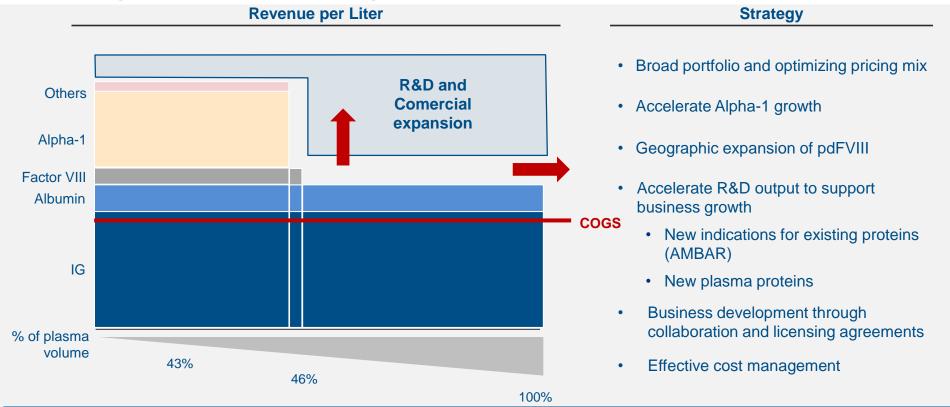
### The Paradigm of Balancing the Liter





### **Balancing the liter**

### Maximizing the Grifols' Gross Margin and Profit



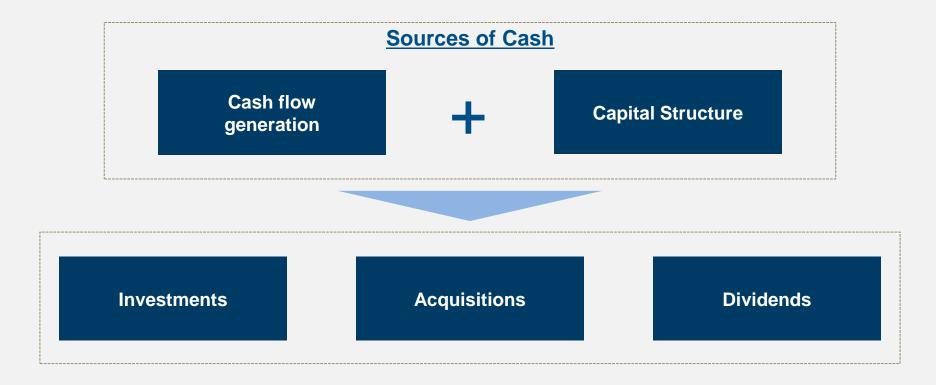


# **Capital Allocation**



# **Capital Allocation Supports Growth**

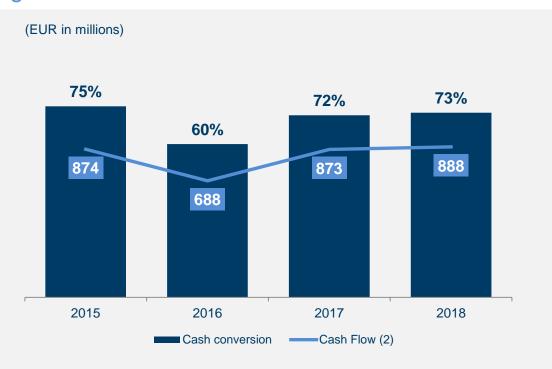
# Capital Discipline Focused on Creating Value





### **Strong Cash Flow Generation**

### High Conversion of EBITDA Into Cash Flow<sup>1,2</sup>

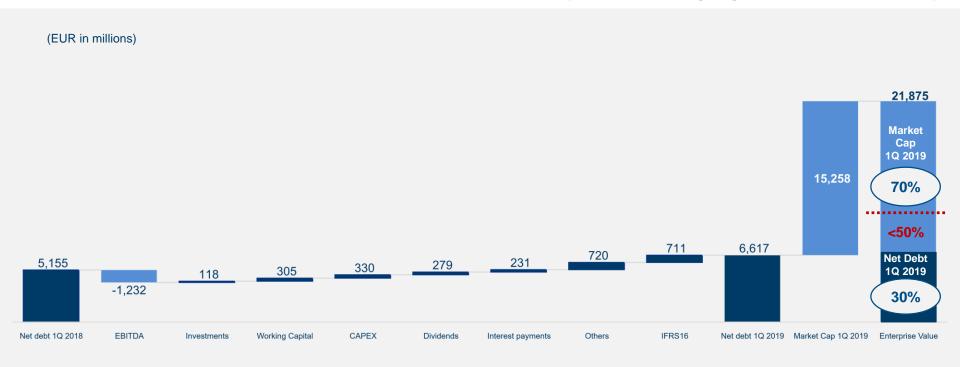


- Strong net cash flow from operating activities of EUR 962M, up from the average in 2014-2018 period
- Continued focus on cash flow from operating activities expansion driven by working capital management, operating performance and disciplined CAPEX

1.- Cash flow conversion defined as (EBITDA – Capex – Change in Working Capital) / EBITDA – Capex – Change in Working Capital

### **Capital Structure Breakdown**

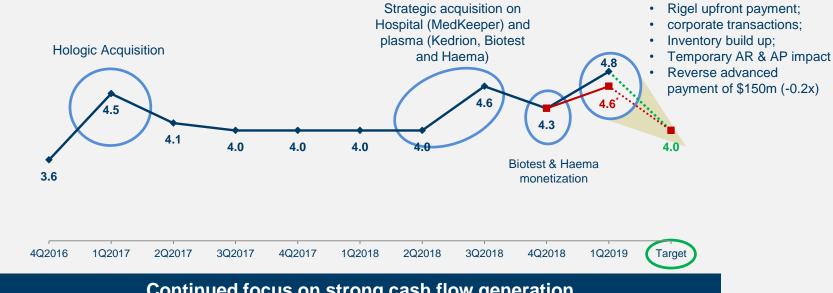
Debt to EV Ratio Enables Balance Sheet Flexibility. Deleveraging remains a priority



Market cap source: Bloomberg at March 31, 2019

### **Evolution of Leverage Ratio**

### Strategic Investments Lead Higher Leverage Ratio. Deleveraging Remains a Priority



Continued focus on strong cash flow generation

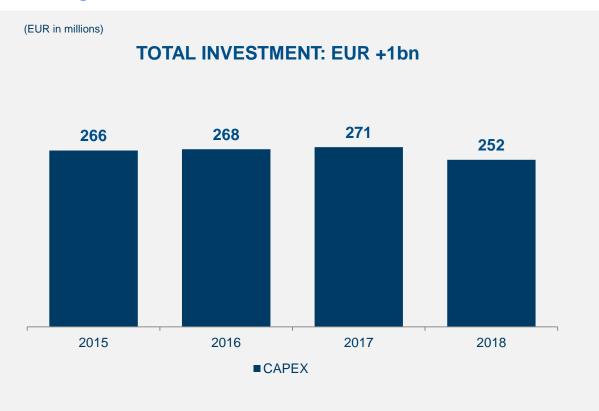
Leverage management remains among the company's top priorities Our target leverage ratio is below 4.0

> Leverage Ratio defined as Net Financial Debt to EBITDA excluding any IFRS 16 impact \* 4.6x excludes payment of \$150m



### **CAPEX Continues to Support Long-Term Growth**

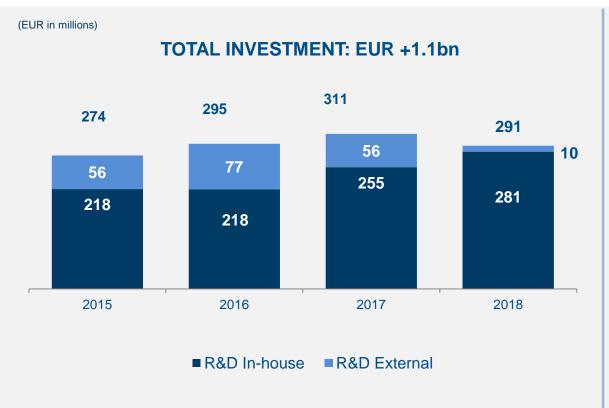
Allocating EUR 1,400M over 2018-2022



- Significant investments to support growth initiatives and innovation
- Continued emphasis on execution and capital allocation efficacy and return

### **R&D Continues to Support Long-Term Growth**

### Commitment to an Integrated Approach



- R&D drives long-term growth and profitability
- Includes strategic collaborations: leveraging internal and external expertise

### **Strategic Investment to Secure Growth**

### Haema and Biotest Transactions

Rationale

- To monetize recent acquisitions since there are still significant 3rd party supply agreements in place
- To strength the financial position while reducing financial leverage

Key terms

- Same price, terms and conditions existing at the time of Grifols acquisition (June 8 and August 2018)
- Call option to re-acquire exclusively and irrevocably at any time and at original acquisition price
- Current plasma agreements remain in place, extended to 30 years

### **Strategic Investment to Secure Growth**

### Haema and Biotest Transactions

Accounting treatment

- Transaction that results in change of ownership interest while retaining control
- Call Option >> potential voting rights >> control >> consolidation
- Full consolidation (P&L and BS) reversed in minorities

Business Management Agreement

- Covers all the existing key management services provided by Group companies
- Term in line with Plasma Supply Agreement at 30 years

### **Grifols Next Growth Engine**

### Strategic Alliance with Shanghai RAAS (SRAAS) in China



- Grifols will contribute 45% economic rights in Grifols Diagnostic Solutions (GDS) Group into SRAAS and 40% voting rights in GDS
- In exchange, Grifols will acquire 26.2% stake in SRAAS (voting and economic rights);
- Grifols' protections as shareholder include:
  - Grifols will have the right to appoint 2 non-independent directors, out of 6 non-independent directors and 3 independent directors existing in SRAAS in total at the board of directors;
  - SRAAS articles of association will include 75% reinforced quorum to issue shares, disposal of material assets, mergers and modify the articles of associations (Grifols having veto on all these);
  - Grifols will have a right of first refusal upon sale of shares of Creat and pre-emption rights on issuance of new shares in SRAAS, to avoid dilution

### **Grifols Next Growth Engine**

### Strategic Alliance with Shanghai RAAS (SRAAS) in China



- Grifols and SRAAS will enter into an exclusive Strategic Alliance Agreement whereby:
  - Grifols and SRAAS shall enter into a Quality Agreement to set up high International quality standards;
  - To appoint a quality person and manufacturing person to follow compliance with such agreed quality agreement;
  - SRAAS become the exclusive distributor of Grifols in China;
  - In exchange of royalties to be agreed upon, Grifols shall provide technology owned or controlled by Grifols to SRAAS for use in China;
  - Grifols shall provide engineering services to SRAAS in exchange of fees to be agreed upon; and
  - SRAAS commits to use GDS NAT technology in its plasma collection business

### **Grifols Next Growth Engine**

### Strategic Alliance with Shanghai RAAS (SRAAS) in China

Key Transaction<sup>(\*)</sup> Terms

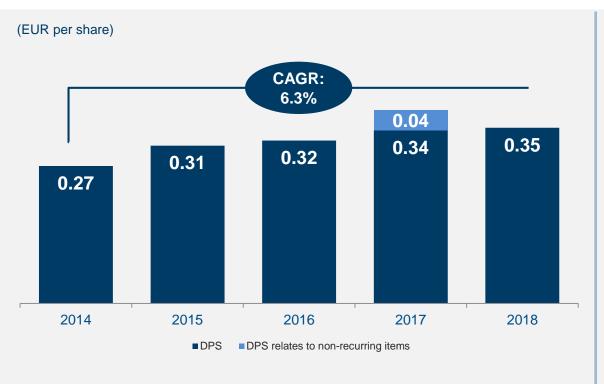
- Shanghai RAAS will have the right to appoint 1 director at GDS and will have similar antidilution protections as Grifols in SRAAS;
- Creat will be prevented from selling its stake in SRAAS to a competitor of Grifols and Grifols will be prevented from selling its stake in GDS to any Chinese entity

Accounting treatment

- **Grifols will fully consolidate GDS Group**, adjusting the 45% of GDS as a minority;
- Grifols will include the 26.2% of SRAAS net profit at EBITDA level

### **Return to Shareholders**

### Sharing success with shareholders



- Strong earnings profile
- Accumulated annual dividend up by 6.3% over the last 5 years
- More than EUR 1,000M returned to shareholders over the last 5 years
- Pay-out ratio 40% of reported consolidated profits



# **Key Takeaways**



### **Key Takeaways**

### Continuous Focus on Long-Term Growth

- Continued support to fund long-term growth
- Steady focus on business fundamentals and global expansion
- Capital allocation efficient and focused on strategic opportunities, industrial and plasma capacity needs and R&D initiatives
- Working capital management to optimize growth while maximizing cash flow generation
- Expected gradual reduction of leverage ratio to c.4.5x by 2019; while target is set to below 4.0x. Company is very mindful of its leverage – remains a key priority
- Shanghai RAAS: the right partner and agreement into the fastest-growing market to create long-term value
- Constant shareholders reward through 40% pay-out

# **Grifols: a Socially Responsible Company**Within Grifols' DNA Since Our Origins

Teresa Rioné VP, Corporate Communications

**GRIFOLS** 

"The right to live in society entails the duty to work to improve it."



Josep Antoni Grífols i Roig, 1976











"At Grifols, we believe in responsible management to generate social, economic and environmental value."

Raimon Grífols Roura Víctor Grífols Deu 2018



### **MISSION**

Our mission is to improve the health and well-being of patients around the world

#### **VISION**

We strive to be a global leader in our markets and a constant industry reference for innovation, quality and safety



Donors and Patients at the Heart of Grifols' Business Model





# **Corporate Social Responsibility**

**Key Areas** 









# **Corporate Social Responsibility**



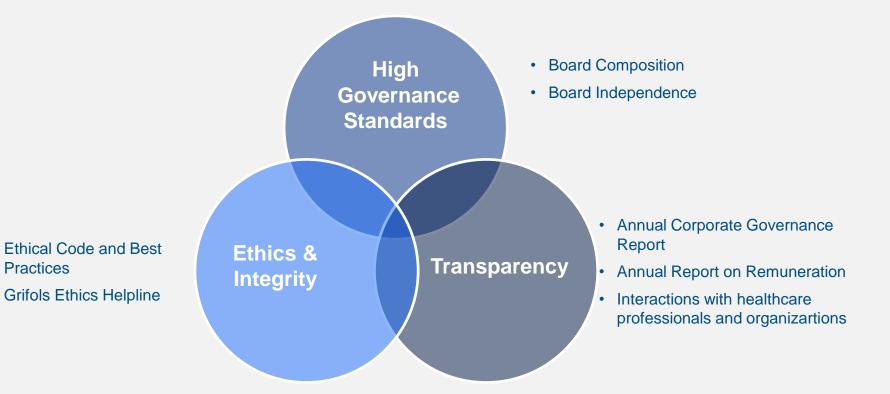






### **Corporate Social Responsibility**

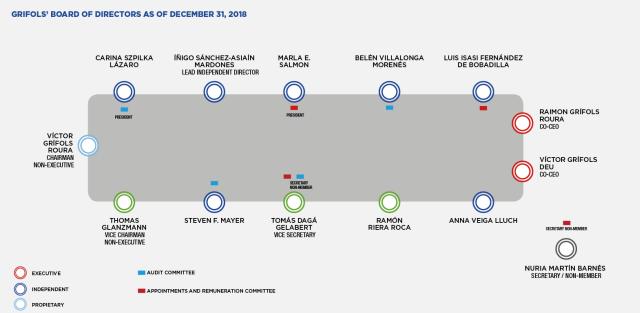
### **Key Principles**





**Practices** 

### High Governance Standards: Composition of Board of Directors



#### **Diverse and well-balanced Board:**

- Even distribution by number of years on Board (31% 1-3 years, 38% 4-10 years, 31% +10 years)
- 31% of Board members are women
- Board members represented in all age groups (39% aged <55, 46% aged 55-65, 15% aged >65)
- Diverse professional experience and career paths (financial, healthcare, research, law)



OTHER EXTERNAL

High Governance Standards: Independence

### **Boards of Directors**

Differentiated roles of President and CEOs

Lead Independent Director

85% Non-Executive

>50% Independent

# **Board of Directors' Committees**

Independent President

Non-Executive only

≥2 out of 3 Independent



### Transparency: Our Commitments Beyond Compliance



### **Interactions with Healthcare Professionals and Organizations**

Indicates interactions with the medical porthesion have a ser- ment has value in incentent it, for interpret provides an interaction as unappe, independent insights, and export knowledge in solvenings of the exportate policy a cotonic role in informing a patient care and instrument optimism. For these research, CP profiscionals and invalidations, as well as that this services the	to patient behavior and situace management. The ability and goving instituting efforts to impose the goldity of this advocates for compensation for healthcome.
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Choose Country	

#### The United States

Sunshine Act

### **2019 Plans:**

- New Transparency Training Program (Employees)
- Quarterly sub-certification process

#### **EU** countries

- All relevant country-specific transparency standards
- Practices voluntarily adopted
  - EFPIA Disclosure Code
  - MedTech Europe Code of Business Practice
  - Grifols Global Compliance Program

Ethics & Integrity: Robust Corporate Policies





## **Corporate Governance**

Ethics & Integrity: Our Commitments Beyond Legal Compliance

## **Compliance Function**

- Ensures Grifols complies with all applicable anticorruption laws, rules and regulations
- Applies best practices within the organization
- Strict and immediate response to possible violations

# **Employee Training and Helpline**

- Employees are trained in anticorruption practices
- Grifols Ethics Helpline to confidentially raise concerns of non-compliance or misconduct

# Third Party Anticorruption Management Practices

- Exhaustive screening
- Due Diligence
- Ethical standards and monitoring



## **Corporate Social Responsibility**

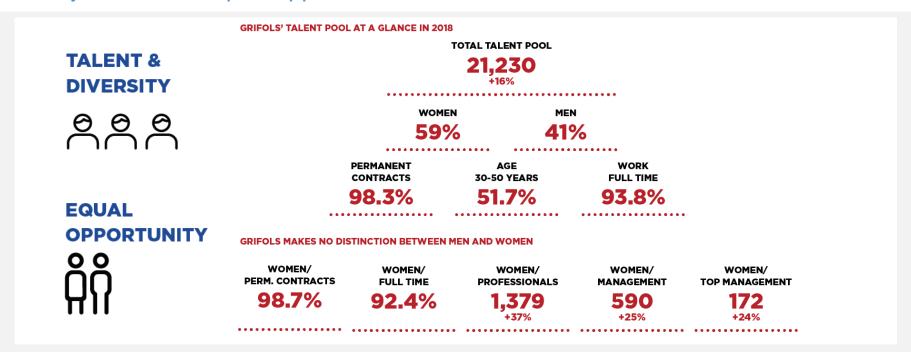








Diversity, Inclusion, Equal Opportunities and Non-Discrimination



Grifols has reduced the gender pay gap, which is below international benchmarks in all professional categories



#### **Talent Attraction and Retention**



Competitive Remuneration package

Career Development Opportunities

Annual and Systematic **Performance review** 

## **Talent Development**



#### **Continuous Training and Development**

- Leadership Development
  - Executive Education Programs (ESADE, Georgetown)
- Ongoing Professional Development
  - Grifols Academy Programs
- Onboarding Programs

2.5 million hours
Total training in 2018

138 hours
Training per employee in 2018

## **Talent Development**



#### THE GRIFOLS ACADEMY PROFESSIONAL DEVELOPMENT



## PLASMAPHERESIS



#### THE GRIFOLS ACADEMY THE GRIFOLS ACADEMY TRANSFUSION MEDICINE

- . Training and professional development for Grifols employees.
- · Aimed at strengthening specific competencies and fostering Grifols' corporate culture.
- · Programs fall into three core areas: scientific-technical knowledge. skills development and leadership competencies.
- Offers advanced training on all plasmapheresis procedures; collection, analysis and control of plasma; manufacture of plasma-derived medicines and other ethical and quality issues framed within the area of human health.
- · Allows the company to transmit its knowledge, standardize procedures and increase employee engagement, while fostering its corporate culture in Grifols' U.S.-based facilities
- · Offers educational programs on transfusion medicine to global professionals.
- . Designed to enhance patient care by contributing to the advancement of knowledge in this field.

## **Corporate Social Responsibility**









#### Recognized for our Sustainability Initiatives









## Carbon Disclosure Project

Businesses leading on Managing Climate Change (level B)

#### **FTSE4Good Index**

Companies that meet globally recognized Corporate Responsibility Standards

#### ISO 14001 Certified

International Standards for Effective Environmental Management

#### **LEED**

Clayton Plant recognition for its sustainable design in two new buildings

#### **Environmental Management**



#### **ENVIRONMENTAL POLICY**

Defines the company-wide principles and commitments common to the entire company aimed at monitoring and improving Grifols' environmental impact.



**ENERGY POLICY** 

Defines the company-wide principles to the entire company to optimize its energy resources.



## CORPORATE ENVIRONMENT MANUAL

Reference manual applicable to all ISO-14001-certified manufacturing facilities or in process. It serves as a reference manual for the company's environmental performance.



## ENVIRONMENTAL PROGRAM

Defines the specific action lines for each business area. The 2017-2019 Environmental Program is currently in force.



### NVIRONMENIAL COMMITTEES

Monitor the environmental management system of all Grifols' companies. Among other functions, they monitor the progress of environmental objectives, review of follow-up indicators, application of corrective measures and compliance with current legislation and identification of opportunities for improvement.

#### 18 million euros allocated to environmental initiatives in 2018

## Our Commitments – Key Achievements in 2019



**Clayton Facilities** 

## 2019 "ZERO WASTE TO LANDFILL" GOLD CERTIFICATION

- Zero Waste Policy in place
- Over 99% waste recovery
- Only 5% of incineration with energy recovery



#### Our Commitments



#### **Directive (EU) 2018/2001**

On the promotion of the use of energy from renewable sources

**2030 target:** ≥ **32%** of energy sources must be renewable in the EU

We are already investing in renewable energy sources:

- A new onsite photovoltaic generation plant in Murcia (Spain)
- Establishing PPAs (Power Purchase Agreement) in order to gradually shift towards green energy.

Grifols is committed to continue investing in environmental sustainability beyond the international regulatory requirements

## **Corporate Social Responsibility**









## Transparency



Patient Organizations



Local
Communities
Plasma Donor
Centers



Research



**Foundations** 



**Special Projects** 

33.3 million euros allocated to social initiatives in 2018

#### Our Commitments with Patients

#### **Educational Programs**

#### For patients and their families

- U.S. Patient Community
   Open Houses
- ES Alpha-1 antitripsin deficiency Organization awareness trekking experience

#### **Support and Patient Care**

ALFACARE, a patientassistance program for patients with alpha-1 antitrypsin deficiency in Spain

- Personalized Support (emotional, psychological)
- Complements Standard Care

Similar programs have been rolled out in other countries (US, DE, CA)

#### **Access to Treatment**

**U.S. PatientCare Program** for Patients with Hemophilia or Primary Immunodeficiency (since 2006)

25 million International Units of Clotting factor donated to WFH Humanitarian Aid Program in 2018

 Treatment for 6,000 patients per year until 2021

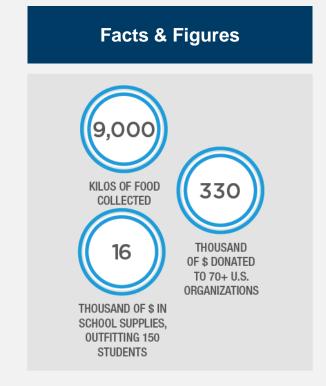


#### Our Commitments with Plasma Donors and Local Communities

#### **Plasma Donors - Recognition**

- Compensation for Donors' time and commitment
- Same Remuneration for All Donors (no age, weight or gender distinction)
- Complete Health Screening
- Plasma Possibilities
   Program (waiving fee for Charity Organizations) since 2017





#### Awards

#### Research



Grifols has a long-standing commitment to the scientific community and promotes awards in research related to our core business areas

Martín Villar Haemostasis Awards

SPIN, Scientific Progress Immunoglobulins In Neurology Award

ALTA, Alpha-1-antitrypsin Laurell's Training Award

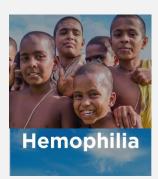
Albus, **Albumin** Awards Program

GATRA\*, Grifols Antithrombin Research Awards

\* There were no GATRA granted in 2018

#### **Awareness**



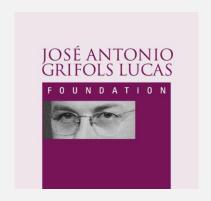


# Grifols supports education and access to treatment in developing countries

- Outreach diagnosis
- Improve education
- Facilitate access to treatment.



#### **Foundations**







The Foundation pays tribute to the memory of Dr. José Antonio Grifols Lucas, who developed the plasmapheresis technique, and recognizes the indispensable value of the donors.

José Antonio Grifols Lucas Foundation **contributes to the communities** where Grifols operates its plasma donation centers **through health**, **wellbeing and educational programs**.

#### **Foundations**







The Víctor Grífols i Lucas Foundation was established in 1998 to spark cross-disciplinary debate on **bioethics**.

The Foundation seeks to foster **ethical attitudes and create new ideas** and insights in organizations, companies and professionals in the field of human health.

Main activities include conferences, seminars and courses as forums to exchange different perspectives.

#### **Foundations**





GLI Global Laboratory Initiative



RAI
Child Nutrition Support
Program



SIT
Health Innovation and
Therapies



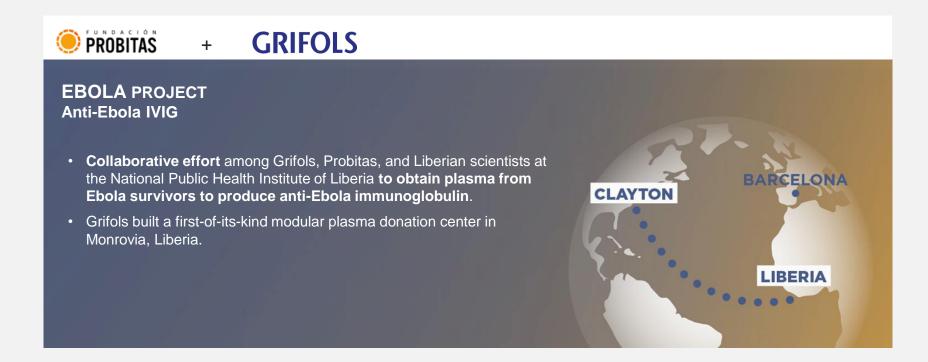
PCI
International Partnership
Program

Fundación Probitas was created in 2008 to leverage **Grifols' expertise in the healthcare field and contribute to enhancing medical care in areas with limited resources.** 

Probitas **combines in-house programs and external collaborations** with NGOs in the humanitarian sector (Spanish Red Cross, Save the Children, World Food Program).

**0.7% of Grifols corporate profits** go to support this private foundation.

#### **Foundations**





## Corporate Social Responsibility

Within Grifols' DNA since our origins

22 21 YEARS OF YEARS OF **VÍCTOR GRIFOLS GRIFOLS MUSEUM** I LUCAS FOUNDATION **Ethics** Heritage 10 10 YEARS OF YEARS OF **JOSÉ ANTONIO PROBITAS FOUNDATION GRIFOLS I LUCAS** Social **FOUNDATION Donors** 

# **Executing a New Chapter of Continued Growth and Success**

Víctor Grífols Deu Co-CEO

**GRIFOLS** 

## **Executing a New Chapter of Continued Growth and Success**

OVER THE PAST TWO YEARS, EVERY DECISION HAS BEEN A BUILDING BLOCK THAT HAS PAVED OUR WAY INTO THE FUTURE...



...WE BELIEVE EVERYTHING IS NOW IN PLACE, NOW IS A MATTER OF RIGHT EXECUTION...

Key priorities moving forward



...TO ENSURE OUR CONTINUED GROWTH & SUCCESS

Corporate Focus areas





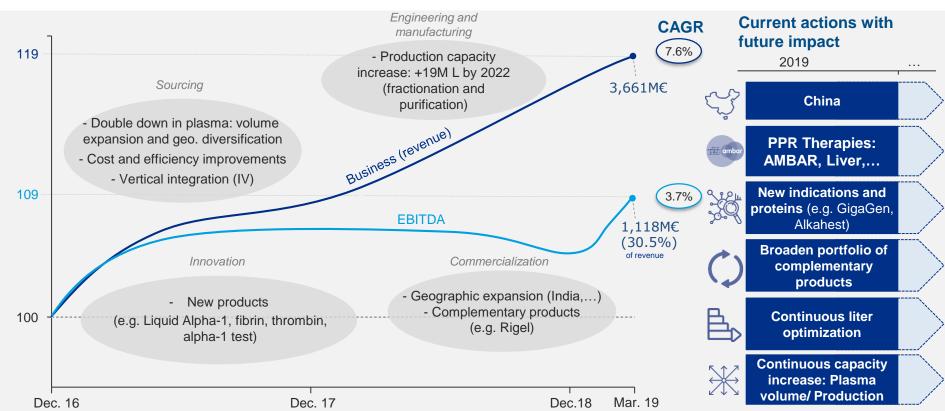
## Business Evolution – Bioscience (LTM, at CC)<sup>1</sup>





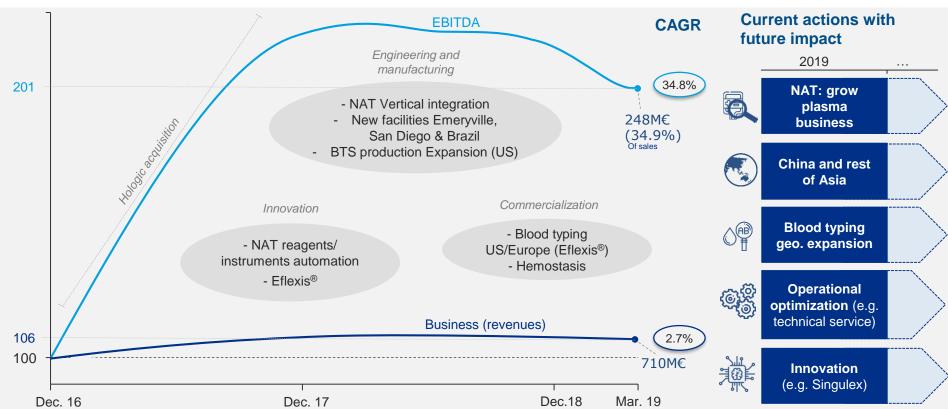






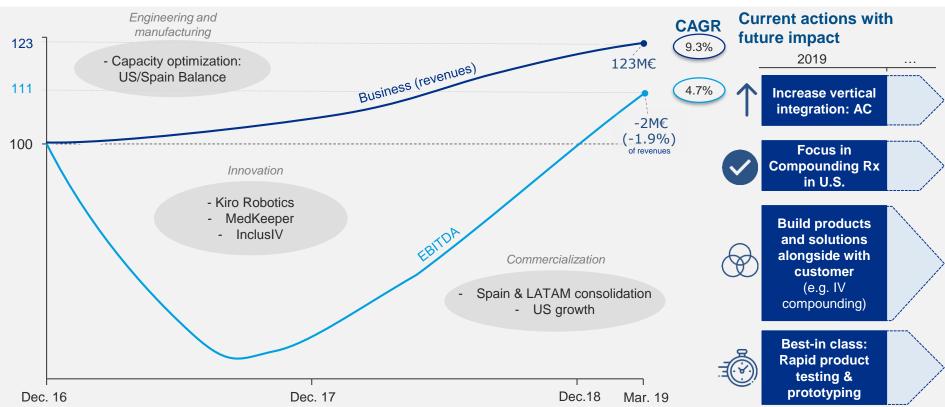
## Business Evolution – Diagnostic (LTM, at CC)<sup>1</sup>





## Business Evolution – Hospital (LTM, at CC)<sup>1</sup>





## Business Evolution – Group (LTM, at CC)<sup>1</sup>









